

Physical interventions to interrupt or reduce the spread of respiratory viruses

Author

Jefferson, Tom, Del Mar, Chris B, Dooley, Liz, Ferroni, Eliana, Al-Ansary, Lubna A, Bawazeer, Ghada A, van Driel, Mieke L, Jones, Mark A, Thorning, Sarah, Beller, Elaine M, Clark, Justin, Hoffmann, Tammy C, Glasziou, Paul P, Conly, John M

Published

2020

Journal Title

Cochrane Database of Systematic Reviews

Version

Version of Record (VoR)

DOI

<https://doi.org/10.1002/14651858.CD006207.pub5>

Copyright Statement

© 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd. This review is published as a Cochrane Review in the Cochrane Database of Systematic Reviews 2020, Issue 11. Art.No.: CD006207. Cochrane Reviews are regularly updated as new evidence emerges and in response to comments and criticisms, and the Cochrane Database of Systematic Reviews should be consulted for the most recent version of the Review.

Downloaded from

<http://hdl.handle.net/10072/399941>

Griffith Research Online

<https://research-repository.griffith.edu.au>



Cochrane
Library

Cochrane Database of Systematic Reviews

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Jefferson T, Del Mar CB, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, van Driel ML, Jones MA, Thorning S, Beller EM, Clark J, Hoffmann TC, Glasziou PP, Conly JM

Jefferson T, Del Mar CB, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, van Driel ML, Jones MA, Thorning S, Beller EM, Clark J, Hoffmann TC, Glasziou PP, Conly JM.

Physical interventions to interrupt or reduce the spread of respiratory viruses.

Cochrane Database of Systematic Reviews 2020, Issue 11. Art. No.: CD006207.

DOI: [10.1002/14651858.CD006207.pub5](https://doi.org/10.1002/14651858.CD006207.pub5).

www.cochranelibrary.com

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Copyright © 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

WILEY

TABLE OF CONTENTS

HEADER	1
ABSTRACT	1
PLAIN LANGUAGE SUMMARY	3
SUMMARY OF FINDINGS	5
BACKGROUND	9
OBJECTIVES	9
METHODS	9
RESULTS	11
Figure 1.	12
Figure 2.	14
Figure 3.	15
DISCUSSION	25
AUTHORS' CONCLUSIONS	28
ACKNOWLEDGEMENTS	30
REFERENCES	31
CHARACTERISTICS OF STUDIES	47
DATA AND ANALYSES	201
Analysis 1.1. Comparison 1: Randomised trials: medical/surgical masks versus no masks, Outcome 1: Viral illness	201
Analysis 1.2. Comparison 1: Randomised trials: medical/surgical masks versus no masks, Outcome 2: Influenza-like illness in healthcare workers	202
Analysis 2.1. Comparison 2: Randomised trials: N95 respirators compared to medical/surgical masks, Outcome 1: Viral illness ..	203
Analysis 2.2. Comparison 2: Randomised trials: N95 respirators compared to medical/surgical masks, Outcome 2: Viral illness in healthcare workers	204
Analysis 3.1. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 1: Viral illness	206
Analysis 3.2. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 2: ARI or ILI or influenza (including outcome with most events from each study)	207
Analysis 3.3. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 3: Influenza or ILI: sensitivity analysis including outcomes with the most precise and unequivocal definitions	207
Analysis 3.4. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 4: ARI or ILI or influenza: subgroup analysis	208
Analysis 3.5. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 5: Absenteeism	208
Analysis 4.1. Comparison 4: Randomised trials: hand hygiene + medical/surgical masks compared to control, Outcome 1: Viral illness	209
Analysis 5.1. Comparison 5: Randomised trials: hand hygiene + medical/surgical masks compared to hand hygiene, Outcome 1: Viral illness	210
Analysis 6.1. Comparison 6: Randomised trials: gargling compared to control, Outcome 1: Viral illness	210
ADDITIONAL TABLES	210
APPENDICES	296
WHAT'S NEW	307
HISTORY	308
CONTRIBUTIONS OF AUTHORS	309
DECLARATIONS OF INTEREST	309
SOURCES OF SUPPORT	310
DIFFERENCES BETWEEN PROTOCOL AND REVIEW	310
NOTES	310
INDEX TERMS	311

[Intervention Review]

Physical interventions to interrupt or reduce the spread of respiratory viruses

Tom Jefferson¹, Chris B Del Mar², Liz Dooley², Eliana Ferroni³, Lubna A Al-Ansary⁴, Ghada A Bawazeer⁵, Mieke L van Driel^{6,7}, Mark A Jones², Sarah Thorning⁸, Elaine M Beller², Justin Clark², Tammy C Hoffmann², Paul P Glasziou², John M Conly^{9,10,11}

¹Centre for Evidence Based Medicine, University of Oxford, Oxford, UK. ²Institute for Evidence-Based Healthcare, Bond University, Gold Coast, Australia. ³Epidemiological System of the Veneto Region, Regional Center for Epidemiology, Veneto Region, Padova, Italy. ⁴Department of Family and Community Medicine, King Saud University, Riyadh, Saudi Arabia. ⁵Department of Clinical Pharmacy, College of Pharmacy, King Saud University, Riyadh, Saudi Arabia. ⁶Primary Care Clinical Unit, Faculty of Medicine, The University of Queensland, Brisbane, Australia. ⁷Department of Public Health and Primary Care, Ghent University, Ghent, Belgium. ⁸GCUH Library, Gold Coast Hospital and Health Service, Southport, Australia. ⁹Cumming School of Medicine, University of Calgary, Room AGW5, SSB, Foothills Medical Centre, Calgary, Canada. ¹⁰O'Brien Institute for Public Health and Synder Institute for Chronic Diseases, Cumming School of Medicine, University of Calgary, Calgary, Canada. ¹¹Calgary Zone, Alberta Health Services, Calgary, Canada

Contact address: John M Conly, john.conly@albertahealthservices.ca.

Editorial group: Cochrane Acute Respiratory Infections Group.

Publication status and date: New search for studies and content updated (conclusions changed), published in Issue 11, 2020.

Citation: Jefferson T, Del Mar CB, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, van Driel ML, Jones MA, Thorning S, Beller EM, Clark J, Hoffmann TC, Glasziou PP, Conly JM. Physical interventions to interrupt or reduce the spread of respiratory viruses. *Cochrane Database of Systematic Reviews* 2020, Issue 11. Art. No.: CD006207. DOI: [10.1002/14651858.CD006207.pub5](https://doi.org/10.1002/14651858.CD006207.pub5).

Copyright © 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

ABSTRACT

Background

Viral epidemics or pandemics of acute respiratory infections (ARIs) pose a global threat. Examples are influenza (H1N1) caused by the H1N1pdm09 virus in 2009, severe acute respiratory syndrome (SARS) in 2003, and coronavirus disease 2019 (COVID-19) caused by SARS-CoV-2 in 2019. Antiviral drugs and vaccines may be insufficient to prevent their spread. This is an update of a Cochrane Review published in 2007, 2009, 2010, and 2011. The evidence summarised in this review does not include results from studies from the current COVID-19 pandemic.

Objectives

To assess the effectiveness of physical interventions to interrupt or reduce the spread of acute respiratory viruses.

Search methods

We searched CENTRAL, PubMed, Embase, CINAHL on 1 April 2020. We searched ClinicalTrials.gov, and the WHO ICTRP on 16 March 2020. We conducted a backwards and forwards citation analysis on the newly included studies.

Selection criteria

We included randomised controlled trials (RCTs) and cluster-RCTs of trials investigating physical interventions (screening at entry ports, isolation, quarantine, physical distancing, personal protection, hand hygiene, face masks, and gargling) to prevent respiratory virus transmission. In previous versions of this review we also included observational studies. However, for this update, there were sufficient RCTs to address our study aims.

Data collection and analysis

We used standard methodological procedures expected by Cochrane. We used GRADE to assess the certainty of the evidence. Three pairs of review authors independently extracted data using a standard template applied in previous versions of this review, but which was revised

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

1

Copyright © 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

to reflect our focus on RCTs and cluster-RCTs for this update. We did not contact trialists for missing data due to the urgency in completing the review. We extracted data on adverse events (harms) associated with the interventions.

Main results

We included 44 new RCTs and cluster-RCTs in this update, bringing the total number of randomised trials to 67. There were no included studies conducted during the COVID-19 pandemic. Six ongoing studies were identified, of which three evaluating masks are being conducted concurrent with the COVID pandemic, and one is completed.

Many studies were conducted during non-epidemic influenza periods, but several studies were conducted during the global H1N1 influenza pandemic in 2009, and others in epidemic influenza seasons up to 2016. Thus, studies were conducted in the context of lower respiratory viral circulation and transmission compared to COVID-19. The included studies were conducted in heterogeneous settings, ranging from suburban schools to hospital wards in high-income countries; crowded inner city settings in low-income countries; and an immigrant neighbourhood in a high-income country. Compliance with interventions was low in many studies.

The risk of bias for the RCTs and cluster-RCTs was mostly high or unclear.

Medical/surgical masks compared to no masks

We included nine trials (of which eight were cluster-RCTs) comparing medical/surgical masks versus no masks to prevent the spread of viral respiratory illness (two trials with healthcare workers and seven in the community). There is low certainty evidence from nine trials (3507 participants) that wearing a mask may make little or no difference to the outcome of influenza-like illness (ILI) compared to not wearing a mask (risk ratio (RR) 0.99, 95% confidence interval (CI) 0.82 to 1.18). There is moderate certainty evidence that wearing a mask probably makes little or no difference to the outcome of laboratory-confirmed influenza compared to not wearing a mask (RR 0.91, 95% CI 0.66 to 1.26; 6 trials; 3005 participants). Harms were rarely measured and poorly reported. Two studies during COVID-19 plan to recruit a total of 72,000 people. One evaluates medical/surgical masks (N = 6000) (published *Annals of Internal Medicine*, 18 Nov 2020), and one evaluates cloth masks (N = 66,000).

N95/P2 respirators compared to medical/surgical masks

We pooled trials comparing N95/P2 respirators with medical/surgical masks (four in healthcare settings and one in a household setting). There is uncertainty over the effects of N95/P2 respirators when compared with medical/surgical masks on the outcomes of clinical respiratory illness (RR 0.70, 95% CI 0.45 to 1.10; very low-certainty evidence; 3 trials; 7779 participants) and ILI (RR 0.82, 95% CI 0.66 to 1.03; low-certainty evidence; 5 trials; 8407 participants). The evidence is limited by imprecision and heterogeneity for these subjective outcomes. The use of a N95/P2 respirator compared to a medical/surgical mask probably makes little or no difference for the objective and more precise outcome of laboratory-confirmed influenza infection (RR 1.10, 95% CI 0.90 to 1.34; moderate-certainty evidence; 5 trials; 8407 participants). Restricting the pooling to healthcare workers made no difference to the overall findings. Harms were poorly measured and reported, but discomfort wearing medical/surgical masks or N95/P2 respirators was mentioned in several studies. One ongoing study recruiting 576 people compares N95/P2 respirators with medical surgical masks for healthcare workers during COVID-19.

Hand hygiene compared to control

Settings included schools, childcare centres, homes, and offices. In a comparison of hand hygiene interventions with control (no intervention), there was a 16% relative reduction in the number of people with ARIs in the hand hygiene group (RR 0.84, 95% CI 0.82 to 0.86; 7 trials; 44,129 participants; moderate-certainty evidence), suggesting a probable benefit. When considering the more strictly defined outcomes of ILI and laboratory-confirmed influenza, the estimates of effect for ILI (RR 0.98, 95% CI 0.85 to 1.13; 10 trials; 32,641 participants; low-certainty evidence) and laboratory-confirmed influenza (RR 0.91, 95% CI 0.63 to 1.30; 8 trials; 8332 participants; low-certainty evidence) suggest the intervention made little or no difference. We pooled all 16 trials (61,372 participants) for the composite outcome of ARI or ILI or influenza, with each study only contributing once and the most comprehensive outcome reported. The pooled data showed that hand hygiene may offer a benefit with an 11% relative reduction of respiratory illness (RR 0.89, 95% CI 0.84 to 0.95; low-certainty evidence), but with high heterogeneity. Few trials measured and reported harms.

There are two ongoing studies of handwashing interventions in 395 children outside of COVID-19.

We identified one RCT on quarantine/physical distancing. Company employees in Japan were asked to stay at home if household members had ILI symptoms. Overall fewer people in the intervention group contracted influenza compared with workers in the control group (2.75% versus 3.18%; hazard ratio 0.80, 95% CI 0.66 to 0.97). However, those who stayed at home with their infected family members were 2.17 times more likely to be infected.

We found no RCTs on eye protection, gowns and gloves, or screening at entry ports.

Authors' conclusions

The high risk of bias in the trials, variation in outcome measurement, and relatively low compliance with the interventions during the studies hamper drawing firm conclusions and generalising the findings to the current COVID-19 pandemic.

There is uncertainty about the effects of face masks. The low-moderate certainty of the evidence means our confidence in the effect estimate is limited, and that the true effect may be different from the observed estimate of the effect. The pooled results of randomised trials did not show a clear reduction in respiratory viral infection with the use of medical/surgical masks during seasonal influenza. There were no clear differences between the use of medical/surgical masks compared with N95/P2 respirators in healthcare workers when used in routine care to reduce respiratory viral infection. Hand hygiene is likely to modestly reduce the burden of respiratory illness. Harms associated with physical interventions were under-investigated.

There is a need for large, well-designed RCTs addressing the effectiveness of many of these interventions in multiple settings and populations, especially in those most at risk of ARIs.

PLAIN LANGUAGE SUMMARY

Do physical measures such as hand-washing or wearing masks stop or slow down the spread of respiratory viruses?

What are respiratory viruses?

Respiratory viruses are viruses that infect the cells in your airways: nose, throat, and lungs. These infections can cause serious problems and affect normal breathing. They can cause flu (influenza), severe acute respiratory syndrome (SARS), and COVID-19.

How do respiratory viruses spread?

People infected with a respiratory virus spread virus particles into the air when they cough or sneeze. Other people become infected if they come into contact with these virus particles in the air or on surfaces on which they have landed. Respiratory viruses can spread quickly through a community, through populations and countries (causing epidemics), and around the world (causing pandemics).

How can we stop the spread of respiratory viruses?

Physical measures to try to stop respiratory viruses spreading between people include:

- washing hands often;
- not touching your eyes, nose, or mouth;
- sneezing or coughing into your elbow;
- wiping surfaces with disinfectant;
- wearing masks, eye protection, gloves, and protective gowns;
- avoiding contact with other people (isolation or quarantine);
- keeping a certain distance away from other people (distancing); and
- examining people entering a country for signs of infection (screening).

Why we did this Cochrane Review

We wanted to find out whether physical measures stop or slow the spread of respiratory viruses.

What did we do?

We searched for studies that looked at physical measures to stop people catching a respiratory virus infection.

We were interested in how many people in the studies caught a respiratory virus infection, and whether the physical measures had any unwanted effects.

Search date: This is an update of a review first published in 2007. We included evidence published up to 1 April 2020.

What we found

We identified 67 relevant studies. They took place in low-, middle-, and high-income countries worldwide: in hospitals, schools, homes, offices, childcare centres, and communities during non-epidemic influenza periods, the global H1N1 influenza pandemic in 2009, and epidemic influenza seasons up to 2016. No studies were conducted during the COVID-19 pandemic. We identified six ongoing, unpublished studies; three of them evaluate masks in COVID-19.

One study looked at quarantine, and none eye protection, gowns and gloves, or screening people when they entered a country.

We assessed the effects of:

- medical or surgical masks;
- N95/P2 respirators (close-fitting masks that filter the air breathed in, more commonly used by healthcare workers than the general public); and
- hand hygiene (hand-washing and using hand sanitiser).

What are the results of the review?

Medical or surgical masks

Seven studies took place in the community, and two studies in healthcare workers. Compared with wearing no mask, wearing a mask may make little to no difference in how many people caught a flu-like illness (9 studies; 3507 people); and probably makes no difference in how many people have flu confirmed by a laboratory test (6 studies; 3005 people). Unwanted effects were rarely reported, but included discomfort.

N95/P2 respirators

Four studies were in healthcare workers, and one small study was in the community. Compared with wearing medical or surgical masks, wearing N95/P2 respirators probably makes little to no difference in how many people have confirmed flu (5 studies; 8407 people); and may make little to no difference in how many people catch a flu-like illness (5 studies; 8407 people) or respiratory illness (3 studies; 7799 people). Unwanted effects were not well reported; discomfort was mentioned.

Hand hygiene

Following a hand hygiene programme may reduce the number of people who catch a respiratory or flu-like illness, or have confirmed flu, compared with people not following such a programme (16 studies; 61,372 people). Few studies measured unwanted effects; skin irritation in people using hand sanitiser was mentioned.

How reliable are these results?

Our confidence in these results is generally low for the subjective outcomes related to respiratory illness, but moderate for the more precisely defined laboratory-confirmed respiratory virus infection, related to masks and N95/P2 respirators. The results might change when further evidence becomes available. Relatively low numbers of people followed the guidance about wearing masks or about hand hygiene, which may have affected the results of the studies.

Key messages

We are uncertain whether wearing masks or N95/P2 respirators helps to slow the spread of respiratory viruses.

Hand hygiene programmes may help to slow the spread of respiratory viruses.

SUMMARY OF FINDINGS

Summary of findings 1. Medical/surgical masks compared to no masks for preventing the spread of viral respiratory illness

Randomised studies: medical/surgical masks compared to no masks for preventing the spread of viral respiratory illness

Patient or population: general population and healthcare workers
Setting: community and hospitals
Intervention: medical/surgical masks
Comparison: no masks

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	Nº of participants (studies)	Certainty of the evidence (GRADE)	Comments
	Risk with no masks	Risk with randomised studies: masks				
Viral illness - influenza-like illness	Study population		RR 0.99 (0.82 to 1.18)	3507 (9 RCTs)	⊕⊕⊕⊕ LOW ^{a,b}	
	160 per 1000	158 per 1000 (131 to 189)				
Viral illness - laboratory-confirmed influenza	Study population		RR 0.91 (0.66 to 1.26)	3005 (6 RCTs)	⊕⊕⊕⊕ MODERATE ^b	
	40 per 1000	36 per 1000 (26 to 50)				
Influenza-like illness in healthcare workers	Study population		RR 0.37 (0.05 to 2.50)	1070 (2 RCTs)	⊕⊕⊕⊕ LOW ^{a,b}	Studies in healthcare workers only
	40 per 1000	15 per 1000 (2 to 100)				
Adverse events	-		-	(3 RCTs)	⊕⊕⊕⊕ VERY LOW ^{a,c}	Adverse events were not reported consistently and could not be meta-analysed. Adverse events reported for masks included warmth, discomfort, respiratory difficulties, humidity, pain, and shortness of breath, in up to 45% of participants.

***The risk in the intervention group** (and its 95% confidence interval) is based on the median observed risk in the comparison group of included studies and the **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; **RCT:** randomised controlled trial; **RR:** risk ratio

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect.

Moderate certainty: We are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: Our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low certainty: We have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

^aStudy limitations (lack of blinding).

^bImprecision (wide confidence intervals).

^cImprecision: 2 steps (only 3 studies enumerated adverse events; another study mentioned no adverse events).

Summary of findings 2. N95 respirators compared to medical/surgical masks for preventing the spread of viral respiratory illness

Randomised studies: N95 respirators compared to medical/surgical masks for preventing the spread of viral respiratory illness

Patient or population: healthcare workers and general population

Setting: hospitals and households

Intervention: N95 masks

Comparison: medical/surgical masks

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	N° of participants (studies)	Certainty of the evidence (GRADE)	Comments
	Risk with medical masks	Risk with randomised studies: N95				
Viral illness - clinical respiratory illness	Study population		RR 0.70 (0.45 to 1.10)	7799 (3 RCTs)	⊕⊕⊕⊕ VERY LOW ^{a,b,c}	All studies were conducted in hospital settings with healthcare workers.
	120 per 1000	84 per 1000 (54 to 132)				
Viral illness - influenza-like illness	Study population		RR 0.82 (0.66 to 1.03)	8407 (5 RCTs)	⊕⊕⊕⊕ LOW ^{a,b}	1 study was conducted in households (MacIntyre 2009).
	50 per 1000	41 per 1000 (33 to 52)				
Viral illness - laboratory-confirmed influenza	Study population		RR 1.10 (0.90 to 1.34)	8407 (5 RCTs)	⊕⊕⊕⊕ MODERATE ^b	1 study was conducted in households (MacIntyre 2009).
	70 per 1000	77 per 1000 (63 to 94)				

Adverse events	-	-	(5 RCTs)	⊕⊕⊕⊕ VERY LOW ^{a,b,c}	There was insufficient consistent reporting of adverse events to enable meta-analysis. Only 1 study reported detailed adverse events: discomfort was reported in 41.9% of N95 wearers versus 9.8% of medical mask wearers (P < 0.001); headaches were more common with N95 (13.4% versus 3.9%; P < 0.001); difficulty breathing was reported more often in the N95 group (19.4% versus 12.5%; P = 0.01); and N95 caused more problems with pressure on the nose (52.2% versus 11.0%; P < 0.001). 4 RCTs either reported no adverse events or only reported on comfort wearing masks.
----------------	---	---	----------	--------------------------------------	---

***The risk in the intervention group** (and its 95% confidence interval) is based on the median risk in the comparison group and the observed **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; **RCT:** randomised controlled trial; **RR:** risk ratio

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect.

Moderate certainty: We are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: Our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low certainty: We have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

^aStudy limitations (lack of blinding).

^bImprecision (wide confidence interval or no meta-analysis conducted).

^cInconsistency of results (heterogeneity).

Summary of findings 3. Hand hygiene compared to control for preventing the spread of viral respiratory illness

Hand hygiene compared to control for preventing the spread of viral respiratory illness

Patient or population: prevention of spread of viral respiratory illness

Setting: schools, childcare centres, homes, offices

Intervention: hand hygiene

Comparison: control

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	Nº of participants (studies)	Certainty of the evidence (GRADE)	Comments
	Risk with control	Risk with hand hygiene				
Acute respiratory illness	Study population		RR 0.84 (0.82 to 0.86)	44,129 (7 RCTs)	⊕⊕⊕⊕ MODERATE ^a	
	380 per 1000	319 per 1000 (312 to 327)				

Influenza-like illness	Study population		RR 0.98 (0.85 to 1.13)	32,641 (10 RCTs)	⊕⊕⊕⊖ LOW ^{a,b}	
	90 per 1000	88 per 1000 (77 to 102)				
Laboratory-confirmed influenza	Study population		RR 0.91 (0.63 to 1.30)	8332 (8 RCTs)	⊕⊕⊕⊖ LOW ^{b,c}	
	80 per 1000	73 per 1000 (50 to 104)				
Composite of acute respiratory illness, influenza-like illness, influenza	Study population		RR 0.89 (0.84 to 0.95)	61,372 (16 RCTs)	⊕⊕⊕⊖ LOW ^{a,b}	
	200 per 1000	178 per 1000 (168 to 190)				
Adverse events	-		-	(2 RCTs)	⊕⊕⊕⊖ VERY LOW ^{a,b,c}	Data were insufficient to conduct meta-analysis. 1 study reported that no adverse events were observed, and another study reported that skin reaction was recorded for 10.4% of participants in the hand sanitiser group versus 10.3% in the control group.

***The risk in the intervention group** (and its 95% confidence interval) is based on the median observed risk in the comparison groups of included studies and the **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; **RCT:** randomised controlled trial; **RR:** risk ratio

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect.

Moderate certainty: We are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: Our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low certainty: We have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

^aStudy limitation (majority of studies were unblinded, with participant-assessed outcome).

^bInconsistent results across studies.

^cImprecision (wide confidence interval or no meta-analysis conducted).

BACKGROUND

Description of the condition

Epidemic and pandemic viral infections pose a serious threat to people worldwide. Epidemics of note include severe acute respiratory syndrome (SARS) in 2003 and the Middle East respiratory syndrome (MERS), which began in 2012. Major pandemics include the H1N1 influenza caused by the H1N1pdm09 virus in 2009 and the coronavirus disease 2019 (COVID-19) caused by SARS-CoV-2.

Even non-epidemic acute respiratory infections (ARIs) place a huge burden on healthcare systems around the world, and are a prominent cause of morbidity (WHO 2017). Furthermore, ARIs are often antecedents to lower respiratory tract infections caused by bacterial pathogens (i.e. pneumonia), which cause millions of deaths worldwide, mostly in low-income countries (Schwartz 2018).

High viral load, high levels of transmissibility, susceptible populations, and symptomatic patients are considered to be the drivers of such epidemics and pandemics (Jefferson 2006a). Preventing the spread of respiratory viruses from person to person may be effective at reducing the spread of outbreaks. Physical interventions, such as the use of masks and physical distancing measures, might prevent the spread of respiratory viruses which are transmitted by large droplets from infected to susceptible people. This review assumes that physical interventions used to prevent transmission of respiratory viruses are similar for most viral ARIs.

Description of the intervention

Single measures of intervention (Demicheli 2018a; Demicheli 2018b; Jefferson 2014; Jefferson 2018; Thomas 2010), such as the use of vaccines or antivirals, may be insufficient to contain the spread of influenza, but combinations of interventions may reduce the reproduction number to below 1. For some respiratory viruses there are no licensed interventions, and a combination of social and physical interventions may be the only option to reduce the spread of outbreaks, particularly those that may be capable of becoming epidemic or pandemic in nature (Luby 2005). Such interventions were emphasised in the World Health Organization's latest Global Influenza Strategy 2019 to 2030, and have several possible advantages over other methods of suppressing ARI outbreaks since they may be instituted rapidly and may be independent of any specific type of infective agent, including novel viruses. In addition, the possible effectiveness of public health measures during the Spanish flu pandemic of 1918 to 1919 in US cities supports the impetus to investigate the existing evidence on the effectiveness of such interventions (Bootsma 2007), including quarantine (such as isolation, physical distancing) and the use of disinfectants. We also considered the major societal implications for any community adopting these measures (CDC 2005a; CDC 2005b; WHO 2006b; WHO 2020a; WHO 2020b).

How the intervention might work

Epidemics and pandemics are more likely during antigenic change (changes in the viral composition) in the virus or transmission from animals (domestic or wild) when there is no natural human immunity (Bonn 1997). High viral load, high levels of

transmissibility, and symptomatic patients are considered to be the drivers of such epidemics and pandemics (Jefferson 2006b).

Physical interventions, such as the use of masks, physical distancing measures, school closures, and limitations of mass gatherings, might prevent the spread of the virus transmitted by large droplets or aerosols from infected to susceptible individuals. The use of hand hygiene, gloves, and protective gowns can also prevent the spread by limiting the transfer of viral particles onto and from fomites (inanimate objects such as flat surfaces, tabletops, utensils, porous surfaces, or nowadays cell phones, which can transmit the agent if contaminated). Such public health measures were widely adopted during the Spanish flu pandemic and have been the source of considerable debate (Bootsma 2007).

Why it is important to do this review

Although the benefits of physical interventions seem self-evident, given the global importance of interrupting viral transmission, having up-to-date estimates of their effectiveness is necessary to inform planning, decision-making, and policy. The outbreak of COVID-19 has prompted this update. Physical methods have several possible advantages over other methods of suppressing ARI outbreaks, including their rapid deployment and ability to be independent of the infective agent, including novel viruses.

The last update of this review in 2011, Jefferson 2011, identified 23 trials on physical interventions that might interrupt or reduce the spread of respiratory viruses. Because of poor reporting and heterogeneity, and the relatively small number of included trials, it was not possible to perform a meta-analysis. Case-control studies were sufficiently homogenous to permit meta-analysis, which provided evidence that hand-washing for a minimum of 11 times daily prevented cases of SARS during the 2003 epidemic (odds ratio 0.54, 95% confidence interval 0.44 to 0.67). Many randomised trials have been published in the past decade, prompting us to focus only on these for the current update.

This is the fourth update of a Cochrane Review first published in 2007 (Jefferson 2007; Jefferson 2009; Jefferson 2010; Jefferson 2011).

OBJECTIVES

To assess the effectiveness of physical interventions to interrupt or reduce the spread of acute respiratory viruses.

METHODS

Criteria for considering studies for this review

Types of studies

For this 2020 update we only considered individual-level RCTs, or cluster-RCTs, or quasi-RCTs for inclusion.

In previous versions of the review we also included observational studies (cohorts, case-controls, before-after, and time series studies). However, for this update there were sufficient randomised studies to address our study aims, so we excluded observational studies (which are known to be at a higher risk of bias).

Types of participants

People of all ages.

Types of interventions

We included randomised controlled trials (RCTs) and cluster-RCTs of trials investigating physical interventions (screening at entry ports, isolation, quarantine, physical distancing, personal protection, hand hygiene, face masks, and gargling) to prevent respiratory virus transmission compared with doing nothing or with another intervention.

Types of outcome measures

For this 2020 update we added one outcome: adverse events related to the intervention, and we split the outcomes into primary and secondary outcomes.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, influenza-like illness (ILI), and laboratory-confirmed influenza, or other viral pathogens).
2. Adverse events related to the intervention.

Secondary outcomes

1. Deaths.
2. Severity of viral illness as reported in the studies.
3. Absenteeism.
4. Hospital admissions.
5. Complications related to the illness, e.g. pneumonia.

Search methods for identification of studies

Electronic searches

For this 2020 update, we refined the original search strategy using a combination of previously included studies and automation tools (Clark 2020). We converted this search using the Polyglot Search Translator (Clark 2020), and ran the searches in the following databases:

1. the Cochrane Central Register of Controlled Trials (CENTRAL) (2020, Issue 3), which includes the Acute Respiratory Infections Group's Specialised Register (searched 1 April 2020) (Appendix 1);
2. PubMed (2010 to 1 April 2020) (Appendix 2);
3. Embase (2010 to 1 April 2020) (Appendix 3);
4. CINAHL (Cumulative Index to Nursing and Allied Health Literature) (2010 to 1 April 2020) (Appendix 4);
5. US National Institutes of Health Ongoing Trials Register ClinicalTrials.gov (January 2010 to 16 March 2020); and
6. World Health Organization International Clinical Trials Registry Platform (January 2010 to 16 March 2020).

We combined the database searches with the Cochrane Highly Sensitive Search Strategy for identifying randomised trials in MEDLINE: sensitivity- and precision-maximising version (2008 revision) (Lefebvre 2011). Details of previous searches are available in Appendix 5.

Searching other resources

We conducted a backwards-and-forwards citation analysis in Scopus on all newly included studies to identify other potentially relevant studies.

Data collection and analysis

Selection of studies

The search and citation analysis results were initially screened via the RobotSearch tool (Marshall 2018) to exclude all studies that were obviously not RCTs. We scanned the titles and abstracts of studies identified by the searches. We obtained the full-text articles of studies that either appeared to meet our eligibility criteria or for which there was insufficient information to exclude it. We then used a standardised form to assess the eligibility of each study based on the full article.

Data extraction and management

Three pairs of review authors (MJ/EF, LA/GB, EB/TOJ) independently applied the inclusion criteria to all identified and retrieved articles, and extracted data using a standard template that had been developed for and applied to previous versions of the review, but was revised to reflect our focus on RCTs and cluster-RCTs for this update. Any disagreements were resolved through discussion. We extracted and reported descriptions of interventions using the Template for Intervention Description and Replication (TIDieR) template (Table 1).

Assessment of risk of bias in included studies

Three pairs of review authors (TOJ/EB, LA/GB, MJ/EF) independently assessed risk of bias for the method of random sequence generation and allocation concealment (selection bias), blinding of participants and personnel (performance bias), blinding of outcome assessment (detection bias), outcome reporting (attrition bias), and selective reporting (reporting bias). We used the Cochrane 'Risk of bias' tool to assess risk of bias, classifying each 'Risk of bias' domain as 'low', 'high', or 'unclear'. The following were indications for low risk of bias:

1. method of random sequence generation: the method was well-described and is likely to produce balanced and truly random groups;
2. allocation concealment: the next treatment allocation was not known to participant/cluster or treating staff until after consent to join the study;
3. blinding of participants and personnel: the method is likely to maintain blinding throughout the study;
4. blinding of outcome assessors: all outcome assessors were unaware of treatment allocation;
5. outcome reporting: participant attrition throughout the study is reported, and reasons for loss are appropriately described; and
6. selective reporting: all likely planned and collected outcomes have been reported.

Measures of treatment effect

When possible, we performed meta-analysis and summarised effectiveness as risk ratio (RR) using 95% confidence intervals (CIs). For studies that could not be pooled, we used the effect measures reported by the trial authors (such as RR or incidence rate ratio (IRR) with 95% CI or, when these were not available, relevant P values).

Unit of analysis issues

Many of the included studies were cluster-RCTs. To avoid any unit of analysis issues, we only included treatment effect estimates that were based on methods that were appropriate for the analysis of

cluster trials, such as mixed models and generalised estimating equations. Given this restriction, we used the generalised inverse-variance method of meta-analysis. Some cluster-RCTs that did not report cluster-adjusted treatment effects provided sufficient data (number of events and participants by treatment group and intraclass correlations) for us to calculate appropriate treatment effect estimates and standard errors. For studies with multiple treatment groups but only one control group, where appropriate, we adjusted standard errors upwards to avoid unit of analysis errors in the meta-analyses.

Dealing with missing data

Given the urgency of this update, we did not contact authors of studies with significant missing data. Previously, whenever details of studies were unclear, or studies were only known to us by abstracts or communications at meetings, we corresponded with first or corresponding authors.

Assessment of heterogeneity

Aggregation of data was dependent on types of comparisons, sensitivity and homogeneity of definitions of exposure, populations and outcomes used. We calculated the I^2 statistic and Chi^2 test for each pooled estimate to assess the presence of statistical heterogeneity (Higgins 2002; Higgins 2003).

Assessment of reporting biases

Given the widely disparate nature of our evidence base, we limited our assessment of possible reporting biases to funnel plot visual inspection if we had > 10 included studies.

Data synthesis

If possible and appropriate, we combined studies in a meta-analysis. We used the generalised inverse-variance random-effects model. We chose the random-effects model because we expected clinical heterogeneity due to differences in pooled interventions and outcome definitions, and methodological heterogeneity due to pooling of RCTs and cluster-RCTs.

Subgroup analysis and investigation of heterogeneity

We conducted two post hoc subgroup analyses:

1. healthcare workers for the comparison of masks versus control; and

2. children for the comparison of hand hygiene versus control.

We did not conduct further investigation of heterogeneity due to insufficient numbers of studies included in the comparisons.

Sensitivity analysis

We conducted a sensitivity analysis for hand hygiene versus control where we included the most precise and unequivocal measure of viral illness reported for each included study.

Summary of findings and assessment of the certainty of the evidence

We created three 'Summary of findings' tables using the following outcomes: numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza) and adverse events related to the intervention ([Summary of findings 1](#); [Summary of findings 2](#); [Summary of findings 3](#)). We planned to include the secondary outcomes of deaths; severity of viral illness as reported in the studies; absenteeism; hospital admissions; and complications related to the illness (e.g. pneumonia). However, this data were poorly reported in the included studies. We used the five GRADE considerations (study limitations, consistency of effect, imprecision, indirectness, and publication bias) to assess the certainty of evidence as it related to the studies which contributed data to the meta-analyses for the prespecified outcomes (Atkins 2004). We used the methods and recommendations described in Section 8.5 and Chapter 12 of the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011), employing GRADEpro GDT software (GRADEpro GDT 2015). We justified all decisions to down- or upgrade the certainty of the evidence in footnotes, and made comments to aid the reader's understanding of the review where necessary.

RESULTS

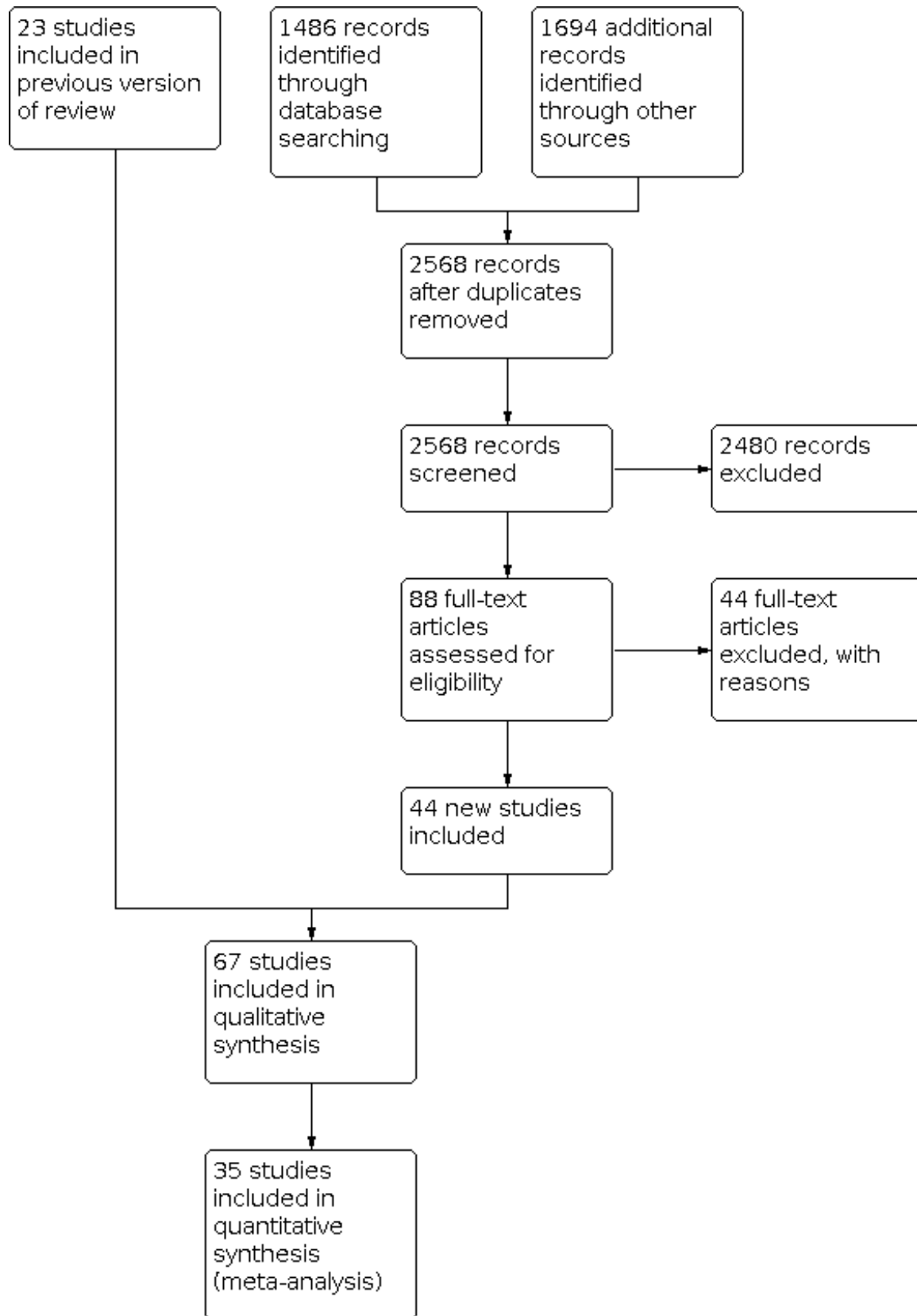
Description of studies

See [Characteristics of included studies](#) and [Characteristics of excluded studies](#) tables.

Results of the search

We identified a total of 3180 titles in this 2020 update. We excluded 3092 titles and retrieved the full papers of 88 studies, to include 44 new studies. See [Figure 1](#).

Figure 1. Study flow diagram.



Included studies

The 44 newly included studies were all RCTs ($n = 11$) or cluster-RCTs ($n = 33$) published between 2010 and 2019. We included 23 RCTs in the 2011 version of the review. For detailed descriptions of the interventions of the included studies, see [Table 1](#).

Fifteen trials focused on using masks ([Aiello 2010](#); [Aiello 2012](#); [Barasheed 2014](#); [Canini 2010](#); [Cowling 2008](#); [Ide 2016](#); [Jacobs 2009](#); [Loeb 2009](#); [MacIntyre 2009](#); [MacIntyre 2011](#); [MacIntyre 2013](#); [MacIntyre 2015](#); [MacIntyre 2016](#); [Radonovich 2019](#); [Suess 2012](#)). Ten of the 15 trials compared medical/surgical masks to no mask (control) ([Aiello 2010](#); [Aiello 2012](#); [Barasheed 2014](#); [Canini 2010](#); [Cowling 2008](#); [Jacobs 2009](#); [MacIntyre 2009](#); [MacIntyre 2015](#); [MacIntyre 2016](#); [Suess 2012](#)). One study compared catechin-treated masks to no mask ([Ide 2016](#)), and one study included cloth masks versus control (third arm in [MacIntyre 2015](#)). Three of the 15 trials were in healthcare workers ([Ide 2016](#); [Jacobs 2009](#); [MacIntyre 2015](#)), whilst the remaining trials were in non-healthcare workers (students, households, families, or pilgrims). Only one trial was conducted during H1N1 pandemic season ([Suess 2012](#)).

Five of the 15 trials compared N95 masks or P2 masks to medical/surgical masks ([Loeb 2009](#); [MacIntyre 2009](#); [MacIntyre 2011](#); [MacIntyre 2013](#); [Radonovich 2019](#)). All of these trials, except for one study that was conducted on household individuals ([MacIntyre 2009](#)), included healthcare workers either in a hospital setting, [Loeb 2009](#); [MacIntyre 2011](#); [MacIntyre 2013](#), or an outpatient setting ([MacIntyre 2009](#); [Radonovich 2019](#)).

One trial evaluated the effectiveness of quarantining workers of one of two sibling companies in Japan whose family members had developed an ILI during the 2009 to 2010 H1N1 influenza pandemic ([Miyaki 2011](#)).

Fifteen trials compared hand hygiene interventions with no hand hygiene (control) and provided data suitable for meta-analysis. The populations in these trials included adults, children, and families, in settings such as schools ([Biswas 2019](#); [Stebbins 2011](#)), childcare centres ([Azor-Martinez 2018](#); [Correa 2012](#); [Roberts 2000](#); [Zomer 2015](#)), homes/households ([Cowling 2009](#); [Larson 2010](#); [Little 2015](#); [Nicholson 2014](#); [Ram 2015](#); [Sandora 2005](#); [Simmerman 2011](#)), offices ([Hubner 2010](#)), and military trainees ([Millar 2016](#)). None of the trials was conducted during a pandemic, although some of the studies were conducted during peak influenza seasons.

A further 10 trials that compared a variety of hand hygiene modalities to control provided insufficient information to include in meta-analyses. Three trials were in children: one was conducted in daycare centres in Denmark examining a multimodal hygiene programme ([Ladegaard 1999](#)), and two trials compared a hand hygiene campaign or workshop in an elementary school environment in Saudi Arabia, [Alzaher 2018](#), and Egypt, [Talaat 2011](#). Three trials tested virucidal hand treatment in an experimental setting, [Gwaltney 1980](#); [Turner 2004a](#), and in a community, [Turner 2012](#), in the USA. [Feldman 2016](#) compared hand-washing with chlorhexidine gluconate amongst Israeli sailors. One trial compared hand sanitiser packaged in a multimodal hygiene programme amongst office employees in the USA ([Arbogast 2016](#)). Two trials were conducted in a long-term facility setting: one trial examined the effect of a bundle hand hygiene programme on infectious risk in nursing home residents in France ([Temime 2018](#)), and the other trial compared the effect of using hand sanitisers in

healthcare workers on the rate of infections (including respiratory infections) in nursing home residents in Hong Kong ([Yeung 2011](#)).

Five trials compared different hand hygiene interventions in a variety of settings such as schools ([Morton 2004](#) in kindergartens and elementary schools in the USA; [Priest 2014](#) in primary schools in New Zealand; and [Pandejpong 2012](#) in kindergartens in Thailand). One study was conducted in low-income neighbourhoods in Karachi, Pakistan ([Luby 2005](#)), and one was conducted in a workplace environment in Finland ([Savolainen-Kopra 2012](#)). A variety of interventions were used across these trials such as soap and water ([Luby 2005](#); [Savolainen-Kopra 2012](#)), hand sanitiser ([Morton 2004](#); [Pandejpong 2012](#); [Priest 2014](#); [Savolainen-Kopra 2012](#)), body wash ([Luby 2005](#)), and alcohol-based hand wipes ([Morton 2004](#)), with or without additional hygiene education. There was considerable variation in interventions, and the information in the trial reports was insufficient to permit meta-analysis.

Seven trials compared a combined intervention of hand hygiene and face masks with control. Four of these trials were carried out in households in Germany ([Suess 2012](#)), Thailand ([Simmerman 2011](#)), Hispanic immigrant communities in the USA ([Larson 2010](#)), and households in Hong Kong ([Cowling 2009](#)). Two trials were conducted amongst university student residences ([Aiello 2010](#); [Aiello 2012](#)), and one trial in a group of pilgrims at the annual Hajj ([Aelami 2015](#)). Moreover, six trials evaluated the incremental benefit of combining surgical mask in addition to hand hygiene with soap, [Simmerman 2011](#), hand sanitiser, [Aiello 2010](#); [Aiello 2012](#); [Larson 2010](#); [Suess 2012](#), or both, [Cowling 2009](#), versus mask or hand hygiene alone on the outcomes of ILI and influenza. [Aelami 2015](#) investigated a hygienic package (alcohol-based handrub (gel or spray), surgical masks, soap, and paper handkerchiefs) with a control group.

Seven trials compared a multimodal combination of hand hygiene and disinfection of surfaces, toys, linen, or other components of the environment with a control ([Ban 2015](#); [Carabin 1999](#); [Ibfelt 2015](#); [Kotch 1994](#); [McConeghy 2017](#); [Sandora 2008](#); [White 2001](#)). Variation in scope and type of interventions and insufficient data in trial reports precluded meta-analysis. All studies except for one were in children ([McConeghy 2017](#) was in nursing population).

Three trials included in two papers investigated the role of virucidal tissues in interrupting transmission of naturally occurring respiratory infections in households ([Farr 1988a](#); [Farr 1988b](#); [Longini 1988](#)). Four cluster-RCTs implemented complex, multimodal sanitation, education, cooking, and hygiene interventions ([Chard 2019](#); [Hartinger 2016](#); [Huda 2012](#); [Najnin 2019](#)). All four of these trials were conducted in low-income countries in settings with minimal to no access to basic sanitation.

Three trials assessed the effect of gargling on the incidence of upper respiratory tract infections (URTIs) or influenza: gargling with povidone-iodine ([Satomura 2005](#)), green tea ([Ide 2014](#)), and tap water ([Goodall 2014](#)).

Ongoing studies

We identified six ongoing studies. Two assess hand hygiene measures ([NCT03454009](#); [NCT04267952](#)), and four assess face masks ([NCT04471766](#); [NCT04296643](#); [NCT04337541](#); [Wang 2015](#)) one of which – [NCT04337541](#) – published as this review update was going to press.

Excluded studies

We excluded a total of 160 studies. We identified 12 new studies for exclusion at the data extraction stage of this 2020 update, all of which appeared to be eligible at screening. Six of the 12 studies were ineligible due to only reporting composite outcomes that included other infections besides those caused by respiratory viruses (Azor-Martinez 2014; Bowen 2007; Chami 2012; Denbak 2018; Stedman-Smith 2015; Vessey 2007); two trials measured absenteeism due to non-specific infection (Lennell 2008; Rosen

2006); one trial only had two clusters (Nandrup-Bus 2009); one study was not an RCT (Patel 2012); one study evaluated a hand hygiene intervention that was antibacterial rather than antiviral (Slayton 2016); and one study had no respiratory illness data for extraction (Uhari 1999).

Risk of bias in included studies

The overall risk of bias is presented graphically in Figure 2 and summarised in Figure 3. Details on risk of bias for the included studies are provided below.

Figure 2. 'Risk of bias' graph: review authors' judgements about each risk of bias item presented as percentages across all included trials.

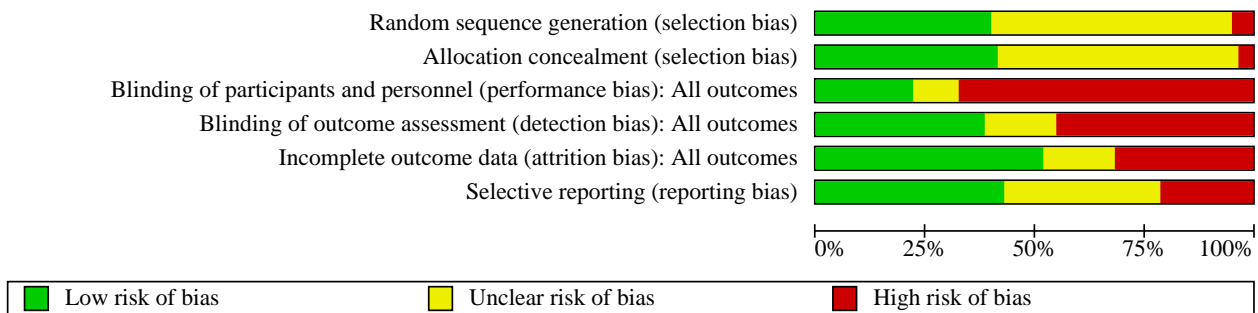


Figure 3. 'Risk of bias' summary: review authors' judgements about each risk of bias item for each included trial.

	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of participants and personnel (performance bias): All outcomes	Blinding of outcome assessment (detection bias): All outcomes	Incomplete outcome data (attrition bias): All outcomes	Selective reporting (reporting bias)
Aelami 2015	?	?	-	?	?	?
Aiello 2010	?	-	-	+	+	-
Aiello 2012	+	+	-	+	+	+
Alzaher 2018	?	+	-	-	+	?
Arbogast 2016	?	?	-	-	+	?
Azor-Martinez 2016	+	+	-	-	-	?
Azor-Martinez 2018	+	+	-	-	+	?
Ban 2015	-	?	-	-	-	-
Barasheed 2014	?	?	+	?	+	+
Biswas 2019	+	+	-	-	-	?
Canini 2010	+	+	-	+	+	+
Carabin 1999	?	?	-	-	-	-
Chard 2019	?	+	-	-	+	+
Correa 2012	+	?	-	-	+	?
Cowling 2008	+	+	-	-	-	-
Cowling 2009	+	+	-	?	-	?
DiVita 2011	?	?	?	?	?	?
Farr 1988a	?	?	+	+	-	+
Farr 1988b	?	?	+	+	-	+
Feldman 2016	?	?	-	?	?	?
Goodall 2014	?	+	+	+	+	+
Gwaltney 1980	?	?	+	?	?	?
Hartinger 2016	?	?	-	-	+	+

Figure 3. (Continued)

Gwaltney 1980	?	?	+	?	?	?
Hartinger 2016	?	?	-	-	+	+
Hubner 2010	?	?	-	-	+	?
Huda 2012	?	?	-	-	-	?
Ibfelt 2015	?	?	-	+	?	+
Ide 2014	+	+	-	-	+	?
Ide 2016	?	+	+	+	+	+
Jacobs 2009	?	?	-	-	+	-
Kotch 1994	?	?	-	-	-	-
Ladegaard 1999	?	?	-	-	-	-
Larson 2010	?	?	-	?	-	?
Little 2015	?	+	-	-	-	+
Loeb 2009	?	+	-	+	+	+
Longini 1988	?	+	+	+	?	-
Luby 2005	+	+	+	+	?	+
MacIntyre 2009	?	?	-	+	+	+
MacIntyre 2011	?	+	-	-	+	+
MacIntyre 2013	?	?	+	+	+	+
MacIntyre 2015	+	+	-	-	+	+
MacIntyre 2016	+	-	-	-	+	+
McConeghy 2017	?	?	-	-	?	-
Millar 2016	+	?	-	+	-	-
Miyaki 2011	?	?	+	+	+	?
Morton 2004	?	?	?	?	?	?
Najnin 2019	+	?	-	-	-	-
Nicholson 2014	-	+	-	-	-	?
Pandejpong 2012	?	?	?	+	+	+
Priest 2014	+	+	+	+	?	+
Radonovich 2019	+	+	+	+	+	+
Ram 2015	+	+	-	-	+	+
Roberts 2000	+	?	-	+	?	+
Sandora 2005	+	+	-	-	+	?
Sandora 2008	+	?	-	+	+	?
Satomura 2005	+	+	-	-	+	?
Savolainen-Kopra 2012	?	+	-	-	-	+
Simmerman 2011	+	?	+	+	+	+
Stebbins 2011	+	+	-	+	-	?
Suess 2012	+	+	?	+	+	+
Talaat 2011	+	?	?	?	-	?
Temime 2018	-	?	-	-	-	+
Turner 2004a	?	?	?	?	+	-
Turner 2004b	?	?	?	?	+	-
Turner 2012	+	+	+	+	+	+
White 2001	?	?	+	+	-	-
Yeung 2011	?	?	-	-	+	?
Zomer 2015	+	?	-	-	+	+

Allocation

For this 2020 review, information on sequence generation was overall poorly reported in most of the newly included studies. Twenty-one newly included studies provided adequate information on randomisation scheme and were judged as at low risk of bias (Aiello 2012; Azor-Martinez 2016; Azor-Martinez 2018; Biswas 2019; Canini 2010; Correa 2012; Goodall 2014; Ide 2014; MacIntyre 2015; MacIntyre 2016; Millar 2016; Najnin 2019; Priest 2014; Radonovich 2019; Ram 2015; Simmerman 2011; Stebbins 2011; Suess 2012; Talaat 2011; Turner 2012; Zomer 2015). Nine studies described the use of computerised sequence generation program/software (Aiello 2012; Azor-Martinez 2018; Biswas 2019; Canini 2010; Millar 2016; Najnin 2019; Radonovich 2019; Talaat 2011; Turner 2012). One study used random number tables for sequence generation (Azor-Martinez 2016). Three studies described using the random function in Microsoft Excel (Correa 2012; MacIntyre 2016; Suess 2012). Two studies used statistical software to generate a randomisation allocation (MacIntyre 2015; Priest 2014). Two studies reported using block randomisation: Ram 2015 used block randomisation, and an independent investigator generated the list of random assignments, whilst Simmerman 2011 performed block randomisation. Stebbins 2011 used constrained randomisation, and Zomer 2015 reported using stratified randomisation by means of computer generation with a 1:1 ratio in each of the strata.

Fourteen studies reported insufficient information to permit a judgement on the adequacy of the process to minimise selection bias (Aelami 2015; Alzاهر 2018; Arbogast 2016; Barasheed 2014; Chard 2019; DiVita 2011; Feldman 2016; Hubner 2010; Ibfelt 2015; McConeghy 2017; Miyaki 2011; Pandepong 2012; Savolainen-Kopra 2012; Yeung 2011). Six studies provided some description about sequence generation, but it was still unclear (Hartinger 2016; Huda 2012; Ide 2016; Little 2015; MacIntyre 2011; MacIntyre 2013). Huda 2012 mentioned random number tables, but it was unclear if this was for random selection or randomisation. Ide 2016 used computer-generated randomisation, but the method was not stated. Hartinger 2016 used covariate-constrained randomisation, but the method was not described. In Little 2015, participants were automatically randomly assigned by the intervention software, but the sequence generation was not described. Two studies used a secure computerised randomisation program (MacIntyre 2011; MacIntyre 2013), but the sequence generation was not described.

Three of the newly included studies were poorly randomised (Ban 2015; Nicholson 2014; Temime 2018). Ban 2015 included only two clusters, and the randomisation scheme was not reported. Nicholson 2014 used coin tossing, which can lead to a large imbalance. Temime 2018 used "simple randomisation" with no further description.

For the RCTs included in previous versions of the review, three were poorly reported with no description of randomisation sequence or concealment of allocation (Gwaltney 1980; Turner 2004a; Turner 2004b). The quality of the cluster-RCTs varied, with four studies not providing a description of the randomisation procedure (Carabin 1999; Kotch 1994; Morton 2004; White 2001). We rated seven studies as at low risk of bias for sequence generation (Cowling 2008; Cowling 2009; Luby 2005; Roberts 2000; Sandora 2005; Sandora 2008; Satomura 2005), and a further six studies as at unclear risk of bias (Farr 1988a; Farr 1988b; Ladegaard 1999; Loeb 2009; Longini 1988; MacIntyre 2009).

Many of the newly included cluster-RCTs did not report adequately on allocation concealment. Twenty-one of these studies reported adequate allocation and were judged as at low risk of bias (Aiello 2012; Alzاهر 2018; Azor-Martinez 2016; Azor-Martinez 2018; Biswas 2019; Canini 2010; Chard 2019; Goodall 2014; Ide 2014; Ide 2016; Little 2015; MacIntyre 2011; MacIntyre 2015; Nicholson 2014; Priest 2014; Radonovich 2019; Ram 2015; Savolainen-Kopra 2012; Stebbins 2011; Suess 2012; Turner 2012). Aiello 2012 randomised all residence houses in each of the residence halls prior to the intervention implementation. Alzاهر 2018 allocated schools prior to all schoolgirls attending selected schools being invited to participate. Azor-Martinez 2016 allocated schools/classes prior to children's recruitment. Azor-Martinez 2018 assigned clusters prior to recruitment. Biswas 2019 completed the allocation prior to individuals being recruited. Chard 2019 allocated schools prior to individuals being recruited. Goodall 2014 used opaque, sealed, serially numbered envelopes that were only accessed when two study personnel were present. Ide 2014 also reported using individual drawing of sealed, opaque envelopes to randomly assign participants to the study groups. MacIntyre 2011 randomised hospitals prior to inclusion of participants. In MacIntyre 2015, hospital wards were randomised prior to recruitment of individuals. Nicholson 2014 used coin tossing to assign communities to intervention or control arms. Radonovich 2019 used constrained randomisation to resolve any potential imbalance between covariates between the trial arms. Four studies reported the use of central randomisation: Canini 2010 used central randomisation employing an interactive voice response system; Ide 2016 used central randomisation services; in Little 2015 participants were automatically randomly assigned by the intervention software; and Ram 2015 described a central allocation through data collectors notifying the field research officer, who consulted the block randomisation list to make the assignment of the household compound to intervention or control. Savolainen-Kopra 2012 randomised clusters by matching prior to the onset of the interventions. Four studies reported that allocation was assigned by personnel (investigator, physician, or statistician) unaware of the randomisation sequence (Priest 2014; Stebbins 2011; Suess 2012; Turner 2012). Twenty-two studies reported insufficient information to permit a judgement on the adequacy of the process to minimise selection bias (Aelami 2015; Arbogast 2016; Ban 2015; Barasheed 2014; Correa 2012; DiVita 2011; Feldman 2016; Hartinger 2016; Hubner 2010; Huda 2012; Ibfelt 2015; MacIntyre 2013; McConeghy 2017; Millar 2016; Miyaki 2011; Najnin 2019; Pandepong 2012; Simmerman 2011; Talaat 2011; Temime 2018; Yeung 2011; Zomer 2015). Two studies provided some information about allocation, but it was not enough to permit a judgement on risk of bias (Barasheed 2014; Simmerman 2011). Barasheed 2014 randomised pilgrim tents using an independent study co-ordinator who was not an investigator, but did not describe how this was done. Simmerman 2011 described using a study co-ordinator to assign households to study arm (after consent was obtained). Only one of the newly added studies was judged as at high risk of bias, where random assignment was allocated by doctors enrolling the participants (MacIntyre 2016). Of the previously included RCTs, 14 provided no or an insufficient description of concealment of allocation (Carabin 1999; Farr 1988a; Farr 1988b; Gwaltney 1980; Kotch 1994; Ladegaard 1999; Larson 2010; MacIntyre 2009; Morton 2004; Roberts 2000; Sandora 2008; Turner 2004a; Turner 2004b; White 2001). We assessed all of the remaining studies as at low risk of bias (Canini 2010; Cowling 2008; Cowling 2009; Loeb 2009; Longini 1988; Luby 2005; Sandora

2005; Satomura 2005). Aiello 2010 used the drawing of a uniform ticket with the name of each hall out of a container and was rated as at high risk of bias.

Blinding

Although blinding is less of a concern in cluster-RCTs, the risk of bias is substantial when the outcomes are subjective and the outcome assessor is not blinded. We judged 26 studies to have a high risk of bias (Aiello 2012; Alzahr 2018; Arbogast 2016; Azor-Martinez 2016; Azor-Martinez 2018; Ban 2015; Biswas 2019; Carabin 1999; Chard 2019; Correa 2012; Cowling 2008; Ide 2014; Kotch 1994; Ladegaard 1999; Little 2015; MacIntyre 2011; MacIntyre 2015; MacIntyre 2016; McConeghy 2017; Najnin 2019; Nicholson 2014; Ram 2015; Sandora 2008; Savolainen-Kopra 2012; Temime 2018; Zomer 2015). We assessed five cluster-RCTs as at low risk of bias. Farr 1988a and Farr 1988b were double-blinded studies and were judged as at low risk of bias. MacIntyre 2013 and Simmerman 2011 reported laboratory-confirmed influenza, and blinding would not have affected the result. In Miyaki 2011 the self-reported respiratory symptoms were confirmed by a physician. We judged three cluster-RCTs to have a low risk of detection bias because the outcome was laboratory-confirmed influenza, Barasheed 2014; Suess 2012, or physician-confirmed ILI, Pandejpong 2012. Two cluster-RCTs provided insufficient data to judge the effect of non-blinding. Talaat 2011 included outcomes that were both self-reported ILI and laboratory-confirmed influenza. In Yeung 2011 the detection of cases was based on record for hospitalisation related to infection (including pneumonia). Eleven cluster-RCTs were not blinded, but we judged the primary outcome to be unaffected by non-blinding. Seven trials reported laboratory-confirmed influenza (Aiello 2012; Cowling 2009; Larson 2010; Loeb 2009; MacIntyre 2009; Millar 2016; Stebbins 2011). Four studies reported self-reported outcome (Canini 2010; Priest 2014; Roberts 2000; Sandora 2008), but outcome assessors were not aware of the intervention assignment. Five RCTs were double-blinded and were judged as at low risk of bias (Goodall 2014; Ide 2016; Longini 1988; Luby 2005; White 2001), whilst two studies were single-blinded where investigators, Radonovich 2019, or laboratory personnel, Turner 2012, were blinded. Four RCTs were not blinded and were judged as at high risk of bias given the subjective nature of the outcome assessed (Hubner 2010; Ibfelt 2015; Jacobs 2009; Satomura 2005). Turner 2004a and Turner 2004b were double-blind studies, but insufficient information was provided to assess risk of bias.

Incomplete outcome data

In this 2020 review, we assessed 26 newly included trials as having a low risk of attrition bias, with sufficient evidence from the participant flow chart, and explanation of loss to follow-up (which was minimal) similar between groups (Aiello 2012; Alzahr 2018; Arbogast 2016; Azor-Martinez 2018; Barasheed 2014; Canini 2010; Chard 2019; Correa 2012; Goodall 2014; Hartinger 2016; Hubner 2010; Ide 2014; Ide 2016; MacIntyre 2011; MacIntyre 2013; MacIntyre 2015; MacIntyre 2016; Miyaki 2011; Pandejpong 2012; Radonovich 2019; Ram 2015; Simmerman 2011; Suess 2012; Turner 2012; Yeung 2011; Zomer 2015). Seven studies did not report sufficient information on incomplete data (attrition bias) (Aelami 2015; DiVita 2011; Feldman 2016; Hartinger 2016; Ibfelt 2015; McConeghy 2017; Priest 2014). Twelve studies had a high risk of attrition bias (Azor-Martinez 2016; Ban 2015; Biswas 2019; Huda 2012; Little 2015; Millar 2016; Najnin 2019; Nicholson 2014; Savolainen-Kopra 2012;

Stebbins 2011; Talaat 2011; Temime 2018). In Azor-Martinez 2016, attrition levels were high and differed between the two groups. Ban 2015 did not report on reasons for loss to follow-up. Biswas 2019 did not provide information on missing participants (28 children in the control schools and two children in the intervention schools). Huda 2012 did not provide a flow diagram of study participants. Little 2015 had high attrition that differed between the two groups. Attrition in Millar 2016 differed amongst the three groups. In addition, ARI cases were captured utilising clinic-based medical records for those participants who sought hospital care only. In Najnin 2019, there was high migration movement during the study, which could have distorted the baseline characteristics even more. There was no description of how such migration and changes in the intervention group were dealt with. In Nicholson 2014, households were removed from the study if they provided no data for five consecutive weeks. Although attrition was reported in Savolainen-Kopra 2012, and 76% of volunteers who were recruited at the beginning of the reporting period completed the study, new recruits were added during the study to replace volunteers lost in most clusters. The total number of reporting participants at the end of the trial was 626 (91.7%) compared to the beginning, meaning that 15.7% of participants were replaced during the study. In Stebbins 2011 reasons for episodes of absence in 66% of the study participants were not reported. Talaat 2011 did not provide a flow chart of clusters flow during the study period and provided no information on withdrawal. Temime 2018 was greatly biased due to underreporting of outcomes in the control groups. Furthermore, no study flow chart was provided, and there was no reporting on any exclusions.

Selective reporting

In this 2020 review, 22 newly included studies reported all specified outcomes and were judged as at low risk of reporting bias (Aiello 2012; Barasheed 2014; Canini 2010; Chard 2019; Goodall 2014; Hartinger 2016; Ibfelt 2015; Ide 2016; Little 2015; MacIntyre 2011; MacIntyre 2013; MacIntyre 2015; MacIntyre 2016; Pandejpong 2012; Priest 2014; Radonovich 2019; Savolainen-Kopra 2012; Simmerman 2011; Suess 2012; Temime 2018; Turner 2012; Zomer 2015). For 18 studies, it is unlikely that other outcomes were measured and not reported, although no protocol was available to assess reporting bias (Aelami 2015; Alzahr 2018; Arbogast 2016; Azor-Martinez 2016; Azor-Martinez 2018; Ban 2015; Biswas 2019; Correa 2012; DiVita 2011; Feldman 2016; Hubner 2010; Huda 2012; Ide 2014; Miyaki 2011; Nicholson 2014; Stebbins 2011; Talaat 2011; Yeung 2011). Three studies were at high risk of reporting bias (McConeghy 2017; Millar 2016; Najnin 2019). In McConeghy 2017, URTI was mentioned in the methods (the intervention presumably would have targeted these), but only lower respiratory tract infection (LRTI) and overall infection were reported. Millar 2016 was originally conducted for another purpose; we could not find the respiratory outcomes reported in the study as part of the original study protocol. In Najnin 2019, the published study protocol did not include respiratory illness as an outcome.

Other potential sources of bias

An additional consideration for cluster-RCTs is identification/recruitment bias, where individuals are recruited in the trial after clusters are randomised. Such bias can introduce an imbalance amongst groups. Of the cluster-RCTs included in our 2020 review, we judged 13 to have a low risk of identification/recruitment bias (Arbogast 2016; Biswas 2019; Canini 2010; Cowling 2008; Longini

1988; Luby 2005; MacIntyre 2015; MacIntyre 2016; Roberts 2000; Sandora 2005; Suess 2012; Temime 2018; White 2001). In Arbogast 2016, all identified individuals (office workers) were included in the assigned cluster. Schools were identified and then randomised to the clusters; students were then randomly selected from each classroom and school. Nine studies described identification of participants, consenting/enrolling, and then randomising to the clusters (Canini 2010; Cowling 2008; Longini 1988; Luby 2005; MacIntyre 2015; MacIntyre 2016; Roberts 2000; Sandora 2005; White 2001). Suess 2012 identified and consented patients, then recruitment was performed by physicians unaware of cluster assignment. In Temime 2018, directors of the included nursing homes agreed to participate in the study before randomisation, and written consent was not required from the residents. We judged 11 cluster-RCTs as at high risk of identification/recruitment bias (Aiello 2010; Aiello 2012; Azor-Martinez 2018; Chard 2019; Correa 2012; Cowling 2009; Larson 2010; McConeghy 2017; Nicholson 2014; Priest 2014; Savolainen-Kopra 2012). In Aiello 2010 and Aiello 2012, recruitment continued for two weeks after start of the study, which could have introduced bias. Six trials identified and recruited participants after cluster randomisation (Azor-Martinez 2018; Chard 2019; Cowling 2009; Larson 2010; McConeghy 2017; Nicholson 2014). Three trials recruited new participants after the start of the study to replace those lost to follow-up (Correa 2012; Priest 2014; Savolainen-Kopra 2012). We judged five cluster-RCTs to have probable identification/recruitment bias (Alzahrer 2018; Barasheed 2014; MacIntyre 2011; Najnin 2019; Radonovich 2019), whereas in 19 studies there were insufficient details to permit a judgement of risk of bias (Carabin 1999; DiVita 2011; Feldman 2016; Hartinger 2016; Huda 2012; Ibfelt 2015; Kotch 1994; Ladegaard 1999; MacIntyre 2009; MacIntyre 2013; Millar 2016; Miyaki 2011; Pandepong 2012; Radonovich 2019; Sandora 2008; Stebbins 2011; Talaat 2011; Yeung 2011; Zomer 2015).

Twenty-six cluster-RCTs reported intracluster correlation coefficient (ICC) to adjust sample size, taking into consideration clustering effects, and described adjusting outcomes for clustering effect using different statistical methods, or provided justification for not performing adjusted analysis for clustering (Aiello 2010; Aiello 2012; Arbogast 2016; Canini 2010; Carabin 1999; Correa 2012; Cowling 2008; Cowling 2009; Hartinger 2016; Huda 2012; Little 2015; Luby 2005; MacIntyre 2009; MacIntyre 2011; MacIntyre 2013; MacIntyre 2015; MacIntyre 2016; McConeghy 2017; Priest 2014; Radonovich 2019; Ram 2015; Roberts 2000; Stebbins 2011; Suess 2012; Talaat 2011; Temime 2018). Five cluster-RCTs did not report the ICC but described adjusting outcomes for clustering effect using different statistical methods, or explained why adjusted analysis for clustering was not performed (Biswas 2019; Chard 2019; McConeghy 2017; Simmerman 2011; Zomer 2015). Thirteen cluster-RCTs provided insufficient details on ICC and/or did not perform adjusted analysis or justified the absence of it (Alzahrer 2018; Azor-Martinez 2016; Azor-Martinez 2018; Barasheed 2014; Feldman 2016; Larson 2010; Millar 2016; Miyaki 2011; Najnin 2019; Nicholson 2014; Pandepong 2012; Savolainen-Kopra 2012; Yeung 2011). Two cluster-RCTs reported the ICC but did not perform adjusted analysis or justified the absence of it (Sandora 2005; Sandora 2008).

Effects of interventions

See: **Summary of findings 1** Medical/surgical masks compared to no masks for preventing the spread of viral respiratory illness; **Summary of findings 2** N95 respirators compared to medical/

surgical masks for preventing the spread of viral respiratory illness; **Summary of findings 3** Hand hygiene compared to control for preventing the spread of viral respiratory illness

Comparison 1: Medical/surgical masks compared to no masks

We included nine trials (eight of which were cluster-RCTs) comparing medical/surgical masks versus no masks (Aiello 2012; Barasheed 2014; Canini 2010; Cowling 2008; Jacobs 2009; MacIntyre 2009; MacIntyre 2015; MacIntyre 2016; Suess 2012). Two trials were conducted with healthcare workers (HCWs) (Jacobs 2009; MacIntyre 2015), whilst the other seven studies included people living in the community. All trials were conducted in non-pandemic settings.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Pooling of all nine trials found an estimate of effect for the outcomes of ILI cases (risk ratio (RR) 0.99, 95% confidence interval (CI) 0.82 to 1.18; low-certainty evidence; [Analysis 1.1](#)) suggesting that wearing a medical/surgical mask may make little or no difference for this outcome. Similarly, the estimate of effect for laboratory-confirmed influenza cases (RR 0.91, 95% CI 0.66 to 1.26; 6 trials; moderate-certainty evidence; [Analysis 1.1](#)) suggests that wearing a medical/surgical mask probably makes little or no difference compared to not wearing a mask for this outcome. We downgraded the certainty of the evidence two levels for ILI due to inconsistency of the effect across studies and wide CI of the pooled effect. Sixty-five per cent of the weight of the ILI analysis was carried by one study (Aiello 2012).

A separate analysis of the two trials in healthcare workers for the outcome ILI (RR 0.37, 95% CI 0.05 to 2.50; low-certainty evidence; [Analysis 1.2](#)) suggests that there is considerable uncertainty as to whether there is any benefit (Jacobs 2009; MacIntyre 2015). The effect estimate was downgraded due to very wide CI interval of the pooled effect.

The design of most trials assessed whether masks protected the wearer. Four trials were cluster-RCTs, with all participants in the intervention clusters required to wear masks, thus assessing both source control and personal protection. In two trials the clusters were households with a member with new influenza; neither of these studies found any protective effect (RR 1.03 in 105 households (Canini 2010); RR 1.21 in 145 households (MacIntyre 2009)). In two trials the clusters were college dormitories during the influenza season; neither study found any reduction (RR 1.10 in 37 dormitories (Aiello 2012); RR 0.90 in three dormitories (Aiello 2010)). We excluded Aiello 2010 from the meta-analysis since we did not consider 'randomisation' of three clusters to three arms to be a proper randomised trial.

2. Adverse events related to the intervention

Canini 2010 reported that 38 (75%) of participants in the intervention arm experienced discomfort with the mask use due to warmth (45%), respiratory difficulties (33%), and humidity (33%). Children reported feeling pain more frequently (3/12) than other participants wearing adult face masks (1/39; $P = 0.04$). In MacIntyre 2015, adverse events associated with face mask use were reported in 40.4% (227/562) of HCWs in the medical-mask arm. General discomfort (35.1%; 397/1130) and breathing problems (18.3%;

207/1130) were the most frequently reported adverse events. [Suess 2012](#) reported that the majority of participants (107/172; 62%) did not report any problems with mask-wearing. More adults reported no problems (71%) compared to children (36/72; 50%; $P = 0.005$). The main issues when wearing a face mask for adults as well as for children were "heat/humidity" (18/34; 53% of children; 10/29; 35% of adults; $P = 0.1$), followed by "pain" and "shortness of breath". [Cowling 2008](#) mentioned that no adverse events were reported. The other trials did not report measuring adverse outcomes.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

[Jacobs 2009](#) reported that participants in the mask group were significantly more likely to experience more days with headache and feeling bad. They found no significant differences between the two groups for symptom severity scores. None of the other trials reported this outcome.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 2: N95/P2 respirators compared to medical/surgical masks

We included five trials comparing medical/surgical masks with N95/P2 respirators ([Loeb 2009](#); [MacIntyre 2009](#); [MacIntyre 2011](#); [MacIntyre 2013](#); [Radonovich 2019](#)). All of these trials except [MacIntyre 2009](#) included HCWs. [MacIntyre 2009](#) included carers and household members of children with a respiratory illness recruited from a paediatric outpatient department and a paediatric primary care practice in Sydney, Australia.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Pooling of three trials found an estimate of effect suggesting considerable uncertainty as to whether an N95/P2 respirator provides any benefit compared to medical/surgical masks for the outcome of clinical respiratory illness (RR 0.70, 95% CI 0.45 to 1.10; very low-certainty evidence; [Analysis 2.1](#)) ([MacIntyre 2011](#); [MacIntyre 2013](#) (2 arms); [Radonovich 2019](#)). Based on five trials conducted in four healthcare settings and one household ([Loeb 2009](#); [MacIntyre 2009](#); [MacIntyre 2011](#); [MacIntyre 2013](#); [Radonovich 2019](#)), the estimates of effect for the outcome of ILI (RR 0.82, 95% CI 0.66 to 1.03; low-certainty evidence; [Analysis 2.1](#)) suggest that N95/P2 respirators may make little or no difference for this outcome. The estimate of the effect for the outcome of laboratory-confirmed influenza infection (RR 1.10, 95% CI 0.90 to 1.34; moderate-certainty evidence; [Analysis 2.1](#)) suggests that the use of a N95/P2 respirator compared to a medical/surgical mask probably makes little or no difference for this more precise and objective outcome.

The outcomes clinical respiratory illness and ILI were reported separately. Considering how these outcomes were defined, it is highly likely that there was considerable overlap between the two, therefore these outcomes were not combined into a single clinical outcome ([Analysis 2.1](#)). The laboratory-confirmed viral respiratory infection outcome included influenza primarily but multiple other common viral respiratory pathogens were also included in several studies. The laboratory-confirmed viral infection outcome was considered more precise and objective in comparison to the clinical outcomes, which were more subjective and considered to be less precise. The findings did not change when we restricted the evidence to HCWs ([Analysis 2.2](#)).

2. Adverse events related to the intervention

Harms were poorly reported, but generally discomfort wearing medical/surgical masks and N95/P32 respirators was mentioned in several studies. [Radonovich 2019](#) mentioned that participants wearing the N95 respirator reported skin irritation and worsening of acne. [MacIntyre 2011](#) reported that adverse events were more common with N95 respirators; in particular, discomfort was reported in 41.9% of N95 wearers versus 9.8% of medical-mask wearers ($P < 0.01$); headaches were more common with N95 (13.4% versus 3.9%; $P < 0.01$); difficulty breathing was reported more often in the N95 group (19.4% versus 12.5%; $P = 0.01$); and N95 caused more problems with pressure on the nose (52.2% versus 11.0%; $P < 0.01$). In [MacIntyre 2013](#), fewer participants using the N95 respirator reported problems (38% (195/512) versus 48% (274/571) of participants in the medical-mask arm; $P = 0.001$). [Loeb 2009](#) mentioned that no adverse events were reported.

The one trial conducted in the community mentioned that more than 50% of participants reported concerns with both types of masks, mainly that wearing them was uncomfortable, but there were no significant differences between the P2 (N95) and surgical-mask groups ([MacIntyre 2009](#)).

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

[Loeb 2009](#) reported that 42 participants (19.8%) in the surgical-mask group reported an episode of work-related absenteeism compared with 39 (18.6%) of participants in the N95 respiratory group (absolute risk difference -1.24%, 95% CI -8.75% to 6.27%; $P = 0.75$).

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

[Loeb 2009](#) reported that there were no episodes of LRTIs.

Comparison 3: Hand hygiene compared to control

Sixteen trials compared hand hygiene interventions with control and provided sufficient data to include in meta-analyses ([Azor-Martinez 2018](#); [Biswas 2019](#); [Correa 2012](#); [Cowling 2008](#); [Cowling](#)

2009; Hubner 2010; Larson 2010; Little 2015; Millar 2016; Nicholson 2014; Ram 2015; Roberts 2000; Sandora 2005; Simmerman 2011; Stebbins 2011; Zomer 2015). The populations of these studies included adults, children, and families, in settings such as schools, childcare centres, homes, and offices. None of the studies was conducted during a pandemic, although a few studies were conducted during peak influenza seasons. A further 16 trials comparing hand hygiene to a control had other outcomes or insufficient information to include in meta-analyses (Alzaher 2018; Arbogast 2016; Azor-Martinez 2016; DiVita 2011; Feldman 2016; Gwaltney 1980; Ladegaard 1999; Luby 2005; Morton 2004; Priest 2014; Savolainen-Kopra 2012; Talaat 2011; Temime 2018; Turner 2012; White 2001; Yeung 2011). The results of these trials were consistent with the findings of our meta-analyses. The results for all outcomes from the 16 trials that were meta-analysed and the 16 trials that were not meta-analysed are shown in Table 2.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Pooling of seven trials for the broad outcome of ARI showed a 16% relative reduction in the numbers of participants with ARI (RR 0.84, 95% CI 0.82 to 0.86; moderate-certainty evidence; Analysis 3.1.1) in the hand hygiene group (Analysis 3.1), suggesting a probable benefit (Azor-Martinez 2018; Correa 2012; Larson 2010; Little 2015; Millar 2016; Nicholson 2014; Sandora 2005). When considering the more strictly defined outcomes of ILI, Biswas 2019; Cowling 2008; Cowling 2009; Hubner 2010; Larson 2010; Little 2015; Ram 2015; Roberts 2000; Simmerman 2011; Zomer 2015, and laboratory-confirmed influenza, Biswas 2019; Cowling 2008; Cowling 2009; Hubner 2010; Larson 2010; Ram 2015; Simmerman 2011; Stebbins 2011, the estimates of the effect were heterogeneous, suggesting that hand hygiene made little or no difference (RR 0.98, 95% CI 0.85 to 1.13 for ILI; low-certainty evidence; Analysis 3.1.2) (RR 0.91, 95% CI 0.63 to 1.30 for laboratory-confirmed influenza; low-certainty evidence; Analysis 3.1.3) (Analysis 3.1). All 16 trials could be pooled for analysis of the composite outcome 'ARI or ILI or influenza', with each study only contributing once with the most comprehensive outcome (in terms of number of events) reported showing an 11% relative reduction in participants with a respiratory illness, suggesting that hand hygiene may offer a benefit (RR 0.89, 95% CI 0.84 to 0.95; low-certainty evidence; Analysis 3.2), but with high heterogeneity. In a sensitivity analysis we used only the most precise and unequivocal (with laboratory confirmed considered the most precise and an undefined ARI considered the least precise) outcome reported in each of 11 studies identified by JMC, an infectious disease physician, and found an estimate of effect in favour of hand hygiene, but with wider CIs (RR 0.92, 95% CI 0.80 to 1.05; Analysis 3.3).

We considered that studies in children might have a different effect than studies in adults, so we conducted subgroup analysis by age group. We found no evidence of a difference in treatment effect by age group ($P = 0.21$; Analysis 3.4).

2. Adverse events related to the intervention

Correa 2012 reported that no adverse events were observed; in the study by Priest 2014, skin reaction was recorded for 10.4% of participants in the hand sanitiser group versus 10.3% in the control group (RR 1.01, 95% CI 0.78 to 1.30).

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Three trials measured absenteeism from school or work and demonstrated a 36% relative reduction in the numbers of participants with absence in the hand hygiene group (RR 0.64, 95% CI 0.58 to 0.71; Analysis 3.5) (Azor-Martinez 2016; Hubner 2010; Nicholson 2014).

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 4: Hand hygiene + medical/surgical masks compared to control

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Six trials (Aelami 2015; Aiello 2012; Cowling 2009; Larson 2010; Simmerman 2011; Suess 2012) were able to be pooled to compare the use of the combination of hand hygiene and medical/surgical masks with control. Four of these trials were in households, two in university student residences, and one at the annual Hajj pilgrimage. For both outcomes (ILI and influenza), pooling demonstrated an estimate of effect suggesting little or no difference between the hand hygiene and medical/surgical mask combination and control. The number of trials and events was lower than for comparisons of hand hygiene alone, or medical/surgical masks alone, and the confidence interval was wide. For ILI, the RR for intervention compared to control was 1.03 (95% CI 0.77 to 1.37; Analysis 4.1.1), and for influenza it was 0.97 (95% CI 0.69 to 1.36; Analysis 4.1.2) (Analysis 4.1). Full results of these trials are shown in Table 3

2. Adverse events related to the intervention

Adverse events related to mask wearing in the study by Suess 2012 are reported under Comparison 1 (medical/surgical masks). There was no mention of adverse events related to hand hygiene.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness, e.g. pneumonia

Not reported.

Comparison 5: Hand hygiene + medical/surgical masks compared to hand hygiene

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI and laboratory-confirmed influenza)

Three trials studied the addition of medical/surgical masks to hand hygiene (Cowling 2009; Larson 2010; Simmerman 2011). All three trials had three arms, and are also included in the comparison of hand hygiene plus medical/surgical mask versus control (Comparison 4). All three studies showed no difference between hand hygiene plus medical/surgical mask groups and hand hygiene alone, for all outcomes. The estimates of effect suggested little or no difference when adding masks to hand hygiene compared to hand hygiene alone: for the outcome ILI (RR 1.03, 95% CI 0.69 to 1.53; 3 trials) and the outcome laboratory-confirmed influenza (RR 0.99, 95% CI 0.69 to 1.44), the estimates of effect were not different and the CIs were relatively wide, suggesting little or no difference (Analysis 5.1). However, the CIs around the estimates were wide and do not rule out an important benefit.

2. Adverse events related to the intervention

Not reported.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 6: Medical/surgical masks compared to other (non-N95) masks

One trial compared medical/surgical masks with cloth masks in hospital healthcare workers (MacIntyre 2015), and another trial compared catechin-treated masks versus control masks in healthcare workers and staff of hospitals, rehabilitation centres, and nursing homes in Japan (Ide 2016).

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

MacIntyre 2015 found that the rate of ILI was higher in the cloth mask arm compared to the medical/surgical masks arm (RR 13.25, 95% CI 1.74 to 100.97).

Ide 2016 did not find a benefit from the catechin-treated masks over untreated masks on influenza infection rates (adjusted odds ratio (OR) 2.35, 95% CI 0.40 to 13.72; $P = 0.34$).

2. Adverse events related to the intervention

In MacIntyre 2015 adverse events associated with face mask use were reported in 40.4% (227/562) of HCWs in the medical/surgical mask arm and 42.6% (242/568) in the cloth mask arm ($P = 0.45$). The most frequently reported adverse events were general discomfort (35.1%; 397/1130) and breathing problems (18.3%; 207/1130). Laboratory tests showed the penetration of particles through the cloth masks to be very high (97%) compared with medical/surgical masks (44%). Ide 2016 reported that there were no serious adverse events associated with the intervention.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 7: Soap + water compared to sanitiser, and comparisons of different types of sanitiser

Two trials compared soap and water with sanitiser (Azor-Martinez 2018; Savolainen-Kopra 2012). Another trial compared different types of hand sanitiser in a virus challenge study (Turner 2004a; Turner 2004b), and one trial studied the frequency of use of hand sanitiser (Pandepong 2012). The full results of these four trials are shown in Table 4.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

In the trial by Azor-Martinez 2018, ARI incidence was significantly higher in the soap-and-water group compared with the hand sanitiser group (rate ratio 1.21, 95% CI 1.06 to 1.39). In contrast, there was no significant difference between interventions in Savolainen-Kopra 2012. In the rhinovirus challenge study (Turner 2004a; Turner 2004b), all hand sanitisers tested led to a significant lowering of infection rates, but no differences between sanitisers were observed. The study sample size was small.

2. Adverse events related to the intervention

Two trials stated that no adverse events were observed (Pandejpong 2012; Savolainen-Kopra 2012).

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

The authors of Azor-Martinez 2018 also observed a significant benefit for hand sanitiser in reduction in days absent, whereas there was no difference between intervention groups in the Savolainen-Kopra 2012 trial. The study on frequency of use of sanitiser found that use of sanitiser every hour significantly reduced days absent compared with use every two hours or with use only before the lunch break (Pandejpong 2012).

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 8: Surface/object disinfection (with or without hand hygiene) compared to control

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Six trials contributed data to this comparison (Ban 2015; Carabin 1999; Ibfelt 2015; Kotch 1994; McConeghy 2017; Sandora 2008). Full results of these trials are shown in Table 5. Five of the six trials combined disinfection with other interventions such as hand hygiene education, provision of hand hygiene products, and audits. Ban 2015 utilised a combination of provision of hand hygiene products, and cleaning and disinfection of surfaces, and demonstrated a significant reduction in ARI in the intervention group (OR 0.47, 95% CI 0.48 to 0.65). A similar result was seen in Carabin 1999, with a significant reduction in episodes of ARI. Two studies tested multicomponent interventions and observed no significant difference in ARI outcomes (Kotch 1994; McConeghy 2017).

One trial compared disinfection alone to usual care (Ibfelt 2015). This study demonstrated a significant reduction in some viruses detected on surfaces in the childcare centres (adenovirus, rhinovirus, respiratory syncytial virus (RSV), and metapneumovirus), but not in other viruses, including coronavirus.

2. Adverse events related to the intervention

Not reported.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Only one study measured this outcome (Sandora 2008), observing no significant difference between groups for the outcome of absence due to respiratory illness (rate ratio for intervention to control 1.10, 95% CI 0.97 to 1.24).

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 9: Complex interventions compared to control

Complex interventions are either multifaceted environmental programmes (such as those in low-income countries) or combined interventions including hygiene measures and gloves, gowns, and masks.

Four trials studied complex hygiene and sanitation interventions in low-income country settings (Chard 2019; Hartinger 2016; Huda 2012; Najnin 2019). Full results from these studies are given in Table 6.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

All four trials of complex interventions observed no significant differences between groups in rates of viral respiratory illness.

2. Adverse events related to the intervention

Not reported

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 10: Physical distancing/quarantine compared to control

We found one quasi-cluster-RCT assessing the effectiveness of quarantining workers of one of two sibling companies in Japan whose family members developed an ILI during the 2009 to 2010 H1N1 influenza pandemic (Miyaki 2011). Workers in the intervention group were asked to stay home on full pay until

five days after the household member(s) showed resolution of symptoms or two days after alleviation of fever.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

Compliance with the intervention was 100%. In the intervention group 2.75% of workers contracted influenza, compared with 3.18% in the control group (Cox hazard ratio 0.799, 95% CI 0.66 to 0.97; $P = 0.02$), indicating that the rate of infection was reduced by 20% in the intervention group. However, the risk of a worker being infected was 2.17-fold higher in the intervention group where workers stayed at home with their infected family members. The authors concluded that quarantining workers who have infected household members could be a useful additional measure to control the spread of respiratory viruses in an epidemic setting.

2. Adverse events related to the intervention

Not reported.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 11: Eye protection compared to control

We did not find any randomised studies investigating the effect of eye protection compared to control.

Comparison 12: Gargling compared to control

Three trials investigated the effect of gargling. [Satomura 2005](#) compared throat gargling with povidone-iodine versus tap water in healthy adults. [Ide 2014](#) compared gargling with green tea versus tap water in high school students, and [Goodall 2014](#) compared gargling with tap water with no gargling in university students.

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

[Satomura 2005](#) reported that gargling with tap water reduced the incidence of URTIs compared to the control group (usual care) (hazard ratio (HR) 0.60, 95% CI 0.39 to 0.95). Gargling with povidone-iodine did not reduce the incidence of URTIs compared to the control group (HR 0.88, 95% CI 0.58 to 1.34).

[Goodall 2014](#) found no difference in laboratory-confirmed URTIs between the gargling (tap water) and no-gargling groups (RR for gargling versus no gargling 0.82, 95% CI 0.53 to 1.26; $P = 0.36$).

In a meta-analysis of gargling versus control based on two trials ([Goodall 2014](#); [Satomura 2005](#)), the pooled estimate of effect suggested little or no difference for the outcome of clinical URTI due to gargling (RR 0.91, 95% CI 0.63 to 1.31; [Analysis 6.1](#)).

There was no difference in the incidence of laboratory-confirmed influenza between high school students gargling with green tea compared with those using tap water (adjusted OR 0.69, 95% CI 0.37 to 1.28; $P = 0.24$) ([Ide 2014](#)). There was also no difference in the incidence of clinically defined influenza (adjusted OR 0.75, 95% CI 0.50 to 1.13; $P = 0.17$). However, the authors reported that adherence to the interventions amongst students was low.

2. Adverse events related to the intervention

[Satomura 2005](#) reported no adverse events during the 60-day intervention period. [Ide 2014](#) also did not observe any adverse events during the study. [Goodall 2014](#) did not report on adverse effects.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

[Satomura 2005](#) reported that the mean peak score in bronchial symptoms was lower in the water gargling group (0.97) than in the povidone-iodine gargling group (1.41) and the control group (1.40), $P = 0.055$. Other symptoms were not significantly different between groups. [Goodall 2014](#) reported that symptom severity was greater in the gargling group for clinical and laboratory-confirmed URTI, but this was not statistically significant (225.3 versus 191.8, and 210.5 versus 191.8, respectively). [Ide 2014](#) did not report symptom or illness severity.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

Comparison 13: Virucidal tissues compared to control

Two reports (three trials) conducted in the USA studied the effect of virucidal tissues ([Farr 1988a](#); [Farr 1988b](#); [Longini 1988](#)). Full results from these studies are given in [Table 7](#).

Primary outcomes

1. Numbers of cases of viral illness (including ARIs, ILI, and laboratory-confirmed influenza)

The three trials of virucidal tissues reported no differences in infection rates between tissues and placebo, and between tissues and no tissues ([Farr 1988a](#); [Farr 1988b](#); [Longini 1988](#)).

2. Adverse events related to the intervention

Farr 1988b reported cough in 4% of participants using virucidal tissues versus 57% in the placebo group, but 24% reported nasal burning in the virucidal tissue group versus 8% in the placebo group. Longini 1988 did not report on adverse effects.

Secondary outcomes

1. Deaths

Not reported.

2. Severity of viral illness as reported in the studies

Not reported.

3. Absenteeism

Not reported.

4. Hospital admissions

Not reported.

5. Complications related to the illness (e.g. pneumonia)

Not reported.

DISCUSSION

Summary of main results

See Table 8.

1. Medical/surgical masks compared to no masks

The pooled estimates of effect from RCTs and cluster-RCTs for wearing medical/surgical masks compared to no masks suggests little or no difference in interrupting the spread of ILI (RR 0.99, 95% CI 0.82 to 1.18; low-certainty evidence) or laboratory-confirmed influenza (RR 0.91, 95% CI 0.66 to 1.26; moderate-certainty evidence) in the combined analysis of all populations from the included trials. We found similar results for ILI in HCWs (RR 0.37, 95% CI 0.05 to 2.50; very low-certainty evidence). Four trials were cluster-RCTs, with all participants in the intervention clusters required to wear masks, thus assessing both source control and personal protection. In two trials the clusters were households with a member with new influenza; neither trial found any protective effect (RR 1.03 in 105 households (Canini 2010); RR 1.21 in 145 households (MacIntyre 2009)). In two trials the clusters were college dormitories during the influenza season; neither trial found any reduction (RR 1.10 in 37 dormitories (Aiello 2012); RR 0.90 in three dormitories (Aiello 2010)). We excluded Aiello 2010 from meta-analysis since we did not consider 'randomisation' of three clusters to three arms was a proper randomised trial.

Less than half of the trials comparing masks with no masks addressed harms of mask wearing (Canini 2010; Cowling 2008; MacIntyre 2015; Suess 2012). Warmth, respiratory difficulties, humidity, and general discomfort were the most frequently reported adverse events. More adults reported no harms compared to children.

In one trial (MacIntyre 2015), cloth masks were associated with a significantly higher risk of both ILI and laboratory-confirmed respiratory virus infection in HCWs. In addition, filtration capacity of the two-ply cotton cloth masks was found to be only 3% and markedly less than with surgical masks based on standardised

particle testing. The authors suggested moisture retention, poor filtration, and penetration of the virus through the mask as plausible explanations for the increased risk of infection.

We did not find any randomised trials assessing the effectiveness of barrier interventions using a combination of masks, gloves, and gowns.

2. N95 respirators compared to medical/surgical masks

Comparisons between N95 respirators and surgical masks for the outcomes of clinical respiratory illness and the outcome of laboratory-confirmed influenza showed estimates of effect suggesting considerable uncertainty for any benefit for the former outcome and probably little or no difference for the latter outcome. Five trials (four in healthcare settings and one in a household setting) compared N95/P2 respirators with surgical masks. Pooling of three of these trials showed an estimate of effect suggesting considerable uncertainty as to whether there was any benefit comparing N95 respirators and medical/surgical face masks for the outcome of clinical respiratory illness (RR 0.70, 95% CI 0.45 to 1.10; very low-certainty evidence), and that N95 respirators may make little or no difference for the outcome ILI (RR 0.82, 95% CI 0.66 to 1.03; low-certainty evidence) and probably little or no difference for the outcome laboratory-confirmed influenza (RR 1.10, 95% CI 0.90 to 1.34; moderate-certainty evidence). The presence of imprecision (wide confidence intervals) and heterogeneity, particularly for the more subjective and less precise outcomes of clinical respiratory illness and ILI compared to laboratory-confirmed influenza infection, makes it difficult to assess whether there may be a benefit of either medical/surgical masks or N95/P2 respirators. Restricting the pooling to HCWs made no difference to the overall findings. The two trials with the largest event rates were quite consistent in their findings of no significant differences between N95 and surgical masks for the outcomes laboratory-confirmed influenza and all laboratory-confirmed viral infections (Loeb 2009; Radonovich 2019). Three of the trials contributing to this analysis were carried out by members of the same group (MacIntyre 2009; MacIntyre 2011; MacIntyre 2013).

In general, harms were poorly reported or not reported at all in trials comparing N95 respirators with surgical masks. General discomfort resulting in reduced wear compliance was the most frequently reported harm.

3. Hand hygiene compared to control

We found that the estimate of effect may offer a benefit for hand hygiene for the composite outcome 'ARI or ILI or influenza' (RR 0.89, 95% CI 0.84 to 0.95; low-certainty evidence), and probably offers a benefit for the outcomes ARI alone (RR 0.84, 95% CI 0.82 to 0.86; moderate-certainty evidence) and absenteeism (RR 0.64, 95% CI 0.58 to 0.71). An observed estimate of effect in favour of hand hygiene for laboratory-confirmed influenza but with wider CIs may be a consequence of smaller sample sizes in conjunction with a more rigorous outcome measure.

4. Hand hygiene + medical/surgical masks compared to control

The estimate of effect of combined hand hygiene and mask interventions compared to control in six (mostly small) trials suggested that the intervention may make little or no difference for the outcomes ILI (RR 1.03, 95% CI 0.77 to 1.37) and laboratory-confirmed influenza (four trials) (RR 0.97, 95% CI 0.69 to 1.36).

5. Hand hygiene + medical/surgical masks compared to hand hygiene

We also found an estimate of effect suggesting that adding masks to hand hygiene compared to hand hygiene alone may make little or no difference for the outcomes ILI (RR 1.03, 95% CI 0.69 to 1.53; 3 trials) and laboratory-confirmed influenza (RR 0.99, 95% CI 0.69 to 1.44).

6. Medical/surgical masks compared to other (non-N95) masks

One trial found that medical/surgical masks were more effective than cloth masks at reducing the rate of ILI (RR 13.25, 95% CI 1.74 to 100.97) (MacIntyre 2015), but the extremely wide CIs make this finding difficult to interpret. One trial did not find a benefit from catechin-treated masks over untreated masks on influenza infection rates (adjusted OR 2.35, 95% CI 0.40 to 13.72; $P = 0.34$) (Ide 2016).

Harms of wearing masks were reported in 40.4% of HCWs using medical/surgical masks, and in 42.6% of those wearing cloth masks ($P = 0.45$) (MacIntyre 2015). The penetration of particles was higher in cloth masks (97%) compared to medical/surgical masks (44%).

7. Soap + water compared to sanitiser, and comparisons of different types of sanitiser

There were too few trials comparing different types of hand hygiene interventions to be certain of any true differences between soap and water, alcohol-based hand sanitisers, or other types of interventions. Also, it is uncertain whether the incremental effect of adding virucidals or antiseptics to hand-washing actually decreased the respiratory disease burden outside the confines of the rather atypical studies. The extra benefit may have been, at least in part, accrued by confounding additional routines.

8. Surface/object disinfection (with or without hand hygiene) compared to control

We identified six trials on surface/object disinfection (with or without hand hygiene), and although they were heterogeneous (and therefore could not be pooled), three of them showed a clear benefit compared to controls (Ban 2015; Carabin 1999; Ibfelt 2015).

We found no RCTs with nose disinfection, or disinfection of living quarters as described in observational studies reported in Jefferson 2011.

9. Complex interventions compared to control

Four trials studied complex hygiene and sanitation interventions, all in low-income country settings (Chard 2019; Hartinger 2016; Huda 2012; Najnin 2019). These trials could not be pooled due to the heterogeneity of the interventions and settings. All four trials found no significant differences between groups in the rates of viral respiratory illness.

10. Physical distancing/quarantine compared to control

A disappointing finding was the lack of proper evaluation of global and highly resource-intensive measures such as screening at entry ports and physical distancing. We identified only one trial that evaluated the effect of quarantine (Miyaki 2011), and found a reduction in influenza transmission to co-workers when those with infected household members stayed home from work. However, staying home increased their risk of being infected two-fold.

11. Eye protection compared to control

We did not find any trials assessing the effectiveness and safety of eye protection.

12. Gargling compared to control

Three trials addressed the use of gargling in preventing respiratory infections (Goodall 2014; Ide 2014; Satomura 2005). Although the trials used a variety of liquids and different outcomes, pooling the results of the two trials that compared gargling with tap water versus control did not show a favourable effect in reducing URTIs (RR 0.91, 95% CI 0.63 to 1.31) (Goodall 2014; Satomura 2005).

13. Virucidal tissues compared to control

Two reports (three trials) identified in Jefferson 2011 studied the effect of virucidal tissues compared to placebo or no tissues (Farr 1988a; Farr 1988b; Longini 1988). These trials found no differences in infection rates and could not be pooled.

Overall completeness and applicability of evidence

Several features need consideration before making generalisations based on the included studies.

The settings of the included studies, which were conducted over four decades, were heterogeneous and ranged from suburban schools, Carabin 1999, to emergency departments, intensive care units, and paediatric wards, Loeb 2009, in high-income countries; slums in low-income countries (Luby 2005); and an upper Manhattan immigrant Latino neighbourhood (Larson 2010). Few attempts were made to obtain socio-economic diversity by (for example) involving more schools in the evaluations of the same programme. We identified only a few studies from low-income countries, where the vast majority of the burden of ARIs lies and where inexpensive interventions are so critical. Additionally, limited availability of over-the-counter medications and national universal comprehensive health insurance provided with consequent physician prescription of symptomatic treatment may further limit the generalisability of findings.

The included trials generally reported few events and were conducted mostly during non-epidemic periods. The large study by Radonovich 2019 is an exception as it crossed over two of the highest reporting years for influenza in the USA between 2010 and 2017 (Elflein 2019). None of the trials were conducted during a pandemic such as SARS-CoV-1, SARS-CoV-2, or Middle East respiratory syndrome (MERS).

Of the trials assessing the effect of masks, six were carried out in those at greater exposure (i.e. HCWs) (Jacobs 2009; Loeb 2009; MacIntyre 2011; MacIntyre 2013; MacIntyre 2015; Radonovich 2019). None of these studies included HCWs undertaking aerosol-generating procedures, for which the World Health Organization (WHO) currently recommends the N95 or equivalent mask. Three trials on hand hygiene interventions were carried out in nursing homes, and included HCWs (McConeghy 2017; Temime 2018; Yeung 2011). The scarcity of RCTs on HCWs limits the generalisability of such results.

The variable quality of the methods of some studies is striking. Incomplete or no reporting of randomisation (Turner 2004a), blinding (Farr 1988a; Farr 1988b), numerators and denominators (Carabin 1999; Kotch 1994), interventions, and cluster coefficients

in the relevant trials (Carabin 1999), led to a considerable loss of information. Potential biases were often not discussed.

Inappropriate placebos caused design problems. In some studies the placebo probably carried sufficient effect to dilute the intervention effects (Longini 1988). Two valiant attempts with virucidal tissues probably failed because placebo handkerchiefs were impregnated with a dummy compound that stung the users' nostrils (Farr 1988a; Farr 1988b).

Some studies used impractical interventions. Volunteers subjected to the intervention hand cleaner (organic acids) were not allowed to use their hands between cleaning and virus challenge, so the effect of normal use of the hands on the intervention remains unknown (Turner 2004a; Turner 2004b). Two per cent aqueous iodine painted on the hands, although a successful antiviral intervention, causes unacceptable cosmetic staining, which is impractical for all but those at the highest risk of epidemic contagion (Gwaltney 1980).

Compliance with interventions, especially educational programmes, was a problem for many studies despite the importance of many such low-cost interventions. Compliance with mask wearing varied; it was generally around 60% to 80%, but was reported to be as low as 40% (see Table 1). Overall, the logistics of carrying out trials that involve sustained behaviour change are demanding, particularly in challenging settings such as immigrant neighbourhoods or students' halls of residence.

The identified trials provided sparse and unsystematic data on adverse effects of the intervention, and few of the RCTs measured or reported compliance with the intervention, which is especially important for the use of medical/surgical masks or N95 respirators. No studies investigated how the level of adherence may have influenced the effect size.

We did not identify any studies assessing the effects of eye protection, and we identified only one study on physical distancing, during the 2009 H1N1 influenza pandemic. The dearth of evidence and predominant setting of seasonal viral circulation limits generalisability of our findings to other contexts such as the COVID-19 pandemic and any future epidemics due to other respiratory viruses.

Quality of the evidence

We found the available evidence base identified through our search processes to be of variable quality. Reporting of sequence generation and allocation concealment were poor in 30% to 50% of studies across the categories of intervention comparisons. Given the nature of the intervention comparison, blinding of treatment allocation after randomisation was rarely achieved. Although blinding of outcome assessment is highly feasible and desirable, most outcomes were assessed by self-reports. Outcomes in some studies were poorly defined, with a lack of clarity as to the possible aetiologic agents (bacterial versus viral). Some studies used laboratory-confirmed outcomes, both adding precision and lowering the risk of bias (see Table 9 for heterogeneity of trial outcome definitions). We found no evidence of selective reporting of outcomes within the included studies. We believe publication bias is unlikely, as the included studies demonstrated a range of effects, both positive and negative, over all study sizes. The variable quality of the studies hampers drawing any firm conclusions.

Potential biases in the review process

The non-drug (and often locally manufactured) nature of most of the interventions in this review, the lack of effective regulation in some settings, and the possible endless number of manufacturers make it difficult to gauge the existence of unpublished data. Non-drug interventions typically have no or very poor regulation.

In this 2020 update, we focused on RCTs and cluster-RCTs, providing a higher level of evidence compared with the previous version of the review, which also meta-analysed observational studies when appropriate (Jefferson 2011). However, many of the trials were small and hence underpowered, and at high or unclear risk of bias due to poor reporting of methods and lack of blinding. The populations, outcomes, comparators, and interventions tested were heterogeneous.

Due to the urgency of this update in the context of the COVID-19 pandemic, we did not contact trial authors to request missing data. This means that we have not considered studies that included other non-respiratory infections and did not provide stratified data by type of infection.

Agreements and disagreements with other studies or reviews

Several reviews of RCTs have found broadly similar results to this review for face masks. In a meta-analysis comparing surgical masks with N95 respirators, Smith 2016 pooled three trials (Loeb 2009; MacIntyre 2011; MacIntyre 2013), and found an estimate of effect suggesting no difference for laboratory-confirmed respiratory infections (OR 0.89, 95% CI 0.64 to 1.24) or ILI (OR 0.51, 95% CI 0.19 to 1.41). A similar meta-analysis, Offeddu 2017, based on two trials, MacIntyre 2011; MacIntyre 2015, concluded that masks (either N95/P2 respirators or medical/surgical masks) were effective against clinical respiratory infections (RR 0.59, 95% CI 0.46 to 0.77) and ILI (RR 0.34, 95% CI 0.14 to 0.82). Pooling of two studies, MacIntyre 2011; MacIntyre 2013, also found an estimate of effect that favoured N95 respirators to medical/surgical masks for clinical respiratory infections (RR 0.47, 95% CI 0.36 to 0.62), but not for ILI based on three studies, Loeb 2009; MacIntyre 2011; MacIntyre 2013 (RR 0.59, 95% CI 0.27 to 1.28) (Offeddu 2017). The outcome of clinical respiratory infection is considered to be the most subjective and least precise outcome.

A recent meta-analysis included five trials comparing N95/P2 respirators with medical/surgical masks and found no difference between groups for either influenza (RR 1.09, 95% CI 0.92 to 1.28) or respiratory viral infections (RR 0.89, 95% CI 0.70 to 1.11) (Long 2020). By excluding Loeb 2009 (an open, non-inferiority RCT that compared surgical masks with N95 respirators in protecting HCWs against influenza), the authors reported a significant protective effect against viral infections (RR 0.61, 95% CI 0.39 to 0.98). The authors do not report a rationale for the exclusion in the sensitivity analysis and do not report on exclusion of the studies with low weighting, which arguably would be more relevant in a sensitivity analysis. The two trials that make up 96% of the weighting, Loeb 2009; Radonovich 2019, demonstrated no significant differences in the outcome events. A recent meta-analysis of four RCTs (Bartoszek 2020), adjusting for clustering, which compared N95 respirators with the use of medical masks, found pooled estimates of effect that did not demonstrate any difference in any laboratory-confirmed viral respiratory infection

(OR 1.06, 95% CI 0.90 to 1.25), laboratory-confirmed influenza (OR 0.94, 95% CI 0.73 to 1.20), or clinical respiratory illness (OR 1.49, 95% CI 0.98 to 2.28), with the evidence profile suggesting that there was greater imprecision and inconsistency in the outcome of clinical respiratory illness. Moreover, in another recent systematic review that assessed the effectiveness of personal protective and environmental measures in non-healthcare settings (funded by the WHO), 10 RCTs reporting estimates of the effectiveness of face masks in reducing laboratory-confirmed influenza virus infections in the community were identified (Xiao 2020). The evidence from these RCTs suggested that the use of face masks either by infected persons or by uninfected persons does not have a substantial effect on influenza transmission.

The findings from several systematic reviews and meta-analyses over the last decade have not demonstrated any difference in the clinical effectiveness of N95 respirators or equivalent compared to the use of surgical masks when used by HCWs in multiple healthcare settings for the prevention of respiratory virus infections, including influenza.

Reviews based on observational studies have usually found a stronger protective effect for face masks, but have important biases. The review by Chu 2020 did not consider RCTs of influenza transmission, but only the observational studies examining impact on SARS, MERS, or SARS-CoV-2. For N95 masks versus no mask in HCWs, there was a large protective effect with an OR of 0.04 (95% CI 0.004 to 0.30); for surgical masks versus no masks, there was an OR of 0.33 (0.17 to 0.61) overall, but four of these studies were in healthcare settings. Chu 2020 has been criticised for several reasons: use of an outdated 'Risk of bias' tool; inaccuracy of distance measures; and not adequately addressing multiple sources of bias, including recall and classification bias and in particular confounding. Confounding is very likely, as preventive behaviours such as mask use, social distancing, and hand hygiene are correlated behaviours, and hence any effect estimates are likely to be overly optimistic.

Also based on observational studies, Jefferson 2011 found a protective effect of wearing surgical masks with hygienic measures compared to not wearing masks in the SARS 2003 outbreak (OR 0.32, 95% CI 0.26 to 0.39). However, the evidence was based on case-control studies carried out during the outbreak. There was some additional but very limited supportive evidence from the cohort studies in Jefferson 2011.

Although the use of eye protection and physical distancing measures are widely believed to be effective in reducing transmission of respiratory viruses and mitigating the impact of an influenza pandemic, we found only one trial investigating the role of self-quarantine in reducing the incidence of H1N1 influenza events in the workplace, and no trials examining the effect of eye protection. The evidence for these measures was derived largely from observational studies and simulation studies, and the overall quality of supporting evidence is relatively low. The finding of limited evidence evaluating these interventions was also consistent with a recent review funded by the WHO for the preparation of its guidelines on the use of non-pharmaceutical interventions for pandemic influenza in non-medical settings (Fong 2020).

There are several previous systematic reviews on hand hygiene and respiratory infections. Five of them reviewed the evidence

in a community setting (Moncion 2019; Rabie 2006; Saunders-Hastings 2017; Warren-Gash 2013; Wong 2014), and three focused on children (Mbakaya 2017; Willmott 2016; Zivich 2018). The earliest review in 2006 included eight studies (Rabie 2006), three of which were RCTs. The pooled estimate of seven studies was described as "indicative" of the effect of hand hygiene, but the studies were of poor quality. The Warren-Gash 2013 review included 16 studies (10 of which were RCTs) and reported mixed and inconclusive results. A 2014 review identified 10 RCTs and reported that the combination of hand hygiene with face masks in high-income countries (five trials) significantly reduced the incidence of laboratory-confirmed influenza and ILI, whilst hand hygiene alone did not (Wong 2014). This significant reduction in laboratory-confirmed influenza and ILI for hand hygiene and face masks may have been based on the raw numbers without adjusting for any clustering effects in the included cluster trials, which produced inappropriately narrow CIs, and possibly biased treatment effect estimates. Moreover, trials from the low-income countries were not included in the review, and this significant effect was not demonstrated when all the trials identified in the review were combined. The Saunders-Hastings 2017 review of studies evaluating the effectiveness of personal protective measures in interrupting pandemic influenza transmission only identified two RCTs (Azor-Martinez 2014; Suess 2012), which reported a significant effect of hand hygiene. The Moncion 2019 review identified seven RCTs of hand hygiene compared to control, with mixed results for preventing the transmission of laboratory-confirmed or possible influenza. Systematic reviews of RCTs of hand hygiene interventions amongst children, Mbakaya 2017; Willmott 2016, or at a non-clinical workplace, Zivich 2018, identified heterogeneous trials with quality problems including small numbers of clusters and participants, inadequate randomisation, and self-reported outcomes. Evidence of impact on respiratory infections was equivocal.

AUTHORS' CONCLUSIONS

Implications for practice

The evidence summarised in this review on the use of masks is largely based on studies conducted during traditional peak respiratory virus infection seasons up until 2016. We will incorporate relevant published studies in COVID-19 when their results are available. The observed lack of effect of mask wearing in interrupting the spread of ILI or influenza in our review has many potential reasons, including: poor study design; insufficiently powered studies arising from low viral circulation in some studies; lower compliance with mask wearing, especially among children; quality of the masks used; self-contamination of the mask by hands; lack of protection from eye exposure from respiratory droplets (allowing a route of entry of respiratory viruses into the nose via the lacrimal duct); saturation of masks with saliva from extended use (promoting virus survival in proteinaceous material); and risk compensation behaviour leading to an exaggerated sense of security (Brosseau 2020; Canini 2010; Cassell 2006; MacIntyre 2015; Rengasamy 2010; Zamora 2006).

Our findings show that hand hygiene has a modest effect as a physical intervention to interrupt the spread of respiratory viruses, but several questions remain. First, the high heterogeneity between studies may suggest that there are differences in the effect of different interventions. The poor reporting limited our ability to extract the information needed to assess any 'dose response'

relationship, and there are few head-to-head trials comparing hand hygiene materials (such as alcohol-based sanitiser or soap and water). Second, the sustainability of hand hygiene is unclear where participants in some studies achieved 5 to 10 hand-washings per day, but compliance may have diminished with time as motivation decreased, or due to adverse effects from frequent hand-washing. Third, there is little evidence about the effectiveness of combinations of hand hygiene with other interventions, and how those are best introduced and sustained. Finally, some interventions were intensively implemented within small organisations, and involved education or training as a component, and the ability to scale these up to broader interventions is unclear.

Our findings with respect to hand hygiene should be considered generally relevant to all viral respiratory infections, given the diverse populations where transmission of viral respiratory infections occurs. The participants were adults, children and families, and multiple congregation settings including schools, childcare centres, homes, and offices. Most respiratory viruses, including the pandemic SARS-CoV-2, are considered to be predominantly spread via respiratory droplets or contact routes, or both (WHO 2020c). Data from studies of SARS-CoV-2 contamination of the environment based on the presence of viral ribonucleic acid (RNA) suggest significant fomite contamination from the virus (Ong 2020; Wu 2020). Hand hygiene would be expected to be beneficial in reducing the spread of SARS-CoV-2 similar to other beta coronaviruses (SARS-CoV-1, Middle East respiratory syndrome (MERS), and human coronaviruses), which are very susceptible to the concentrations of alcohol commonly found in most hand sanitiser preparations (Rabenau 2005; WHO 2020c). Support for this effect is the finding that poor hand hygiene, despite the use of full PPE, was independently associated with an increased risk of SARS-CoV-2 transmission to healthcare workers in a retrospective cohort study in Wuhan, China in both a high-risk and low-risk clinical unit for patients infected with COVID-19 (Ran 2020). The practice of hand hygiene appears to have a consistent effect in all settings, and should be an essential component of other interventions.

The highest-quality cluster-RCTs indicate that the most effect on preventing respiratory virus spread from hygienic measures occurs in younger children. This may be because younger children are least capable of hygienic behaviour themselves (Roberts 2000), and have longer-lived infections and greater social contact, thereby acting as portals of infection into the household (Monto 1969). Additional benefit from reduced transmission from them to other members of the household is broadly supported by the results of other study designs where the potential for confounding is greater.

Routine long-term implementation of some of the interventions covered in this review may be problematic, particularly maintaining strict hygiene and barrier routines for long periods of time. This would probably only be feasible in highly motivated environments, such as hospitals. Many of the trial authors commented on the major logistical burdens that barrier routines imposed at the community level. However, the threat of a looming epidemic may provide stimulus for their inception.

Implications for research

Public health measures and physical interventions can be highly effective to interrupt the spread of respiratory viral infections, especially when they are part of a structured and co-ordinated programme that includes instruction and education, and when

they are delivered together. Our review has provided important insights into research gaps that need to be addressed with respect to these physical interventions and their implementation. The 2014 WHO document 'Infection prevention and control of epidemic- and pandemic-prone acute respiratory infections in health care' identified several research gaps as part of their GRADE assessment of their infection prevention and control recommendations, which remain very relevant (WHO 2014). Research gaps identified during the course of our review and the WHO 2014 document may be considered from the perspective of both general and specific themes.

A general theme identified was the need to provide outcomes with explicitly defined clinical criteria for acute respiratory infections (ARIs) and discrete laboratory-confirmed outcomes of viral ARIs using molecular diagnostic tools which are now widely available. Our review found large disparities between studies with respect to the clinical outcome events, which were imprecisely defined in several studies, and there were differences in the extent to which laboratory-confirmed viruses were included in the studies that assessed them. Another general theme identified was the lack of consideration of sociocultural factors that might affect compliance with the interventions, especially those employed in the community setting. In addition, the cost and resource implications of the physical interventions employed in different settings would have important relevance for low- to middle-income countries. Resources have been a major issue with the COVID-19 pandemic, with global shortages of several components of PPE. Several specific research gaps related to physical interventions were identified within the WHO 2014 document and are congruent with many of the findings of our current update, including the following: transmission dynamics of respiratory viruses from patients to healthcare workers during aerosol-generating procedures; a lack of precision with regards to defining aerosol-generating procedures; the safety of cohorting of patients with the same suspected but unconfirmed diagnosis in a common unit or ward with patients infected with the same known pathogen in healthcare settings; the optimal duration of the use of physical interruptions to prevent spread of ARI viruses; use of spatial separation or physical distancing (in healthcare and community settings, respectively) alone versus spatial separation or physical distancing with the use of other added physical interventions coupled with examining discrete distance parameters (e.g. 1 metre, 2 metres, or > 2 metres); the effectiveness of respiratory etiquette (i.e. coughing/sneezing into tissues or a sleeved bent elbow); the effectiveness of triage and early identification of infected individuals with an ARI in both hospital and community settings; use of frequent disinfection techniques appropriate to the setting (high-touch surfaces in the environment, gargling with oral disinfectants, and virucidal tissues or clothing) alone or in combination with facial masks and hand hygiene; the use of ultraviolet light germicidal irradiation for disinfection of air in healthcare and selected community settings; and the use of widespread compliance with effective vaccination strategies.

There is a clear requirement to conduct large, pragmatic trials to evaluate the best combinations in the community and in healthcare settings with multiple respiratory viruses and in different sociocultural settings. RCTs with a pragmatic design, similar to the Luby 2005 trial, should be conducted whenever possible. Alternately, large population-based cohort studies may

also be considered if individual RCTs prove to be too expensive or less practical, depending on the issue that is being addressed.

Several specific research gaps deserve expedited attention and may be highlighted within the context of the COVID-19 pandemic. The use of facial masks in the community setting represents one of the most pressing needs to address, given the polarised opinions around the world. Both broad-based ecological studies, adjusting for confounding and high-quality randomised trials, may be necessary to determine if there is an independent contribution to their use as a physical intervention, and how they may best be deployed to optimise their contribution. The type of fabric and weave used in the face mask is an equally pressing concern, given that surgical masks with their cotton-polypropylene fabric appear to be effective in the healthcare setting, but there are questions about the effectiveness of simple cotton masks. In addition, these masking intervention studies should focus on measuring not only benefits but also compliance, harms, and risk compensation if the latter may lead to a lower protective effect. In addition, although the use of surgical masks versus N95 respirators demonstrates no differences in clinical effectiveness to date, their use needs to be studied in the setting of a new pandemic such as COVID-19, and with concomitant measurement of harms, which to date have been poorly studied. Physical distancing represents another major research gap which needs to be addressed expediently, especially within the context of the COVID-19 pandemic setting as well as in future epidemic settings. The use of quarantine and screening at entry ports needs to be investigated in well-designed, high-quality studies. We found only one RCT of quarantine, and no trials of screening at entry ports or physical distancing. Given that this is one of the primary strategies applied globally in the face of the COVID-19 pandemic, future trials should be conducted within the context of this pandemic, as well as in future epidemics with other respiratory viruses of less virulence.

The variable quality and small scale of some studies is known from descriptive studies (Aiello 2002; Fung 2006; WHO 2006b), and systematic reviews of selected interventions (Meadows 2004). In summary, more high-quality studies are needed to evaluate the most effective strategies to implement successful physical

interventions in practice, both on a small scale and at a population level. Finally, we emphasise that more attention should be paid to describing and quantifying the harms of the interventions assessed in this review and their relationship with compliance.

ACKNOWLEDGEMENTS

Thanks to the following people for commenting on the draft protocol and review: Anne Lyddiatt, Stephanie Kondos, Tom Sandora, Kathryn Glass, Max Bulsara, Rick Shoemaker, and Allen Cheng. Thanks also to the following people for translating non-English language trials for the draft review: Jørgen Lous for translating a Danish paper and extracting data, Ryuki Kassai who translated a Japanese paper, and Taixiang Wu who translated several Chinese papers. We thank the following people for commenting on the 2009, 2010, and 2011 updated reviews, respectively: for 2009, Anne Lyddiatt, Thomas Sandora, Michael Broderick, Sree Nair, and Allen Cheng; for 2010, Carmem Pessoa da Silva and Sergey Eremin; and for 2011, Amy Zelmer, Tom Sandora, Sree Nair, and Allen Cheng. Drs Aiello and Larson provided additional information on their trials for the 2010 update. Dr V Shukla, Dr J Conly, and K Lee prepared the GRADE tables using GRADEprofiler software. Drs Carmem Pessoa da Silva and Thomas Haustein provided linguistic assistance for the 2010 update.

We gratefully acknowledge funding for the 2009 update from the UK National Institute for Health Research (NIHR) and the National Health and Medical Research Council (NHMRC) of Australia and from WHO for the 2010 update. We also thank Ruth Foxlee and Alex Rivetti for constructing the search strategies for the previous review versions, and Bill Hewak and Adi Prabhala for extracting data in the original review. Finally, we thank Sree Nair, who co-authored previous versions of this review.

For this 2020 update, we thank Dr Elizabeth Gibson for assistance with describing the intervention details of the included trials. We also wish to thank the following people for commenting on the draft update: Dr CJ Cates, Reader in Evidence Based Medicine, SGUL, Dr Teresa Neeman, Dr Robert Ware, Dr Janet Wale, and Prof Allen Cheng.

REFERENCES

References to studies included in this review

Aelami 2015 {published data only}

Aelami MH, Khajehdalouee M, Yagoubian H, Amel Jamahdar S, Pittet D. Implementation of an educational intervention among Iranian hajj pilgrims for the prevention of influenza-like illness. *Antimicrobial Resistance and Infection Control* 2015;**4**(1):48.

Aiello 2010 {published data only}

Aiello AE, Murray GF, Perez V, Coulborn RM, Davis BM, Uddin M, et al. Mask use, hand hygiene, and seasonal influenza-like illness among young adults: a randomized intervention trial. *Journal of Infectious Diseases* 2010;**201**(4):491-8.

Aiello 2012 {published data only}

Aiello AE, Perez V, Coulborn RM, Davis BM, Uddin M, Monto AS. Facemasks, hand hygiene, and influenza among young adults: a randomized intervention trial. *PLOS ONE* 2012;**7**(1):e29744. [CTG: NCT00490633]

Alzahr 2018 {published data only} [10.15537/smj.2018.10.23344](#)

Alzahr AA, Almudarra SS, Mustafa MH, Gosadi IM. The importance of hand hygiene education on primary schoolgirls' absence due to upper respiratory infections in Saudi Arabia. *Saudi Medical Journal* 2018;**39**(10):1044-9. [DOI: [10.15537/smj.2018.10.23344](#)]

Arbogast 2016 {published data only}

Arbogast JW, Moore-Schiltz L, Jarvis WR, Harpster-Hagen A, Hughes J, Parker A. Impact of a comprehensive workplace hand hygiene program on employer health care insurance claims and costs, absenteeism, and employee perceptions and practices. *Journal of Occupational and Environmental Medicine* 2016;**58**(6):e231-40.

Azor-Martinez 2016 {published data only}

Azor-Martinez E, Cobos-Carrascosa E, Seijas-Vazquez ML, Fernández-Sánchez C, Strizzi JM, Torres-Alegre P, et al. Hand hygiene program decreases school absenteeism due to upper respiratory infections. *Journal of School Health* 2016;**86**(12):873-81.

Azor-Martinez 2018 {published data only}

Azor-Martinez E, Yui-Hifume R, Muñoz-Vico FJ, Jimenez-Noguera E, Strizzi JM, Martinez-Martinez I, et al. Effectiveness of a hand hygiene program at child care centers: a cluster randomized trial. *Pediatrics* 2018;**142**(5):e20181245.

Ban 2015 {published data only}

Ban HQ, Li T, Shen J, Li J, Peng PZ, Ye HP, et al. Effects of multiple cleaning and disinfection interventions on infectious diseases in children: a group randomized trial in China. *Biomedical and Environmental Sciences* 2015;**28**(11):779-87.

Barasheed 2014 {published data only} [12613001007729](#)

Barasheed O, Almasri N, Badahdah AM, Heron L, Taylor J, McPhee K, et al, Hajj Research Team. Pilot randomised controlled trial to test effectiveness of facemasks in preventing influenza-like illness transmission among Australian

hajj pilgrims in 2011. *Infectious Disorders Drug Targets* 2014;**14**(2):110-6. [DOI: [10.2174/1871526514666141021112855](#)]

Biswas 2019 {published data only}

Biswas D, Ahmed M, Roguski K, Ghosh PK, Parveen S, Nizame FA, et al. Effectiveness of a behavior change intervention with hand sanitizer use and respiratory hygiene in reducing laboratory-confirmed influenza among schoolchildren in Bangladesh: a cluster randomized controlled trial. *American Journal of Tropical Medicine and Hygiene* 2019;**101**(6):1446-55. [DOI: [10.4269/ajtmh.19-0376](#)]

Canini 2010 {published data only}

Canini L, Andreoletti L, Ferrari P, D'Angelo R, Blanchon T, Lemaitre M, et al. Surgical mask to prevent influenza transmission in households: a cluster randomized trial. *PLOS ONE* 2010;**5**(11):e13998. [DOI: [10.1371/journal.pone.0013998](#)]

Carabin 1999 {published data only}

Carabin H, Gyorkos TW, Soto JC, Joseph L, Payment P, Collet JP. Effectiveness of a training program in reducing infections in toddlers attending day care centers. *Epidemiology* 1999;**10**(3):219-27.

Chard 2019 {published data only} [10.7189/jogh.09.020402](#)

Chard AN, Garn JV, Chang HW, Clasen T, Freeman MC. Impact of a school-based water, sanitation, and hygiene intervention on school absence, diarrhea, respiratory infection, and soil-transmitted helminths: results from the WASH HELPS cluster-randomized trial. *Journal of Global Health* 2019;**9**(2):1-14. [DOI: [10.7189/jogh.09.020402](#)]

Correa 2012 {published data only}

Correa JC, Pinto D, Salas LA, Camaccho JC, Rondon M, Quintero J. A cluster-randomized controlled trial of handrubs for prevention of infectious diseases among children in Colombia. *Journal of Public Health* 2012;**31**(6):476-84.

Cowling 2008 {published data only}

Cowling BJ, Fung RO, Cheng CK, Fang VJ, Chan KH, Seto WH, et al. Preliminary findings of a randomized trial of non-pharmaceutical interventions to prevent influenza transmission in households. *PLOS ONE* 2008;**3**(5):e2101.

Cowling 2009 {published data only}

Cowling BJ, Chan KH, Fang VJ, Cheng CKY, Fung ROP, Wai W, et al. Facemasks and hand hygiene to prevent influenza transmission in households. A randomized trial. *Annals of Internal Medicine* 2009;**151**(7):437-46.

DiVita 2011 {published data only}

DiVita MA, Khatun-e-Jannat K, Islam M, Cercone E, Rook K, Sohel BM, et al. Impact of intensive handwashing promotion on household transmission of influenza in a low income setting: preliminary results of a randomized controlled clinical trial. *American Journal of Tropical Medicine and Hygiene* 2011;**85**(6):381.

Farr 1988a {published data only}

Farr BM, Hendley JO, Kaiser DL, Gwaltney JM. Two randomised controlled trials of virucidal nasal tissues in the prevention of natural upper respiratory infections. *American Journal of Epidemiology* 1988;**128**:1162-72.

Farr 1988b {published data only}

Farr BM, Hendley JO, Kaiser DL, Gwaltney JM. Two randomised controlled trials of virucidal nasal tissues in the prevention of natural upper respiratory infection. *American Journal of Epidemiology* 1988;**128**:1162-72.

Feldman 2016 {published data only}

Feldman L, Galili E, Cohen Y, Hartal M, Yavnai N, Netzer I. Routine chlorhexidine gluconate use onboard navy surface vessels to reduce infection: a cluster randomized controlled trial. *American Journal of Infection Control* 2016;**44**(12):1535-8.

Goodall 2014 {published data only}

Goodall EC, Granados AC, Luinstra K, Pullenayegum E, Coleman BL, Loeb M, et al. Vitamin D3 and gargling for the prevention of upper respiratory tract infections: a randomized controlled trial. *BMC Infectious Diseases* 2014;**14**:273.

Gwaltney 1980 {published data only}

Gwaltney JM Jr, Moskalski PB, Hendley JO. Interruption of experimental rhinovirus transmission. *Journal of Infectious Diseases* 1980;**142**(6):811-5.

Hartinger 2016 {published data only}

Hartinger SM, Lanata CF, Hattendorf J, Verastegui H, Gil AI, Wolf J, et al. Improving household air, drinking water and hygiene in rural Peru: a community-randomized controlled trial of an integrated environmental home-based intervention package to improve child health. *International Journal of Epidemiology* 2016;**45**(6):2089-99.

Hubner 2010 {published data only}

Hubner NO, Hubner C, Wodny M, Kampf G, Kramer A. Effectiveness of alcohol-based hand disinfectants in a public administration: impact on health and work performance related to acute respiratory symptoms and diarrhoea. *BMC Infectious Diseases* 2010;**10**(250):1-8.

Huda 2012 {published data only}

Huda TM, Unicomb L, Johnston RB, Halder AK, Yushuf Sharker MA, Luby SP. Interim evaluation of a large scale sanitation, hygiene and water improvement programme on childhood diarrhea and respiratory disease in rural Bangladesh. *Social Science & Medicine* 2012;**75**(4):604-11.

Ibfelt 2015 {published data only}

Ibfelt T, Engelund EH, Schultz AC, Andersen LP. Effect of cleaning and disinfection of toys on infectious diseases and micro-organisms in daycare nurseries. *Journal of Hospital Infection* 2015;**89**(2):109-15.

Ide 2014 {published data only}

Ide K, Yamada H, Matsushita K, Ito M, Nojiri K, Toyoizumi K, et al. Effects of green tea gargling on the prevention of influenza

infection in high school students: a randomized controlled study. *PLOS ONE* 2014;**9**(5):e96373.

Ide 2016 {published data only}

Ide K, Yamada H, Takuma N, Kawasaki Y, Morohoshi H, Takenaka A, et al. Effects of catechin-treated masks on the prevention of influenza infection: an exploratory randomized study. *Japanese Journal of Clinical Pharmacology and Therapeutics* 2016;**47**(6):229-34.

Jacobs 2009 {published data only}

Jacobs JL, Ohde S, Takahashi O, Tokuda Y, Omata F, Fukui T. Use of surgical face masks to reduce the incidence of the common cold among health care workers in Japan: a randomized controlled trial. *American Journal of Infection Control* 2009;**37**(5):417-9.

Kotch 1994 {published data only}

Kotch JB, Weigle KA, Weber DJ, Clifford RM, Harms TO, Loda FA, et al. Evaluation of an hygienic intervention in child day-care centers. *Pediatrics* 1994;**94**(6 Pt 2):991-4.

Ladegaard 1999 {published data only}

Ladegaard MB, Stage V. Hand hygiene and sickness among small children attending day care centres. An interventional study. *Ukeskrift von Laeger* 1999;**161**:4396-400.

Larson 2010 {published data only}

Larson EL, Ferng Y, Wong-McLoughlin J, Wang S, Haber M, Morse SS. Impact of non-pharmaceutical interventions on URIs and influenza in crowded, urban households. *Public Health Reports* 2010;**125**(2):178-91.

Little 2015 {published data only}

Little P, Stuart B, Hobbs FD, Moore M, Barnett J, Popoola D, et al. An internet-delivered handwashing intervention to modify influenza-like illness and respiratory infection transmission (PRIMIT): a primary care randomised trial. *Lancet* 2015;**386**:1631-9.

Loeb 2009 {published data only}

Loeb M, Dafoe N, Mahony J, John M, Sarabia A, Glavin V, et al. Surgical mask vs N95 respirator for preventing influenza among health care workers: a randomized trial. *JAMA* 2009;**302**(17):1865-71.

Longini 1988 {published data only}

Longini IM Jr, Monto AS. Efficacy of virucidal nasal tissues in interrupting familial transmission of respiratory agents. A field trial in Tecumseh, Michigan. *American Journal of Epidemiology* 1988;**128**(3):639-44.

Luby 2005 {published data only}

Luby SP, Agboatwalla M, Feikin DR, Painter J, Billhimer W, Altaf A, et al. Effect of handwashing on child health: a randomised controlled trial. *Lancet* 2005;**366**(9481):225-33.

MacIntyre 2009 {published data only}

MacIntyre CR, Cauchemez S, Dwyer DE, Seale H, Cheung P, Browne G, et al. Face mask use and control of respiratory virus

transmission in households. *Emerging Infectious Diseases* 2009;**15**(2):233-41.

MacIntyre 2011 {published data only}

* MacIntyre CR, Wang Q, Cauchemez S, Seale H, Dwyer DE, Yang P, et al. A cluster randomized clinical trial comparing fit-tested and non-fit-tested N95 respirators to medical masks to prevent respiratory virus infection in health care workers. *Influenza and Other Respiratory Viruses* 2011;**5**(3):170-9.

MacIntyre CR, Wang Q, Rahman B, Seale H, Ridda I, Gao Z, et al. Efficacy of face masks and respirators in preventing upper respiratory tract bacterial colonization and co-infection in hospital healthcare workers. *Preventive Medicine* 2014;**62**:1-7.

MacIntyre 2013 {published data only}**12609000778280**

MacIntyre CR, Wang Q, Seale H, Yang P, Shi W, Gao Z, et al. A randomized clinical trial of three options for N95 respirators and medical masks in health workers. *American Journal of Respiratory and Critical Care Medicine* 2013;**187**(9):960-6. [DOI: [10.1164/rccm.201207-1164OC](https://doi.org/10.1164/rccm.201207-1164OC)]

MacIntyre 2015 {published data only}

MacIntyre CR, Seale H, Dung TC, Hien NT, Nga PT, Chughtai AA, et al. A cluster randomised trial of cloth masks compared with medical masks in healthcare workers. *BMJ Open* 2015;**5**(4):e006577.

MacIntyre 2016 {published data only}

MacIntyre CR, Zhang Y, Chughtai AA, Seale H, Zhang D, Chu Y, et al. Cluster randomised controlled trial to examine medical mask use as source control for people with respiratory illness. *BMJ Open* 2016;**6**(12):e012330. [DOI: [10.1136/bmjopen-2016-012330](https://doi.org/10.1136/bmjopen-2016-012330)]

McConeghy 2017 {published data only}

McConeghy KW, Baier R, McGrath KP, Baer CJ, Mor V. Implementing a pilot trial of an infection control program in nursing homes: results of a matched cluster randomized trial. *Journal of the American Medical Directors Association* 2017;**18**(8):707-12.

Millar 2016 {published data only}**10.1017/ice.2016.154**

Millar EV, Schlett CD, Law NN, Chen WJ, D'Onofrio MJ, Bennett JW, et al. Reduction in acute respiratory infection among military trainees: secondary effects of a hygiene-based cluster-randomized trial for skin and soft-tissue infection prevention. *Infection Control and Hospital Epidemiology* 2016;**37**(9):1118-20.

Miyaki 2011 {published data only}

Miyaki K, Sakurazawa H, Mikurube H, Nishizaka M, Ando H, Song Y, et al. An effective quarantine measure reduced the total incidence of influenza A H1N1 in the workplace: another way to control the H1N1 flu pandemic. *Journal of Occupational Health* 2011;**53**(4):287-92. [DOI: [10.1539/joh.10-0024-fs](https://doi.org/10.1539/joh.10-0024-fs)]

Morton 2004 {published data only}

Morton JL, Schultz AA. Healthy hands: use of alcohol gel as an adjunct to handwashing in elementary school children. *Journal of School Nursing* 2004;**20**(3):161-7.

Najnin 2019 {published data only}

Najnin N, Leder K, Forbes A, Unicomb L, Winch PJ, Ram PK, et al. Impact of a large-scale handwashing intervention on reported respiratory illness: findings from a cluster-randomized controlled trial. *American Journal of Tropical Medicine and Hygiene* 2019;**100**(3):742-9.

Nicholson 2014 {published data only}

Nicholson JA, Naeeni M, Hoptroff M, Matheson JR, Roberts AJ, Taylor D, et al. An investigation of the effects of a hand washing intervention on health outcomes and school absence using a randomised trial in Indian urban communities. *Tropical Medicine and International Health* 2014;**19**(3):284-92. [DOI: [10.1111/tmi.12254](https://doi.org/10.1111/tmi.12254)]

Pandejpong 2012 {published data only}

Pandejpong D, Danchavijitr S, Vanprapa N, Pandejpong T, Cook EF. Appropriate time-interval application of alcohol hand gel on reducing influenza-like illness among preschool children: a randomized, controlled trial. *American Journal of Infection Control* 2012;**40**(6):507-11. [DOI: [10.1016/j.ajic.2011.08.020](https://doi.org/10.1016/j.ajic.2011.08.020)]

Priest 2014 {published data only}**12609000478213**

Priest P, McKenzie JE, Audas R, Poore M, Brunton C, Reeves L. Hand sanitiser provision for reducing illness absences in primary school children: a cluster randomised trial. *PLOS Medicine* 2014;**11**(8):e1001700.

Radonovich 2019 {published data only}

Radonovich LJ Jr, Simberkoff MS, Bessesen MT, Brown AC, Cummings DAT, Gaydos CA, et al, ResPECT investigators. N95 respirators vs medical masks for preventing influenza among health care personnel: a randomized clinical trial. *JAMA* 2019;**322**(9):824-33.

Ram 2015 {published data only}

Ram PK, DiVita MA, Khatun-e-Jannat K, Islam M, Krytun K, Cercone E, et al. Impact of intensive handwashing promotion on secondary household influenza-like illness in rural Bangladesh: findings from a randomized controlled trial. *PLOS ONE* 2015;**10**(6):e0125200. [DOI: [10.1371/journal.pone.0125200](https://doi.org/10.1371/journal.pone.0125200)]

Roberts 2000 {published data only}

Roberts L, Smith W, Jorm L, Patel M, Douglas RM, McGilchrist C. Effect of infection control measures on the frequency of upper respiratory infection in child care: a randomised controlled study. *Pediatrics* 2000;**105**:738-42.

Sandora 2005 {published data only}

Sandora TJ, Taveras EM, Shih MC, Resnick EA, Lee GM, Ross-Degnan D, et al. A randomized, controlled trial of a multifaceted intervention including alcohol-based hand sanitizer and hand-hygiene education to reduce illness transmission in the home. *Pediatrics* 2005;**116**(3):587-94.

Sandora 2008 {published data only}

Sandora TJ, Shih MC, Goldmann DA. Reducing absenteeism from gastrointestinal and respiratory illness in elementary school students: a randomized, controlled trial of an infection-control intervention. *Pediatrics* 2008;**121**(6):e1555-62.

Satomura 2005 {published data only}

Kitamura T, Satomura K, Kawamura T, Yamada S, Takashima K, Sukanuma N, et al. Can we prevent influenza-like illnesses by gargling? *Internal Medicine* 2007;**46**(18):1623-4.

* Satomura K, Kitamura T, Kawamura T, Shimbo T, Watanabe M, Kamei M, et al. Prevention of upper respiratory tract infections by gargling: a randomized trial. *American Journal of Preventive Medicine* 2005;**29**(4):302-7.

Savolainen-Kopra 2012 {published data only}

Savolainen-Kopra C, Haapakoski J, Peltola PA, Ziegler T, Korpela T, Anttila P, et al. Hand washing with soap and water together with behavioural recommendations prevents infections in common work environment: an open cluster-randomized trial. *Trials* 2012;**13**:10. [DOI: [10.1186/1745-6215-13-10](https://doi.org/10.1186/1745-6215-13-10)]

Simmerman 2011 {published data only}

Simmerman JM, Suntarattiwong P, Levy J, Jarman RG, Kaewchana S, Gibbons RV, et al. Findings from a household randomized controlled trial of hand washing and face masks to reduce influenza transmission in Bangkok, Thailand. *Influenza and Other Respiratory Viruses* 2011;**5**(4):256-67. [DOI: [10.1111/j.1750-2659.2011.00205.x](https://doi.org/10.1111/j.1750-2659.2011.00205.x)]

Stebbins 2011 {published data only}

Stebbins S, Cummings DA, Stark JH, Vukotich C, Mitruka K, Thompson W, et al. Reduction in the incidence of influenza A but not influenza B associated with use of hand sanitizer and cough hygiene in schools: a randomized controlled trial. *Pediatric Infectious Disease Journal* 2011;**30**(11):921-6. [DOI: [10.1097/INF.0b013e3182218656](https://doi.org/10.1097/INF.0b013e3182218656)]

Suess 2012 {published data only}

Suess T, Remschmidt C, Schink SB, Schweiger B, Nitsche A, Schroeder K, et al. The role of facemasks and hand hygiene in the prevention of influenza transmission in households: results from a cluster randomised trial; Berlin, Germany, 2009-2011. *BMC Infectious Disease* 2012;**12**:26. [DOI: [10.1186/1471-2334-12-26](https://doi.org/10.1186/1471-2334-12-26)]

Talaat 2011 {published data only}

Talaat M, Afifi S, Dueger E, El-Ashry N, Marfin A, Kandeel A, et al. Effects of hand hygiene campaigns on incidence of laboratory-confirmed influenza and absenteeism in schoolchildren, Cairo, Egypt. *Emerging Infectious Diseases* 2011;**17**(4):619-25. [DOI: [10.3201/eid1704.101353](https://doi.org/10.3201/eid1704.101353)]

Temime 2018 {published data only} **16474757**

Temime L, Cohen N, Ait-Bouziad K, Denormandie P, Dab W, Hocine MN. Impact of a multicomponent hand hygiene-related intervention on the infectious risk in nursing homes: a cluster randomized trial. *American Journal of Infection Control* 2018;**46**(2):173-9. [DOI: [10.1016/j.ajic.2017.08.030](https://doi.org/10.1016/j.ajic.2017.08.030)]

Turner 2004a {published data only}

Turner RB, Biedermann KA, Morgan JM, Keswick B, Ertel KD, Barker MF. Efficacy of organic acids in hand cleansers for prevention of rhinovirus infections. *Antimicrobial Agents and Chemotherapy* 2004;**48**(7):2595-8.

Turner 2004b {published data only}

Turner RB, Biedermann KA, Morgan JM, Keswick B, Ertel KD, Barker MF. Efficacy of organic acids in hand cleansers for prevention of rhinovirus infections. *Antimicrobial Agents and Chemotherapy* 2004;**48**(7):2595-8.

Turner 2012 {published data only}

Turner RB, Fuls JL, Rodgers ND, Goldfarb HB, Lockhart LK, Aust LB. A randomized trial of the efficacy of hand disinfection for prevention of rhinovirus infection. *Clinical Infectious Diseases* 2012;**54**(10):1422-6. [DOI: [10.1093/cid/cis201](https://doi.org/10.1093/cid/cis201)]

White 2001 {published data only}

White CG, Shinder FS, Shinder AL, Dyer DL. Reduction of illness absenteeism in elementary schools using an alcohol-free instant hand sanitizer. *Journal of School Nursing* 2001;**17**(5):258-65.

Yeung 2011 {published data only}

Yeung WK, Tam WS, Wong TW. Cluster randomized controlled trial of a hand hygiene intervention involving pocket-sized containers of alcohol-based hand rub for the control of infections in long-term care facilities. *Infection Control and Hospital Epidemiology* 2011;**32**:67-76.

Zomer 2015 {published data only} **NTR3000**

Zomer TP, Erasmus V, Looman CW, Tjon-A-Tsien A, Van Beeck EF, De Graaf JM, et al. A hand hygiene intervention to reduce infections in child daycare: a randomized controlled trial. *Epidemiology and Infection* 2015;**143**(12):2494-502. [DOI: [10.1017/S095026881400329X](https://doi.org/10.1017/S095026881400329X)]

References to studies excluded from this review
About El Hassan 2004 {published data only}

About El Hassan MA, van der Meulen-Muileman I, Abbas S, Kruyt FA. Conditionally replicating adenoviruses kill tumor cells via a basic apoptotic machinery-independent mechanism that resembles necrosis-like programmed cell death. *Journal of Virology* 2004;**78**(22):12243-51.

Amirav 2005 {published data only}

Amirav I, Oron A, Tal G, Cesar K, Ballin A, Houry S, et al. Aerosol delivery in respiratory syncytial virus bronchiolitis: hood or face mask? *Journal of Pediatrics* 2005;**147**(5):627-31.

Anderson 2004 {published data only}

Anderson RM, Fraser C, Ghani AC, Donnelly CA, Riley S, Ferguson NM, et al. Epidemiology, transmission dynamics and control of SARS: the 2002-2003 epidemic. *Philosophical Transactions of the Royal Society of London - Series B: Biological Sciences* 2004;**359**(1447):1091-105.

Anonymous 2002 {published data only}

Anon. Antiseptic skin cleansers may prevent rhinovirus transmission. *Clinical Infectious Diseases* 2002;**34**(3):ii.

Anonymous 2004 {published data only}

Anon. Can antiviral tissues prevent the spread of colds? *Consumer Reports* 2004;**69**(12):56.

Anonymous 2005a {published data only}

Anon. Antiviral Kleenex. *Medical Letter on Drugs & Therapeutics* 2005;**47**(1199):3-4.

Anonymous 2005b {published data only}

Anon. Focusing on this year's flu...use healthy handwashing...and when soap and water aren't handy, hand gels work well, too. *Child Health Alert* 2005;**23**:1-2.

Anonymous 2005c {published data only}

How long is a cold contagious? Is there any way to prevent transmitting a cold? Johns Hopkins Medical Letter, Health After 50 2005;**17**(10):8.

Apisarnthanarak 2009 {published data only}

Apisarnthanarak A, Apisarnthanarak P, Cheevakumjorn B, Mundy LM. Intervention with an infection control bundle to reduce transmission of influenza-like illnesses in a Thai preschool. *Infection Control and Hospital Epidemiology* 2009;**30**(9):817-22.

Apisarnthanarak 2010 {published data only}

Apisarnthanarak A, Uyeki TM, Puthavathana P, Kitphati R, Mundy LM. Reduction of seasonal influenza transmission among healthcare workers in an intensive care unit: a 4-year intervention study in Thailand. *Infection Control and Hospital Epidemiology* 2010;**31**(10):996-1003.

Aragon 2005 {published data only}

Aragon D, Sole ML, Brown S. Outcomes of an infection prevention project focusing on hand hygiene and isolation practices. *AACN Clinical Issues* 2005;**16**(2):121-3.

Azor-Martinez 2014 {published data only}

Azor-Martinez E, Gonzalez-Jimenez Y, Seijas-Vazquez ML, Cobos-Carrascosa E, Santisteban-Martinez J, Martinez-Lopez JM, et al. The impact of common infections on school absenteeism during an academic year. *American Journal of Infection Control* 2014;**42**(6):632-7.

Barros 1999 {published data only}

Barros AJ, Ross DA, Fonseca WV, Williams LA, Moreira-Filho DC. Preventing acute respiratory infections and diarrhoea in child care centres. *Acta Paediatrica* 1999;**88**(10):1113-8.

Bauer 2009 {published data only}

Bauer G, Bossi L, Santoalla M, Rodríguez S, Fariña D, Speranza AM. Impacto de un programa de prevención de infecciones respiratorias en lactantes prematuros de alto riesgo: estudio prospectivo y multicéntrico. *Archivos Argentinos de Pediatría* 2009;**107**(2):111-8.

Bell 2004 {published data only}

Bell DM. World Health Organization Working Group on International and Community Transmission of SARS. *Emerging Infectious Diseases* 2004;**10**(11):1900-6.

Bellissimo-Rodrigues 2009 {published data only}

Bellissimo-Rodrigues F, Bellissimo-Rodrigues WT, Viana JM, Teixeira GCA, Nicolini E, Auxiliadora-Martins M, et al. Effectiveness of oral rinse with chlorhexidine in preventing

nosocomial respiratory tract infections among intensive care unit patients. *Infection Control and Hospital Epidemiology* 2009;**30**(10):952-8.

Ben-Abraham 2002 {published data only}

Ben-Abraham R, Keller N, Szold O, Vardi A, Weinberg M, Barzilay Z, et al. Do isolation rooms reduce the rate of nosocomial infections in the pediatric intensive care unit? *Journal of Critical Care* 2002;**17**(3):176-80.

Black 1981 {published data only}

Black RE, Dykes AC, Anderson KE, Wells JG, Sinclair SP, Gary GW, et al. Handwashing to prevent diarrhea in day-care centers. *American Journal of Epidemiology* 1981;**113**(4):445-51.

Borkow 2010 {published data only}

Borkow G, Zhou SS, Page T, Gabbay J. A novel anti-influenza copper oxide containing respiratory face mask. *PLOS ONE* 2010;**5**(6):1-8.

Bouadma 2010 {published data only}

Bouadma L, Mourvillier B, Deiler V, Le Corre B, Lolom I, Régnier B, et al. A multifaceted program to prevent ventilator-associated pneumonia: impact on compliance with preventive measures. *Critical Care Medicine* 2010;**38**(3):789-96.

Bowen 2007 {published data only}

Bowen A, Ma H, Ou J, Billhimer W, Long T, Mintz E, et al. A cluster-randomized controlled trial evaluating the effect of a handwashing-promotion program in Chinese primary schools. *American Journal of Tropical Medicine and Hygiene* 2007;**76**(6):1166-73.

Breugelmans 2004 {published data only}

Breugelmans JG, Zucs P, Porten K, Broll S, Niedrig M, Ammon A, et al. SARS transmission and commercial aircraft. *Emerging Infectious Diseases* 2004;**10**(8):1502-3.

Cai 2009 {published data only}

Cai W, Schweiger B, Buchholz U, Buda S, Littmann M, Heusler J, et al. Protective measures and H5N1-seroprevalence among personnel tasked with bird collection during an outbreak of avian influenza A/H5N1 in wild birds, Ruegen, Germany, 2006. *BMC Infectious Diseases* 2009;**9**:170.

Cantagalli 2010 {published data only}

Cantagalli MR, Alvim VF, Andrade EC, Leite ICG. Associação entre desnutrição energético-proteica e infecção respiratória aguda em crianças na atenção primária à saúde. *Revista de APS* 2010;**13**(1):26-33.

Carbonell-Estrany 2008 {published data only}

Carbonell-Estrany X, Bont L, Doering G, Gouyon JB, Lanari M. Clinical relevance of prevention of respiratory syncytial virus lower respiratory tract infection in preterm infants born between 33 and 35 weeks gestational age. *European Journal of Clinical Microbiology and Infectious Diseases* 2008;**27**(10):891-9.

Carter 2002 {published data only}

Carter JM. Hand washing decreases risk of colds and flu. *Journal of the National Medical Association* 2002;**94**(2):A11.

Castillo-Chavez 2003 {published data only}

Castillo-Chavez C, Castillo-Garsow CW, Yakubu AA. Mathematical models of isolation and quarantine. *JAMA* 2003;**290**(21):2876-7.

Cava 2005a {published data only}

Cava MA, Fay KE, Beanlands HJ, McCay EA, Wignall R. Risk perception and compliance with quarantine during the SARS outbreak. *Journal of Nursing Scholarship* 2005;**37**(4):343-7.

Cava 2005b {published data only}

Cava MA, Fay KE, Beanlands HJ, McCay EA, Wignall R. The experience of quarantine for individuals affected by SARS in Toronto. *Public Health Nursing* 2005;**22**(5):398-406.

CDC 2003a {published data only}

Centers for Disease Control and Prevention. Cluster of severe acute respiratory syndrome cases among protected health-care workers - Toronto, Canada, April 2003. *Morbidity & Mortality Weekly Report* 2003;**52**(19):433-6.

CDC 2003b {published data only}

Centers for Disease Control and Prevention. Use of quarantine to prevent transmission of severe acute respiratory syndrome - Taiwan, 2003. *Morbidity & Mortality Weekly Report* 2003;**52**(29):680-3.

Chai 2005 {published data only}

Chai LY, Ng TM, Habib AG, Singh K, Kumarasinghe G, Tambyah PA. Paradoxical increase in methicillin-resistant *Staphylococcus aureus* acquisition rates despite barrier precautions and increased hand washing compliance during an outbreak of severe acute respiratory syndrome. *Clinical Infectious Diseases* 2005;**40**(4):623-3.

Chami 2012 {published data only}

Chami K, Gavazzi G, Bar-Hen A, Carrat F, de Wazières B, Lejeune B, et al. A short-term, multicomponent infection control program in nursing homes: a cluster randomized controlled trial. *Journal of the American Medical Directors Association* 2012;**13**(6):569.e9-569.e17.

Chaovavanich 2004 {published data only}

Chaovavanich A, Wongsawat J, Dowell SF, Inthong Y, Sangsajja C, Sanguanwongse N, et al. Early containment of severe acute respiratory syndrome (SARS); experience from Bamrasnaradura Institute, Thailand. *Journal of the Medical Association of Thailand* 2004;**87**(10):1182-7.

Chau 2003 {published data only}

Chau PH, Yip PS. Monitoring the severe acute respiratory syndrome epidemic and assessing effectiveness of interventions in Hong Kong Special Administrative Region. *Journal of Epidemiology and Community Health* 2003;**57**(10):766-9.

Chau 2008 {published data only}

Chau JPC, Thompson DR, Twinn S, Lee DTF, Lopez V, Ho LSY. An evaluation of SARS and droplet infection control practices in acute and rehabilitation hospitals in Hong Kong. *Hong Kong Medical Journal* 2008;**14**(Suppl 4):44-7.

Chen 2007 {published data only}

Chen YC, Chiang LC. Effectiveness of hand-washing teaching programs for families of children in paediatric intensive care units. *Journal of Clinical Nursing* 2007;**16**(6):1173-9.

Cheng 2010 {published data only}

Cheng VCC, Tai JWM, Wong LMW, Chan JFW, Li IWS, To KKW, et al. Prevention of nosocomial transmission of swine-origin pandemic influenza virus A/H1N1 by infection control bundle. *Journal of Hospital Infection* 2010;**74**(3):271-7.

Chia 2005 {published data only}

Chia SE, Koh D, Fones C, Qian F, Ng V, Tan BH, et al. Appropriate use of personal protective equipment among healthcare workers in public sector hospitals and primary healthcare polyclinics during the SARS outbreak in Singapore. *Occupational and Environmental Medicine* 2005;**62**(7):473-7.

Clynes 2010 {published data only}

Clynes N. Surgical masks vs N95 respirators for preventing influenza. *JAMA* 2010;**303**(10):937-9.

Cowling 2007 {published data only}

Cowling BJ, Muller MP, Wong IO, Ho LM, Louie M, McGeer A, et al. Alternative methods of estimating an incubation distribution: examples from severe acute respiratory syndrome. *Epidemiology* 2007;**18**(2):253-9.

Daniels 2010 {published data only}

Daniels TL, Talbot TR. Unmasking the confusion of respiratory protection to prevent influenza-like illness in crowded community settings. *Journal of Infectious Diseases* 2010;**201**(4):483-5.

Daugherty 2008 {published data only}

Daugherty EL. Health care worker protection in mass casualty respiratory failure: infection control, decontamination, and personal protective equipment. *Respiratory Care* 2008;**53**(2):201-12; discussion 212-4.

Davies 1994 {published data only}

Davies KJ, Herbert AM, Westmoreland D, Bagg J. Seroepidemiological study of respiratory virus infections among dental surgeons. *British Dental Journal* 1994;**176**(7):262-5.

Day 1993 {published data only}

Day RA, St Arnaud S, Monsma M. Effectiveness of a handwashing program. *Clinical Nursing Research* 1993;**2**(1):24-40.

Day 2006 {published data only}

Day T, Park A, Madras N, Gumel A, Wu J. When is quarantine a useful control strategy for emerging infectious diseases? *American Journal of Epidemiology* 2006;**163**(5):479-85.

Dell'Omodarme 2005 {published data only}

Dell'Omodarme M, Prati MC. The probability of failing in detecting an infectious disease at entry points into a country. *Statistics in Medicine* 2005;**24**(17):2669-79.

Denbak 2018 {published data only}

Denbak AM, Andersen A, Bonnesen CT, Laursen B, Ersbøll AK, Due P, et al. Effect evaluation of a randomized trial to reduce infectious illness and illness-related absenteeism among schoolchildren: The Hi Five Study. *Pediatric Infectious Disease Journal* 2018;**37**(1):16-21.

Desenclos 2004 {published data only}

Desenclos JC, van der Werf S, Bonmarin I, Levy-Bruhl D, Yazdanpanah Y, Hoen B, et al. Introduction of SARS in France, March-April, 2003. *Emerging Infectious Diseases* 2004;**10**(2):195-200.

DiGiovanni 2004 {published data only}

DiGiovanni C, Conley J, Chiu D, Zaborski J. Factors influencing compliance with quarantine in Toronto during the 2003 SARS outbreak. *Biosecurity and Bioterrorism* 2004;**2**(4):265-72.

Doebbeling 1992 {published data only}

Doebbeling BN, Stanley GL, Sheetz CT. Comparative efficacy of alternative hand-washing agents in reducing nosocomial infections in intensive care units. *New England Journal of Medicine* 1992;**327**(2):88-92.

Dwosh 2003 {published data only}

Dwosh HA, Hong HH, Austgarden D, Herman S, Schabas R. Identification and containment of an outbreak of SARS in a community hospital. *Canadian Medical Association Journal* 2003;**168**(11):1415-20.

Edmonds 2010 {published data only}

Edmonds S, Dzyakanava V, Macinga D. Efficacy of hand hygiene products against pandemic H1N1 influenza. *American Journal of Infection Control* 2010;**38**(5):E132-3.

Fendler 2002 {published data only}

Fendler EJ, Ali Y, Hammond BS, Lyons MK, Kelley MB, Vowell NA. The impact of alcohol hand sanitizer use on infection rates in an extended care facility. *American Journal of Infection Control* 2002;**30**(4):226-33.

Flint 2003 {published data only}

Flint J, Burton S, Macey JF, Deeks SL, Tam TW, King A, et al. Assessment of in-flight transmission of SARS - results of contact tracing. *Canadian Communicable Disease Report* 2003;**29**(12):105-10.

Fung 2004 {published data only}

Fung CP, Hsieh TL, Tan KH, Loh CH, Wu JS, Li CC, et al. Rapid creation of a temporary isolation ward for patients with severe acute respiratory syndrome in Taiwan. *Infection Control and Hospital Epidemiology* 2004;**25**(12):1026-32.

Garcia 2010 {published data only}

Garcia MC. The protection against influenza provided by surgical masks is not inferior to what the FFP2 (N95) respiratory protector can give. *FMC Formacion Medica Continuada en Atencion Primaria* 2010;**17**(5):365.

Gaydos 2001 {published data only}

Gaydos JC. Returning to the past: respiratory illness, vaccines, and handwashing. *American Journal of Preventive Medicine* 2001;**21**(2):150-1.

Gensini 2004 {published data only}

Gensini GF, Yacoub MH, Conti AA. The concept of quarantine in history: from plague to SARS. *Journal of Infection* 2004;**49**(4):257-61.

Girou 2002 {published data only}

Girou E, Loyeau S, Legrand P, Oppein F, Brun-Buisson C. Efficacy of handrubbing with alcohol based solution versus standard handwashing with antiseptic soap: randomised clinical trial. *BMJ* 2002;**325**:362.

Glass 2006 {published data only}

Glass K, Becker NG. Evaluation of measures to reduce international spread of SARS. *Epidemiology and Infection* 2006;**134**(5):1092-101.

Goel 2007 {published data only}

Goel S, Gupta AK, Singh A, Lenka SR. Preparations and limitations for prevention of severe acute respiratory syndrome in a tertiary care centre of India. *Journal of Hospital Infection* 2007;**66**(2):142-7.

Gomersall 2006 {published data only}

Gomersall CD, Joynt GM, Ho OM, Ip M, Yap F, Derrick JL, et al. Transmission of SARS to healthcare workers. The experience of a Hong Kong ICU. *Intensive Care Medicine* 2006;**32**(4):564-9.

Gore 2001 {published data only}

Gore J, Lambert JA. Does use of an instant hand sanitizer reduce elementary school illness absenteeism? *Journal of Family Practice* 2001;**50**(1):64.

Gostin 2003 {published data only}

Gostin LO, Bayer R, Fairchild AL. Ethical and legal challenges posed by severe acute respiratory syndrome: implications for the control of severe infectious disease threats. *JAMA* 2003;**290**(24):3229-37.

Gralton 2010 {published data only}

Gralton J, McLaws ML. Protecting healthcare workers from pandemic influenza: N95 or surgical masks? *Critical Care Medicine* 2010;**38**(2):657-67.

Guinan 2002 {published data only}

Guinan M, McGukin M, Ali Y. The effect of a comprehensive handwashing program on absenteeism in elementary schools. *American Journal of Infection Control* 2002;**30**(4):217-20.

Gupta 2005 {published data only}

Gupta AG, Moyer CA, Stern DT. The economic impact of quarantine: SARS in Toronto as a case study. *Journal of Infection* 2005;**50**(5):386-93.

Gwaltney 1982 {published data only}

Gwaltney JM Jr, Hendley JO. Transmission of experimental rhinovirus infection by contaminated surfaces. *American Journal of Epidemiology* 1982;**116**:828-33.

Han 2003 {published data only}

Han H, Li X, Qu W, Shen T, Xu F, Gao D, et al. The building and practice of the emergency isolation radiology information system at the department of radiology in polyclinic during the epidemic outbreak stage of SARS. *Beijing Da Xue Xue Bao [Journal of Peking University]* 2003;**35**(Suppl):86-8.

Hayden 1985 {published data only}

Hayden GF, Hendley JO, Gwaltney JM Jr. The effect of placebo and virucidal paper handkerchiefs on viral contamination of the hand and transmission of experimental rhinoviral infection. *Journal of Infectious Diseases* 1985;**152**(2):403-7.

Hendley 1988 {published data only}

Hendley JO, Gwaltney JM Jr. Mechanisms of transmission of rhinovirus infections. *Epidemiologic Reviews* 1988;**10**:243-58.

Hens 2009 {published data only}

Hens N, Ayele GM, Goeyvaerts N, Aerts M, Mossong J, Edmunds JW, et al. Estimating the impact of school closure on social mixing behaviour and the transmission of close contact infections in eight European countries. *BMC Infectious Diseases* 2009;**9**:187.

Heymann 2009 {published data only}

Heymann AD, Hoch I, Valinsky L, Kokia E, Steinberg DM. School closure may be effective in reducing transmission of respiratory viruses in the community. *Epidemiology and Infection* 2009;**137**(10):1369-76.

Hilburn 2003 {published data only}

Hilburn J, Hammond BS, Fendler EJ, Groziak PA. Use of alcohol hand sanitizer as an infection control strategy in an acute care facility. *American Journal of Infection Control* 2003;**31**(2):109-16.

Hilmarsson 2007 {published data only}

Hilmarsson H, Traustason BS, Kristmundsdottir T, Thormar H. Virucidal activities of medium- and long-chain fatty alcohols and lipids against respiratory syncytial virus and parainfluenza virus type 2: comparison at different pH levels. *Archives of Virology* 2007;**152**(1):2225-36.

Hirsch 2006 {published data only}

Hirsch HH, Steffen I, Francioli P, Widmer AF. Respiratory syncytial virus infections: measures in immunocompromised patients [Respiratorisches syncytial-virus (RSV)-infektion: massnahmen beim immunsupprimierten patienten]. *Praxis* 2006;**95**(3):61-6.

Ho 2003 {published data only}

Ho AS, Sung JJ, Chan-Yeung M. An outbreak of severe acute respiratory syndrome among hospital workers in a community hospital in Hong Kong. *Annals of Internal Medicine* 2003;**139**(7):564-7.

Hsieh 2007 {published data only}

Hsieh YH, King CC, Chen CW, Ho MS, Hsu SB, Wu YC. Impact of quarantine on the 2003 SARS outbreak: a retrospective modeling study. *Journal of Theoretical Biology* 2007;**244**(4):729-36.

Hugonnet 2007 {published data only}

Hugonnet S, Legros D, Roth C, Pessoa-Silva CL. Nosocomial transmission of severe acute respiratory syndrome: better quality of evidence is needed. *Clinical Infectious Diseases* 2007;**45**(12):1651.

Jiang 2003 {published data only}

* Jiang S, Huang L, Chen X, Wang J, Wu W, Yin S. Ventilation of wards and nosocomial outbreak of severe acute respiratory syndrome among healthcare workers. *Chinese Medical Journal* 2003;**116**(9):1293-7.

Jiang SP, Huang LW, Wang JF, Wu W, Yin SM, Chen WX. A study of the architectural factors and the infection rates of healthcare workers in isolation units for severe acute respiratory syndrome. *Chung-Hua Chieh Ho Ho Hu Hsi Tsa Chih [Chinese Journal of Tuberculosis & Respiratory Diseases]* 2003;**26**(10):594-7.

Johnson 2009 {published data only}

Johnson DF, Druce JD, Birch C, Grayson ML. A quantitative assessment of the efficacy of surgical and N95 masks to filter influenza virus in patients with acute influenza infection. *Clinical Infectious Diseases* 2009;**49**(2):275-7.

Jones 2005 {published data only}

Jones EW. "Co-operation in All Human Endeavour": quarantine and immigrant disease vectors in the 1918-1919 influenza pandemic in Winnipeg. *Canadian Bulletin of Medical History* 2005;**22**(1):57-82.

Kaydos-Daniels 2004 {published data only}

Kaydos-Daniels SC, Olowokure B, Chang HJ, Barwick RS, Deng JF, Lee ML, et al. Body temperature monitoring and SARS fever hotline, Taiwan. *Emerging Infectious Diseases* 2004;**10**(2):373-6.

Kelso 2009 {published data only}

Kelso JK, Milne GJ, Kelly H. Simulation suggests that rapid activation of social distancing can arrest epidemic development due to a novel strain of influenza. *BMC Public Health* 2009;**9**:117.

Khaw 2008 {published data only}

Khaw KS, Ngan Kee WD, Tam YH, Wong MK, Lee SW. Survey and evaluation of modified oxygen delivery devices used for suspected severe acute respiratory syndrome and other high-risk patients in Hong Kong. *Hong Kong Medical Journal* 2008;**14**(Suppl 5):27-31.

Kilabuko 2007 {published data only}

Kilabuko JH, Nakai S. Effects of cooking fuels on acute respiratory infections in children in Tanzania. *International Journal of Environmental Research and Public Health* 2007;**4**(4):283-8.

Kosugi 2004 {published data only}

Kosugi Y, Ishikawa T, Chimura Y, Annaka M, Shibasaki S, Adachi K, et al. Control of hospital infection of influenza: administration of neuraminidase inhibitor and cohort isolation of influenza patients. *Kansenshogaku Zasshi [Journal of the Japanese Association for Infectious Diseases]* 2004;**78**(12):995-9.

Lam 2004 {published data only}

Lam BC, Lee J, Lau YL. Hand hygiene practices in a neonatal intensive care unit: a multimodal intervention and impact on nosocomial infection. *Pediatrics* 2004;**114**(5):e565-71.

Lange 2004 {published data only}

Lange JH. Use of disposable face masks for public health protection against SARS. *Journal of Epidemiology and Community Health* 2004;**58**(5):434.

Larson 2004a {published data only}

Larson EL, Lin SX, Gomez-Pichardo C, Della-Latta P. Effect of antibacterial home cleaning and handwashing products on infectious disease symptoms: a randomized, double-blind trial. *Annals of Internal Medicine* 2004;**140**(5):321-9.

Larson 2004b {published data only}

Larson EL, Lin SX, Gomez-Pichardo C, Della-Latta P. Effect of antibacterial home cleaning and handwashing products on infectious disease symptoms: a randomized, double-blind trial. *Annals of Internal Medicine* 2004;**140**(5):321-9.

Larson 2005 {published data only}

Larson EL, Cimiotti J, Haas J, Parides M, Nesin M, Della-Latta P, et al. Effect of antiseptic handwashing vs alcohol sanitizer on health care-associated infections in neonatal intensive care units. *Archives of Pediatrics & Adolescent Medicine* 2005;**159**(4):377-83.

Lau 2004 {published data only}

Lau JT, Yang X, Tsui HY, Pang E. SARS related preventive and risk behaviours practised by Hong Kong-mainland China cross border travellers during the outbreak of the SARS epidemic in Hong Kong. *Journal of Epidemiology and Community Health* 2004;**58**(12):988-96.

Lau 2005 {published data only}

Lau JT, Leung PC, Wong EL, Fong C, Cheng KF, Zhang SC, et al. The use of an herbal formula by hospital care workers during the severe acute respiratory syndrome epidemic in Hong Kong to prevent severe acute respiratory syndrome transmission, relieve influenza-related symptoms, and improve quality of life: a prospective cohort study. *Alternative & Complementary Medicine* 2005;**11**(1):49-55.

Lee 2005 {published data only}

Lee GM, Salomon JA, Friedman JF, Hibberd PL, Ross-Degnan D, Zasloff E, et al. Illness transmission in the home: a possible role for alcohol-based hand gels. *Pediatrics* 2005;**115**(4):852-60.

Lee 2010 {published data only}

Lee VJ, Yap J, Cook AR, Chen MI, Tay JK, Barr I, et al. Effectiveness of public health measures in mitigating pandemic

influenza spread: a prospective sero-epidemiological cohort study. *Journal of Infectious Diseases* 2010;**202**(9):1319-26.

Lennell 2008 {published data only}

Lennell A, Kühlmann-Berenzon S, Geli P, Hedin K, Petersson C, Cars O, et al. Alcohol-based hand-disinfection reduced children's absence from Swedish day care centers. *Acta Paediatrica* 2008;**97**(12):1672-80.

Lipsitch 2003 {published data only}

Lipsitch M, Cohen T, Cooper B, Robins JM, Ma S, James L, et al. Transmission dynamics and control of severe acute respiratory syndrome. *Science* 2003;**300**(5627):1966-70.

Luckingham 1984 {published data only}

Luckingham B. To mask or not to mask: a note on the 1918 Spanish influenza epidemic in Tucson. *Journal of Arizona History* 1984;**25**(2):191-204.

Ma 2004 {published data only}

Ma HJ, Wang HW, Fang LQ, Jiang JF, Wei MT, Liu W, et al. A case-control study on the risk factors of severe acute respiratory syndromes among health care workers. *Chung-Hua Liu Hsing Ping Hsueh Tsa Chih [Chinese Journal of Epidemiology]* 2004;**25**(9):741-4.

MacIntyre 2010 {published data only}

MacIntyre CR. Cluster randomised controlled trial: hand hygiene and face mask use within 36 hours of index patient symptom onset reduces flu transmission to household contacts. *Evidence-Based Medicine* 2010;**15**(2):48-9.

Malone 2009 {published data only}

Malone JD, Brigantic R, Muller GA, Gadgil A, Delp W, McMahon BH, et al. U.S. airport entry screening in response to pandemic influenza: modeling and analysis. *Travel Medicine and Infectious Disease* 2009;**7**(4):181-91.

Marin 1991 {published data only}

Marin J, Dragas AZ, Mavsar B. Virus permeability of protective gloves used in medical practice. *Zentralblatt fur Hygiene und Umweltmedizin [International Journal of Hygiene and Environmental Medicine]* 1991;**191**(5-6):516-22.

McSweeney 2007 {published data only}

McSweeney K, Colman A, Fancourt N, Parnell M, Stantiall S, Rice G, et al. Was rurality protective in the 1918 influenza pandemic in New Zealand? *New Zealand Medical Journal* 2007;**120**(1256):2579.

Mielke 2009 {published data only}

Mielke M, Nassauer A. Pandemic influenza: nonpharmaceutical protective measures in ambulatory care. *Fortschritte der Medizin* 2009;**151**(40):32-4.

Mikolajczyk 2008 {published data only}

Mikolajczyk RT, Akmatov MK, Rastin S, Kretzschmar M. Social contacts of school children and the transmission of respiratory-spread pathogens. *Epidemiology and Infection* 2008;**136**(6):813-22.

- Monsma 1992** {published data only}
 Monsma M, Day R, St Arnaud S. Handwashing. *Journal of School Health* 1992;**62**(3):109-11.
- Nandrup-Bus 2009** {published data only}
 Nandrup-Bus I. Mandatory handwashing in elementary schools reduces absenteeism due to infectious illness among pupils: a pilot intervention study. *American Journal of Infection Control* 2009;**37**(10):820-6.
- Nishiura 2009** {published data only}
 Nishiura H, Wilson N, Baker MG. Quarantine for pandemic influenza control at the borders of small island nations. *BMC Infectious Diseases* 2009;**9**:27.
- O'Callaghan 1993** {published data only}
 O'Callaghan CA. Prevention of nosocomial respiratory syncytial virus infection. *Lancet* 1993;**341**(8838):182.
- Olsen 2003** {published data only}
 Olsen SJ, Chang HL, Cheung TY, Tang AF, Fisk TL, Ooi SP. Transmission of the severe acute respiratory syndrome on aircraft. *New England Journal of Medicine* 2003;**349**(25):2416-22.
- Ooi 2005** {published data only}
 Ooi PL, Lim S, Chew SK. Use of quarantine in the control of SARS in Singapore. *American Journal of Infection Control* 2005;**33**(5):252-7.
- Orellano 2010** {published data only}
 Orellano PW, Grassi A, Reynoso JI, Palmieri A, Uez O, Carlino O. Impact of school closings on the influenza A (H1N1) outbreak in Tierra del Fuego, Argentina. *Pan American Journal of Public Health* 2010;**27**(3):226-9.
- Panchabhai 2009** {published data only}
 Panchabhai TS, Dangayach NS, Krishnan A, Kothari VM, Karnad KR. Oropharyngeal cleansing with 0.2% chlorhexidine for prevention of nosocomial pneumonia in critically ill patients: an open-label randomized trial with 0.01% potassium permanganate as control. *Chest* 2009;**135**(5):1150-6.
- Pang 2004** {published data only}
 Pang XH, Liu DL, Gong XH, Xu FJ, Liu ZJ, Zhang Z, et al. Study on the risk factors related to severe acute respiratory syndrome among close contactors in Beijing. *Chung-Hua Liu Hsing Ping Hsueh Tsa Chih [Chinese Journal of Epidemiology]* 2004;**25**(8):674-6.
- Patel 2012** {published data only}
 Patel MK, Harris JR, Juliao P, Nygren B, Were V, Kola S, et al. Impact of a hygiene curriculum and the installation of simple handwashing and drinking water stations in rural Kenyan primary schools on student health and hygiene practices. *American Journal of Tropical Medicine and Hygiene* 2012;**87**(4):594-601.
- Pittet 2000** {published data only}
 Pittet D, Hugonnet S, Harbarth S, Mourouga P, Sauvan V, Touveneau S, et al. Effectiveness of a hospital-wide programme to improve compliance with hand hygiene. *Infection Control Programme* (Erratum in: *Lancet* 2000 Dec 23-30;356(9248):2196). *Lancet* 2000;**356**(9238):1307-12.
- Prasad 2004** {published data only}
 Prasad GV, Nash MM, Zaltzman JS. Handwashing precautions taken by renal transplant recipients for severe acute respiratory syndrome. *Transplantation* 2004;**77**(12):1917.
- Rabenau 2005** {published data only}
 Rabenau HF, Kampf G, Cinatl J, Doerr HW. Efficacy of various disinfectants against SARS coronavirus. *Journal of Hospital Infection* 2005;**61**(2):107-11.
- Reynolds 2008** {published data only}
 Reynolds DL, Garay JR, Deamond SL, Moran MK, Gold W, Styra R. Understanding, compliance and psychological impact of the SARS quarantine experience. *Epidemiology and Infection* 2008;**136**(7):997-1007.
- Richardson 2010** {published data only}
 Richardson A, Hofacre K. Comparison of filtration efficiency of a N95 filtering facepiece respirator and surgical mask against viral aerosols. *American Journal of Infection Control* 2010;**38**(5):E20.
- Riley 2003** {published data only}
 Riley S, Fraser C, Donnelly CA, Ghani AC, Abu-Raddad LJ, Hedley AJ. Transmission dynamics of the etiological agent of SARS in Hong Kong: impact of public health interventions. *Science* 2003;**300**(5627):1961-6.
- Rodriguez 2009** {published data only}
 Rodriguez CV, Rietberg K, Baer A, Kwan-Gett T, Duchin J. Association between school closure and subsequent absenteeism during a seasonal influenza epidemic. *Epidemiology* 2009;**20**(6):787-92.
- Rosen 2006** {published data only}
 Rosen L, Manor O, Engelhard D, Brody D, Rosen B, Peleg H, et al. Can a handwashing intervention make a difference? Results from a randomized controlled trial in Jerusalem preschools. *Preventive Medicine* 2006;**42**(1):27-32.
- Rosenthal 2005** {published data only}
 Rosenthal VD, Guzman S, Safdar N. Reduction in nosocomial infection with improved hand hygiene in intensive care units of a tertiary care hospital in Argentina. *American Journal of Infection Control* 2005;**33**(7):392-7.
- Safiulin 1972** {published data only}
 Safiulin AA. UV sanitation of the air and surfaces for the purpose of preventing in-hospital viral respiratory infections. *Gigiena i Sanitariia* 1972;**37**(10):99-101.
- Sandrock 2008** {published data only}
 Sandrock C, Stollenwerk N. Acute febrile respiratory illness in the ICU: reducing disease transmission. *Chest* 2008;**133**(5):1221-31.
- Sattar 2000** {published data only}
 Sattar SA, Abebe M, Bueti AJ, Jampani H, Newman J, Hua S. Activity of an alcohol-based hand gel against human adeno-

rhino-, and rotaviruses using the fingerpad method. *Infection Control and Hospital Epidemiology* 2000;**21**(8):516-9.

Schull 2007 {published data only}

Schull MJ, Stukel TA, Vermeulen MJ, Zwarenstein M, Alter DA, Manuel DG, et al. Effect of widespread restrictions on the use of hospital services during an outbreak of severe acute respiratory syndrome. *CMAJ* 2007;**176**(13):1827-32.

Seal 2010 {published data only}

Seal LA, Slade B, Cargill I, Aust D. Persistence factors in handwashes. *American Journal of Infection Control* 2010;**38**(5):E18-9.

Seale 2009 {published data only}

Seale H, Corbett S, Dwyer DE, MacIntyre CR. Feasibility exercise to evaluate the use of particulate respirators by emergency department staff during the 2007 influenza season. *Infection Control and Hospital Epidemiology* 2009;**30**(7):710-2.

Sizun 1996 {published data only}

Sizun J, Baron R, Soupre D, Giroux JD, de Parscau L. Nosocomial infections due to syncytial respiratory virus: which hygienic measures. *Archives de Pédiatrie* 1996;**3**(7):723-7.

Slayton 2016 {published data only}

Slayton RB, Murphy JL, Morris J, Faith SH, Oremo J, Odhiambo A, et al. A cluster randomized controlled evaluation of the health impact of a novel antimicrobial hand towel on the health of children under 2 years old in rural communities in Nyanza Province, Kenya. *American Journal of Tropical Medicine and Hygiene* 2016;**94**(2):437-44.

Stebbins 2009 {published data only}

Stebbins S, Downs JS, Vukotich CJ. Using nonpharmaceutical interventions to prevent influenza transmission in elementary school children: parent and teacher perspectives. *Journal of Public Health Management and Practice* 2009;**15**(2):112-7.

Stedman-Smith 2015 {published data only}

Stedman-Smith M, DuBois CL, Grey SF, Kingsbury DM, Shakya S, Scofield J, et al. Outcomes of a pilot hand hygiene randomized cluster trial to reduce communicable infections among US office-based employees. *Journal of Occupational and Environmental Medicine* 2015;**57**(4):374-80.

Stoner 2007 {published data only}

Stoner MJ, Cohen DM, Fernandez S, Bonsu BK. Physician handwashing: what do parents want? *Journal of Hospital Infection* 2007;**65**(2):112-6.

Stukel 2008 {published data only}

Stukel TA, Schull MJ, Guttman A, Alter DA, Li P, Vermeulen MJ, et al. Health impact of hospital restrictions on seriously ill hospitalized patients: lessons from the Toronto SARS outbreak. *Medical Care* 2008;**46**(9):991-7.

Svoboda 2004 {published data only}

Svoboda T, Henry B, Shulman L, Kennedy E, Rea E, Ng W, et al. Public health measures to control the spread of the severe

acute respiratory syndrome during the outbreak in Toronto. *New England Journal of Medicine* 2004;**350**(23):2353-61.

Tracht 2010 {published data only}

Tracht SM, Del Valle SY, Hyman JM. Mathematical modeling of the effectiveness of facemasks in reducing the spread of novel influenza a (H1N1). *PLOS ONE* 2010;**5**(2):e9018.

Ueno 1990 {published data only}

Ueno T, Saijo K. Prevention of adenovirus infection and antiviral activity of a hand disinfectant, Welpas. *Nippon Ganka Gakkai Zasshi* 1990;**94**(1):44-8.

Uhari 1999 {published data only}

Uhari M, Mottonen M. An open randomized controlled trial of infection prevention in child day-care centers. *Pediatric Infectious Disease Journal* 1999;**18**(8):672-7.

van der Sande 2008 {published data only}

van der Sande M, Teunis P, Sabel R. Professional and home-made face masks reduce exposure to respiratory infections among the general population. *PLOS ONE* 2008;**3**(7):e2618.

Vessey 2007 {published data only}

Vessey JA, Sherwood JJ, Warner D, Clark D. Comparing hand washing to hand sanitizers in reducing elementary school students' absenteeism. *Pediatric Nursing* 2007;**33**(4):368-72.

Viscusi 2009a {published data only}

Viscusi DJ, Bergman M, Sinkule E, Shaffer RE. Evaluation of the filtration performance of 21 N95 filtering face piece respirators after prolonged storage. *American Journal of Infection Control* 2009;**37**(5):381-6.

Viscusi 2009b {published data only}

Viscusi DJ, Bergman MS, Eimer BC, Shaffer RE. Evaluation of five decontamination methods for filtering facepiece respirators. *Annals of Occupational Hygiene* 2009;**53**(8):815-27.

Wang 2003 {published data only}

Wang JX, Feng HY, Liu D, Zhang ZL, Shan AL, Zhu XJ, et al. Epidemiological characteristics of severe acute respiratory syndrome in Tianjin and the assessment of effectiveness on measures of control. *Chung-Hua Liu Hsing Ping Hsueh Tsa Chih [Chinese Journal of Epidemiology]* 2003;**24**(7):565-9.

Wang 2005 {published data only}

Wang HW, He J, Zhang PH, Tang F, Wang TB, Luan YH, et al. A case-control study on the mxA polymorphisms and susceptibility to severe acute respiratory syndromes. *Chung-Hua Liu Hsing Ping Hsueh Tsa Chih [Chinese Journal of Epidemiology]* 2005;**26**(8):574-7.

Weber 2004 {published data only}

Weber JT, Hughes JM. Beyond Semmelweis: moving infection control into the community. *Annals of Internal Medicine* 2004;**140**(5):397-8.

Wen 2010 {published data only}

Wen Z, Lu J, Li J, Li N, Zhao J, Wang J, et al. Determining the filtration efficiency of half-face medical protection mask (N99) against viral aerosol. *Aerobiologia* 2010;**26**(3):245-51.

White 2005 {published data only}

White C, Kolble R, Carlson R, Lipson N. The impact of a health campaign on hand hygiene and upper respiratory illness among college students living in residence halls. *Journal of American College Health* 2005;**53**(4):175-81.

Wilczynski 1997 {published data only}

Wilczynski J, Torbicka E, Brzozowska-Binda A, Szymanska U. Breast feeding for prevention of viral acute respiratory diseases in infants. *Medycyna Doswiadczalna i Mikrobiologia* 1997;**49**(3-4):199-206.

Wilder-Smith 2003 {published data only}

Wilder-Smith A, Paton NI, Goh KT. Low risk of transmission of severe acute respiratory syndrome on airplanes: the Singapore experience. *Tropical Medicine and International Health* 2003;**8**(11):1035-7. [MEDLINE: 14629772]

Wilder-Smith 2005 {published data only}

Wilder-Smith A, Low JG. Risk of respiratory infections in health care workers: lessons on infection control emerge from the SARS outbreak. *Southeast Asian Journal of Tropical Medicine and Public Health* 2005;**36**(2):481-8.

Wong 2005 {published data only}

Wong TW, Tam WW. Handwashing practice and the use of personal protective equipment among medical students after the SARS epidemic in Hong Kong. *American Journal of Infection Control* 2005;**33**(10):580-6.

Yen 2010 {published data only}

Yen M-Y, Lu Y-C, Huang P-H, Chen C-M, Chen Y-C, Lin YE. Quantitative evaluation of infection control models in the prevention of nosocomial transmission of SARS virus to healthcare workers: implication to nosocomial viral infection control for healthcare workers. *Scandinavian Journal of Infectious Diseases* 2010;**42**(6-7):510-5.

Yu 2004 {published data only}

Yu IT, Li Y, Wong TW, Tam W, Chan AT, Lee JH, et al. Evidence of airborne transmission of the severe acute respiratory syndrome virus. *New England Journal of Medicine* 2004;**350**(17):1731-9.

Zamora 2006 {published data only}

Zamora JE, Murdoch J, Simchison B, Day AG. Contamination: a comparison of 2 personal protective systems. *Canadian Medical Association Journal* 2006;**175**(3):249-54.

Zhai 2007 {published data only}

Zhai S, Liu W, Yan B. Recent patents on treatment of severe acute respiratory syndrome (SARS). *Recent Patents on Anti-infective Drug Discovery* 2007;**2**(1):1-10.

Zhao 2003 {published data only}

Zhao Z, Zhang F, Xu M, Huang K, Zhong W, Cai W, et al. Description and clinical treatment of an early outbreak of

severe acute respiratory syndrome (SARS) in Guangzhou, PR China. *Journal of Medical Microbiology* 2003;**52**(8):715-20.

References to ongoing studies
NCT03454009 {published data only}

NCT03454009. Kent State University / Price Chopper Wellness Promotion Study. clinicaltrials.gov/show/NCT03454009 (first date received March 5 2018).

NCT04267952 {published data only}

NCT04267952. Hand hygiene intervention program on primary school students' health outcomes and absenteeism in school. [Clinicaltrials.gov/show/NCT04267952](https://clinicaltrials.gov/show/NCT04267952) (first received Feb 13 2020).

NCT04296643 {published data only}

NCT04296643. Medical masks vs N95 respirators for COVID-19. [Clinicaltrials.gov/show/NCT04296643](https://clinicaltrials.gov/show/NCT04296643) (first received March 5, 2020).

NCT04337541 {published data only}

NCT04337541. Reduction in COVID-19 infection using surgical facial masks outside the healthcare system. clinicaltrials.gov/show/NCT04337541 (first received 7 April 2020).

NCT04471766 {published data only}

NCT04471766. Locally produced cloth face mask and COVID-19 like illness prevention. clinicaltrials.gov/ct2/show/NCT04471766 (first received July 15, 2020). [TRIAL REGISTRY: NCT04471766]

Wang 2015 {published data only}

Wang M, Barasheed O, Rashid H, Booy R, El Bashir H, Haworth E, et al. A cluster-randomised controlled trial to test the efficacy of facemasks in preventing respiratory viral infection among Hajj pilgrims. *Journal of Epidemiology and Global Health* 2015;**5**(2):181-9.

Additional references
ACTRN12610000887077

ACTRN12610000887077. Face masks in the protection of healthcare workers to pandemic influenza and emerging infections [Economic, social and cross cultural issues in non-pharmaceutical protection of front line responders to pandemic influenza and emerging infections]. www.anzctr.org.au/Trial/Registration/TrialReview.aspx?id=336078 (first received 18/10/2010).

Aiello 2002

Aiello AE, Larson EL. What is the evidence for a causal link between hygiene and infections? *Lancet* 2002;**2**(2):103-10.

Atkins 2004

Atkins D, Best D, Briss PA, Eccles M, Falck-Ytter Y, Flottorp S, et al, GRADE Working Group. Grading quality of evidence and strength of recommendations. *BMJ* 2004;**328**(7454):1490.

Bartoszko 2020

Bartoszko JJ, Farooqi MAM, Alhazzani W, Loeb M. Medical masks vs N95 respirators for preventing COVID-19 in healthcare workers: a systematic review and meta-analysis of randomized trials. *Influenza and Other Respiratory Viruses* 2020;**14**(4):365-73. [DOI: [10.1111/irv.12745](https://doi.org/10.1111/irv.12745)] [PMID: 32246890]

Bonn 1997

Bonn D. Spared an influenza pandemic for another year? *Lancet* 1997;**349**(9044):36.

Bootsma 2007

Bootsma MCJ, Ferguson NM. The effect of public health measures on the 1918 influenza pandemic in U.S. cities. Proceedings of the National Academy of Sciences of the United States of America (PNAS) 2007;**E-pub ahead of print**:1-6. [DOI: [10.1073/pnas.06110711104](https://doi.org/10.1073/pnas.06110711104)]

Brosseau 2020

Brosseau LM, Sietsema M. Masks-for-all for COVID-19 not based on sound data. www.cidrap.umn.edu/news-perspective/2020/04/commentary-masks-all-covid-19-not-based-sound-data (accessed 20 April 2020).

Cassell 2006

Cassell MM, Halperin DT, Shelton JD, Stanton D. Risk compensation: the achilles' heel of innovations in HIV prevention? *BMJ* 2006;**332**:605-7.

CDC 2005a

Centers for Disease Control and Prevention. Infection control guidance for the prevention and control of influenza in acute-care facilities. www.cdc.gov/flu/professionals/infectioncontrol/healthcarefacilities.htm (accessed 3 March 2006).

CDC 2005b

Centers for Disease Control and Prevention. 2004-2005 Interim guidance for the use of masks to control influenza transmission. www.cdc.gov/flu/professionals/infectioncontrol/maskguidance.htm (accessed 3 March 2006).

Chan 2020

Chan J, Yuan S, Kok KH, To K, Chu H, Yang J, et al. A familial cluster of pneumonia associated with the 2019 novel coronavirus indicating person-to-person transmission: a study of a family cluster. *Lancet* 2020;**395**(10223):514-24.

Chard 2018

Chard AN, Freeman MC. Design, intervention fidelity, and behavioral outcomes of a school-based water, sanitation, and hygiene cluster-randomized trial in Laos. *International Journal of Environmental Research and Public Health* 2018;**15**(4):570.

Chu 2020

Chu DK, Akl EA, Duda S, Solo K, Yaacoub S, Schünemann HJ. Physical distancing, face masks, and eye protection to prevent person-to-person transmission of SARS-CoV-2 and COVID-19: a systematic review and meta-analysis. *Lancet* 2020;**395**(10242):1973-87. [DOI: [10.1016/S0140-6736\(20\)31142-9](https://doi.org/10.1016/S0140-6736(20)31142-9)]

Claessen 2008

Claessen J-P, Bates S, Sherlock K, Seeparsand F, Wright R. Designing interventions to improve tooth brushing. *International Dental Journal* 2008;**58**(S5):307-20. *International Dental Journal* 2008;**58**(Suppl 5):307-20.

Clark 2020

Clark JM, Sanders S, Carter M, Honeyman D, Cleo G, Auld Y, et al. Improving the translation of search strategies using the Polyglot Search Translator: a randomized controlled trial. *Journal of the Medical Library Association* 2020;**108**(2):195-207.

Curtis 2001

Curtis V, Biran A. Dirt, disgust, and disease: Is hygiene in our genes? *Perspectives in Biology and Medicine* 2001;**44**(1):17-31.

Demicheli 2018a

Demicheli V, Jefferson T, Ferroni E, Rivetti A, Di Pietrantonj C. Vaccines for preventing influenza in healthy adults. *Cochrane Database of Systematic Reviews* 2018, Issue 2. Art. No: CD001269. [DOI: [10.1002/14651858.CD001269.pub6](https://doi.org/10.1002/14651858.CD001269.pub6)]

Demicheli 2018b

Demicheli V, Jefferson T, Di Pietrantonj C, Ferroni E, Thorning S, Thomas RE, et al. Vaccines for preventing influenza in the elderly. *Cochrane Database of Systematic Reviews* 2018, Issue 2. Art. No: CD004876. [DOI: [10.1002/14651858.CD004876.pub4](https://doi.org/10.1002/14651858.CD004876.pub4)]

Donovan 2007

Donovan J, Kudla I, Holness L, Skotnicki-Grant D, Nethercott S, James R. Skin reactions following use of N95 facial masks. *Dermatitis* 2007;**18**(2):104.

Dreibelbis 2013

Dreibelbis R, Winch PJ, Leontsini E, Hulland KR, Ram PK, Unicom L, et al. The integrated behavioural model for water, sanitation, and hygiene: a systematic review of behavioural models and a framework for designing and evaluating behaviour change interventions in infrastructure restricted settings. *BMC Public Health* 2013;**13**:1015.

Elflein 2019

Elflein J. Number of influenza cases in the United States from 2010-2017. www.statista.com/statistics/861113/estimated-number-of-flu-cases-us/ (accessed prior to 2 November 2020).

Ellis 2010

Ellis M. Evaluating strategies to prevent methicillin-resistant Staphylococcus Aureus skin and soft tissue infections in military trainees (Methicillin-resistant Staphylococcus Aureus (MRSA) skin and soft tissue infection (SSTI) prevention in military trainees]. clinicaltrials.gov/ct2/show/study/NCT01105767 April 16, 2010.

Erasmus 2011

Erasmus V, Huis A, Oenema A, Van Empelen P, Boog MC, Van Beeck EH, et al. The ACCOMPLISH study. A cluster randomised trial on the cost-effectiveness of a multicomponent intervention to improve hand hygiene compliance and reduce healthcare associated infections. *BMC Public Health [Internet]* 2011;**11**:721.

Fong 2020

Fong MW, Gao H, Wong JY, Xiao J, Shiu EY, Ryu S, et al. Nonpharmaceutical measures for pandemic influenza in nonhealthcare settings - social distancing measures. *Emerging Infectious Diseases* 2020;**26**(5):976-84.

Fung 2006

Fung IC, Cairncross S. Effectiveness of handwashing in preventing SARS: a review. *Tropical Medicine and International Health* 2006;**11**(11):1749-58.

Glanz 2008

Glanz K, Rimer BK, Viswanath K, editors. Health Behavior and Health Education: Theory, Research, and Practice. 4th edition. San Francisco (CA): Jossey-Bass, 2008.

GRADEpro GDT 2015 [Computer program]

McMaster University (developed by Evidence Prime) GRADEpro GDT. Version accessed 27 April 2020. Hamilton (ON): McMaster University (developed by Evidence Prime), 2015. Available at grade.pro.

Greenhalgh 2020

Greenhalgh T, Schmid MB, Czypionka T, Bassler D, Gruer L. Face masks for the public during the covid-19 crisis. *BMJ* 2020;**369**:m1435.

Hartinger 2011

Hartinger SM, Lanata CF, Hattendorf J, Gil AI, Verastegui H, Ochoa T, et al. A community randomised controlled trial evaluating a home-based environmental intervention package of improved stoves, solar water disinfection and kitchen sinks in rural Peru: Rationale, trial design and baseline findings. *Contemporary Clinical Trials* 2011;**32**(6):864-73.

Hartinger 2012

Hartinger SM, Lanata CF, Gil AI, Hattendorf J, Verastegui H, Mäusezahl D. Combining interventions: Improved chimney stoves, kitchen sinks and solar disinfection of drinking water and kitchen clothes to improve home hygiene in rural Peru. *Field Actions Science Report* <http://factsreports.revues.org/1627> accessed 3 May 2013;**6**:1 SPL.

Higgins 2002

Higgins JP, Thompson SG. Quantifying heterogeneity in a meta-analysis. *Statistics in Medicine* 2002;**21**(11):1539-58.

Higgins 2003

Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. *BMJ* 2003;**327**(7414):557-60.

Higgins 2011

Higgins JP, Green S, editor(s). Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 (updated March 2011). The Cochrane Collaboration, 2011. Available from handbook.cochrane.org.

Howard 2020

Howard J, Huang A, Li Z, Tufekci Z, Zdimal V, van der Westhuizen H, et al. Face masks against COVID-19: an evidence review. www.preprints.org/

manuscript/202004.0203/v1 (accessed 20 April 2020). [DOI: [10.20944/preprints202004.0203.v1](https://doi.org/10.20944/preprints202004.0203.v1)]

Huis 2013

Huis A, Schoonhoven L, Grol R, Donders R, Hulscher M, van Achterberg T. Impact of a team and leaders-directed strategy to improve nurses' adherence to hand hygiene guidelines: a cluster randomised trial. *International Journal of Nursing Studies* 2013;**50**:464-74.

Hulland 2013

Hulland KR, Leontsini E, Dreibelbis R, Unicomb L, Afroz A, Dutta NC, et al. Designing a handwashing station for infrastructure-restricted communities in Bangladesh using the integrated behavioural model for water, sanitation and hygiene interventions (IBM-WASH). *BMC Public Health* 2013;**13**(1):877.

Jefferson 2006b

Jefferson T, Demicheli V, Rivetti D, Jones M, Di Pietrantonj C, Rivetti A. Antivirals for influenza in healthy adults: systematic review. *Lancet* 2006;**367**(9507):303-13. [DOI: [10.1016/S0140-6736\(06\)67970-1](https://doi.org/10.1016/S0140-6736(06)67970-1)]

Jefferson 2014

Jefferson T, Jones M, Doshi P, Del Mar C, Hama R, Thompson M, et al. Neuraminidase inhibitors for preventing and treating influenza in adults and children. *Cochrane Database of Systematic Reviews* 2014, Issue 4. Art. No: CD008965. [DOI: [10.1002/14651858.CD008965.pub4](https://doi.org/10.1002/14651858.CD008965.pub4)]

Jefferson 2018

Jefferson T, Rivetti A, Di Pietrantonj C, Demicheli V. Vaccines for preventing influenza in healthy children. *Cochrane Database of Systematic Reviews* 2018, Issue 2. Art. No: CD004879. [DOI: [10.1002/14651858.CD004879.pub5](https://doi.org/10.1002/14651858.CD004879.pub5)]

Kaewchana 2012

Kaewchana S, Simmerman M, Somrongthong R, Suntarattiwong P, Lertmaharit S, Chotipitayasonondh T. Effect of intensive hand washing education on hand washing behaviors in Thai households with an influenza-positive child in urban Thailand. *Asia-Pacific Journal of Public Health* 2012;**24**(4):577-85.

Kendrick 1994

Kendrick AS. Training to ensure healthy child day care programs. *Pediatrics* 1994;**94**(6 II):1108-110.

Kirscht 1974

Kirscht JP. The Health Belief Model and illness behavior. *Health Education & Behavior* 1974;**2**(4):387-408.

Kitamura 2007

Kitamura T, Satomura K, Kawamura T, Yamada S, Takashima K, Sukanuma N. Can we prevent influenza-like illnesses by gargling? *Japanese Society of Internal Medicine* 2007;**46**(18):1623-4. [DOI: [10.2169/internalmedicine.46.0104](https://doi.org/10.2169/internalmedicine.46.0104)]

Lefebvre 2011

Lefebvre C, Manheimer E, Glanville J. Chapter 6: Searching for studies. In: Higgins JP, Green S, editor(s). Cochrane Handbook

for Systematic Reviews of Interventions Version 5.1.0 (updated March 2011). The Cochrane Collaboration, 2011. Available from handbook.cochrane.org.

Long 2020

Long Y, Hu T, Liu L, Chen R, Guo Q, Yang L, et al. Effectiveness of N95 respirators versus surgical masks against influenza: a systematic review and meta-analysis. *Journal of Evidence-Based Medicine* 2020;**13**(2):93-101. [DOI: [10.1111/jebm.12381](https://doi.org/10.1111/jebm.12381)]

Marshall 2018

Marshall IJ, A Noel-Storr A, Kuiper J, Thomas J, Wallace BD. Machine learning for identifying randomized controlled trials: an evaluation and practitioner's guide [10.1002/jrsm.1287]. *Research Synthesis Methods* 2018;**9**(4):602-14. [DOI: [10.1002/jrsm.1287](https://doi.org/10.1002/jrsm.1287)]

Mbakaya 2017

Mbakaya BC, Lee PH, Lee RL. Hand hygiene intervention strategies to reduce diarrhoea and respiratory infections among schoolchildren in developing countries: a systematic review. *International Journal of Environmental Research and Public Health* 2017;**14**(4):371.

McKenzie 2010

McKenzie JE, Priest P, Audas R, Poore MR, Brunton CR, Reeves LM. Hand sanitisers for reducing illness absences in primary school children in New Zealand: a cluster randomised controlled trial study protocol. *Trials* 2010;**11**:7.

Meadows 2004

Meadows E, Le Saux N. A systematic review of the effectiveness of antimicrobial rinse-free hand sanitizers for prevention of illness-related absenteeism in elementary school children. *BMC Public Health* 2004;**4**:50. [DOI: [10.1186/1471-2458-4-50](https://doi.org/10.1186/1471-2458-4-50)]

Microsoft Excel

Microsoft Corporation. Microsoft Excel. Available from: <https://office.microsoft.com/excel> 2018.

Moncion 2019

Moncion K, Young K, Tunis M, Rempel S, Stirling R, Zhao L. Effectiveness of hand hygiene practices in preventing influenza virus infection in the community setting: a systematic review. *Canada Communicable Disease Report* 2019;**45**(1):12-23.

Monto 1969

Monto AS, Davenport FM, Napier JA, Francis T Jr. Effect of vaccination of a school-age population upon the course of an A2-Hong Kong influenza epidemic. *Bulletin of the World Health Organization* 1969;**41**(3):537-42.

Najnin 2017

Najnin N, Leder K, Qadri F, Forbes A, Unicomb L, Winch PJ, et al. Impact of adding hand-washing and water disinfection promotion to oral cholera vaccination on diarrhoea-associated hospitalization in Dhaka, Bangladesh: Evidence from a cluster randomized control trial International. *Journal of Epidemiology* 2017;**46**(6):2056-66.

Offeddu 2017

Offeddu V, Yung CF, Low MS, Tam CC. Effectiveness of masks and respirators against respiratory infections in healthcare workers: a systematic review and meta-analysis. *Clinical Infectious Diseases* 2017;**65**(11):1934-42. [DOI: [10.1093/cid/cix681](https://doi.org/10.1093/cid/cix681)]

Ong 2020

Ong SWX, Tan YK, Chia PY, Lee TH, Ng OT, Wong MSY, et al. Air, surface environmental, and personal protective equipment contamination by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) from a symptomatic patient. *JAMA* 2020;**323**(16):1610-2.

Perkins 2003

Perkins HW, editor. The Social Norms Approach to Preventing School and College Age Substance Abuse: a Handbook for Educators, Counselors, and Clinicians. San Francisco (CA): Jossey-Bass, 2003.

Qadri 2015

Qadri F, Ali M, Chowdhury F, Khan AI, Saha A, Khan IA, et al. Feasibility and effectiveness of oral cholera vaccine in an urban endemic setting in Bangladesh: A cluster randomised open-label trial. *Lancet* 2015;**386**(10001):1362-71.

Rabie 2006

Rabie T, Curtis V. Handwashing and risk of respiratory infections: a quantitative systematic review. *Tropical Medicine & International Health* 2006;**11**(3):258-67.

Radonovich 2016

Radonovich LJ Jr, Bessesen MT, Cummings DA, Eagan A, Gaydos A, Gibert C, et al. The respiratory protection effectiveness clinical trial (ResPECT): a cluster-randomized comparison of respirator and medical mask effectiveness against respiratory infections in healthcare personnel. *BMC Infectious Diseases* 2016;**16**:243. [DOI: [10.1186/s12879-016-1494-2](https://doi.org/10.1186/s12879-016-1494-2)]

Ran 2020

Ran L, Chen X, Wang Y, Wu W, Zhang L, Tan X. Risk factors of healthcare workers with corona virus disease 2019: a retrospective cohort study in a designated hospital of Wuhan in China. *Clinical Infectious Diseases* 2020 Mar 17 [Epub ahead of print]. [DOI: [10.1093/cid/ciaa287](https://doi.org/10.1093/cid/ciaa287)] [PMID: 32179890]

Rebmann 2013

Rebmann T, Carrico R, Wang J. Physiologic and other effects and compliance with long-term respirator use among medical intensive care unit nurses. *American Journal of Infection Control* 2013;**41**(12):1218-23.

Rengasamy 2010

Rengasamy S, Eimer B, Shaffer RE. Simple respiratory protection - evaluation of the filtration performance of cloth masks and common fabric materials against 20-1000 nm size particles. *Annals of Occupational Hygiene* 2010;**54**(7):789-98.

RIVM 2011

RIVM. Health risks in a child daycare centre or preschool (0 to 4 year olds). Dutch National Institute for Public Health and the Environment (RIVM), Dutch National Centre for Hygiene and Safety (LCHV). Bilthoven (The Netherlands): The Institute 2011.

Saunders-Hastings 2017

Saunders-Hastings P, Crispo JAG, Sikora L, Krewski D. Effectiveness of personal protective measures in reducing pandemic influenza transmission: a systematic review and meta-analysis. *Epidemics* 2017;**20**:1-20.

Savolainen-Kopra 2010

Savolainen-Kopra C, Haapakoski J, Peltola PA, Ziegler T, Korpela T, Anttila P, et al. STOPFLU: is it possible to reduce the number of days off in office work by improved hand-hygiene? *Trials* 2010;**11**:69.

Schwartz 2018

Schwartz JL. The Spanish Flu, epidemics, and the turn to biomedical responses. *American Journal of Public Health* 2018;**108**(11):1455-8.

Sidibe 2003

Sidibe M. Formative Research in Schools: Review Document. London (UK): London School of Hygiene & Tropical Medicine, 2003.

Sidibe 2009

Sidibe M, Shukla N, Rustagi A et al. Inducing the habit of soap usage in an urban community setting. In: Proceedings of the 3rd WHO International Conference on Children's Health and the Environment. Busan (Korea), 2009.

Smith 2016

Smith JD, MacDougall CC, Johnstone J, Copes RA, Schwartz B, Garber GE. Effectiveness of N95 respirators versus surgical masks in protecting health care workers from acute respiratory infection: a systematic review and meta-analysis. *Canadian Medical Association Journal* 2016;**188**(8):567-74. [DOI: [10.1503/cmaj.150835](https://doi.org/10.1503/cmaj.150835)]

Suess 2011

Suess T, Remschmidt C, Schink S, Luchtenberg M, Haas W, Krause G, et al. Facemasks and intensified hand hygiene in a German household trial during the 2009/2010 influenza A(H1N1) pandemic: adherence and tolerability in children and adults. *Epidemiology and Infection* 2011;**139**(12):1895-901.

Tan 2004

Tan KT, Greaves MW. N95 acne. *International Journal of Dermatology* 2004;**43**(7):522-3.

Thomas 2010

Thomas RE, Jefferson T, Lasserson TJ. Influenza vaccination for healthcare workers who work with the elderly. *Cochrane Database of Systematic Reviews* 2010, Issue 2. Art. No: CD005187. [DOI: [10.1002/14651858.CD005187.pub3](https://doi.org/10.1002/14651858.CD005187.pub3)]

Warren-Gash 2013

Warren-Gash C, Fragaszy E, Hayward AC. Hand hygiene to reduce community transmission of influenza and acute respiratory tract infection: a systematic review. *Influenza and Other Respiratory Viruses* 2013;**7**(5):738-49.

WHO 2006a

World Health Organization (WHO). WHO Guidelines on Hand Hygiene in Health Care (advanced draft). Geneva (Switzerland): WHO, 2006.

WHO 2006b

World Health Organization Writing Group. Non pharmaceutical interventions for pandemic influenza, national and community measures. *Emerging Infectious Diseases* 2006;**12**(1):88-94.

WHO 2014

World Health Organization. Infection prevention and control of epidemic- and pandemic-prone acute respiratory infections in health care. apps.who.int/iris/handle/10665/112656 (accessed prior to 2 November 2020).

WHO 2017

World Health Organization. The global impact of respiratory disease. www.who.int/gard/publications/The_Global_Impact_of_Respiratory_Disease.pdf (accessed 15 March 2020).

WHO 2020a

World Health Organization. Coronavirus disease (COVID-19) advice for the public. www.who.int/emergencies/diseases/novel-coronavirus-2019/advice-for-public (accessed 20 March 2020).

WHO 2020b

World Health Organization. Getting your workplace ready for COVID-19. www.who.int/docs/default-source/coronaviruse/getting-workplace-ready-for-covid-19.pdf?sfvrsn=359a81e7_6 (accessed 20 March 2020).

WHO 2020c

World Health Organization. Coronavirus disease (COVID-19) technical guidance: infection prevention and control/WASH. www.who.int/emergencies/diseases/novel-coronavirus-2019/technical-guidance/infection-prevention-and-control (accessed 20 March 2020).

Willmott 2016

Willmott M, Nicholson A, Busse H, MacArthur GJ, Brookes S, Campbell R. Effectiveness of hand hygiene interventions in reducing illness absence among children in educational settings: a systematic review and meta-analysis. *Archives of Disease in Childhood* 2016;**101**(1):42-50.

Wong 2014

Wong VWY, Cowling BJ, Aiello AE. Hand hygiene and risk of influenza virus infections in the community: a systematic review and meta-analysis. *Epidemiology and Infection* 2014;**142**(5):922-32.

Wu 2020

Wu S, Wang Y, Jin X, Tian J, Liu J, Mao Y. Environmental contamination by SARS-CoV-2 in a designated hospital for coronavirus disease 2019. *American Journal of Infection Control* 2020;**48**(8):910-4. [DOI: [10.1016/j.ajic.2020.05.003](https://doi.org/10.1016/j.ajic.2020.05.003)]

Xiao 2020

Xiao J, Shiu EY, Gao H, Wong JY, Fong MW, Ryu S, et al. Nonpharmaceutical measures for pandemic influenza in nonhealthcare settings - personal protective and environmental measures. *Emerging Infectious Diseases* 2020;**26**(5):967-75. [DOI: [10.3201/eid2605.190994](https://doi.org/10.3201/eid2605.190994)]

Zhu 2014

Zhu JH, Lee SJ, Wang DY, Lee HP. Effects of long duration wearing of N 95 respirator and surgical facemask: a pilot study. *Journal of Lung, Pulmonary and Respiratory Research* 2014;**1**(4):97-100.

Zivich 2018

Zivich PN, Gancz AS, Aiello AE. Effect of hand hygiene on infectious diseases in the office workplace: a systematic review. *American Journal of Infection Control* 2018;**46**(4):448-55.

Zomer 2013a

Zomer TP, Erasmus V, Van Beeck EF, Tjon-A-Tsien A, Richardus JH, Voeten HA. Hand hygiene compliance and environmental determinants in child day care centers: an observational study. *American Journal of Infection Control* 2013;**41**(6):497-502.

Zomer 2013b

Zomer TP, Erasmus V, Van Empelen P, Looman C, Van Beeck EF, Tjon-A-Tsien A, et al. Sociocognitive determinants of observed and self-reported compliance to hand hygiene guidelines in child day care centers. *American Journal of Infection Control* 2013;**41**(10):862-7.

References to other published versions of this review
Jefferson 2006a

Jefferson T, Foxlee R, Del Mar C, Dooley L, Ferroni E, Hewak B, et al. Interventions for the interruption or reduction of the spread of respiratory viruses. *Cochrane Database of Systematic Reviews* 2006, Issue 6. Art. No: CD006207. [DOI: [10.1002/14651858.CD006207](https://doi.org/10.1002/14651858.CD006207)]

Jefferson 2007

Jefferson T, Foxlee R, Del Mar C, Dooley L, Ferroni E, Hewak B, et al. Interventions for the interruption or reduction of the spread of respiratory viruses. *Cochrane Database of Systematic Reviews* 2007, Issue 4. Art. No: CD006207. [DOI: [10.1002/14651858.CD006207.pub2](https://doi.org/10.1002/14651858.CD006207.pub2)]

Jefferson 2009

Jefferson T, Del Mar C, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, et al. Physical interventions to interrupt or reduce the spread of respiratory viruses: systematic review. *BMJ* 2009;**339**:b.3657. [DOI: [10.1136/bmj.b367](https://doi.org/10.1136/bmj.b367)]

Jefferson 2010

Jefferson T, Del Mar C, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, et al. Physical interventions to interrupt or reduce the spread of respiratory viruses. *Cochrane Database of Systematic Reviews* 2010, Issue 1. Art. No: CD006207. [DOI: [10.1002/14651858.CD006207.pub3](https://doi.org/10.1002/14651858.CD006207.pub3)]

Jefferson 2011

Jefferson T, Del Mar C, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, et al. Physical interventions to interrupt or reduce the spread of respiratory viruses. *Cochrane Database of Systematic Reviews* 2011, Issue 7. Art. No: CD006207. [DOI: [10.1002/14651858.CD006207.pub4](https://doi.org/10.1002/14651858.CD006207.pub4)]

* Indicates the major publication for the study

CHARACTERISTICS OF STUDIES
Characteristics of included studies [ordered by study ID]

Aelami 2015
Study characteristics

Methods	A prospective cross-sectional study conducted during the Hajj season 2012. Pilgrims were randomised into 2 groups. The intervention group received education on personal hygiene including a hygienic package containing alcohol-based hand rub (gel or spray), surgical masks, soap, paper handkerchiefs, and user instructions; the control group did not receive any intervention. ILI was defined as the presence of at least 2 of the following during their stay: fever, cough, and sore throat. Questionnaires including demographic and clinical information were distributed amongst trained physicians before departure from Iran.
Participants	Total enrolled: 664 Iranian pilgrims (306 in the intervention group and 358 in the control group) Inclusion criteria: not reported Exclusion criteria: not reported

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Aelami 2015 (Continued)

Interventions	Hygiene education and package. See Table 1 for details.
Outcomes	ILI defined as the presence of at least 2 of the following during their stay: fever, cough, and sore throat. No safety outcomes were reported.
Notes	This is an abstract, therefore few details were reported.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient details provided
Allocation concealment (selection bias)	Unclear risk	Insufficient details provided
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Insufficient details provided
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient details provided
Selective reporting (reporting bias)	Unclear risk	Insufficient details provided

Aiello 2010
Study characteristics

Methods	<p>Cluster-RCT assessing the effects of hand sanitiser and masks versus masks or no intervention on ILI symptoms. The trial was conducted in university halls of residence with more than 100 student residents in a US university during the 2006 to 2007 influenza "season". The study lasted 6 weeks.</p> <p>The units of randomisation were 7 of the 15 halls. 1 hall was very large (1240 residents), and the 6 remaining ones, which had between 110 and 830 residents, were combined into 2 clusters roughly equivalent in size. The 3 clusters were then randomised by random extraction of the clustered halls' names out of a container. The largest hall (single-cluster) was randomised to the mask and hand sanitiser arm; the 4-halls cluster received masks; and the remaining 2 halls were assigned as controls.</p>
Participants	<p>A total of 1297 with completed baseline survey and at least 1 weekly survey result were analysed (face mask and hand hygiene group = 367; face mask-only group = 378; control group = 552).</p> <p>Inclusion criteria: aged 18 or more, willing to wear mask and use alcohol-based hand sanitiser, have a throat swab specimen collected when ill, and complete the baseline and weekly surveys over the 6-week study period</p> <p>Exclusion criteria: individuals reporting a skin allergy to alcohol were excluded</p>

Aiello 2010 (Continued)

Recruitment of students began in 26 November, but the trial did not go “live” with distribution of intervention materials until 22 January 2007 when the first case of influenza was confirmed on campus by laboratory tests. Enrolment continued until 16 February 2007, and the study was completed on 16 March 2007. During the study period there was a 1-week break when the majority of residents left campus. There were 1327 eligible participants, 1297 of which had a complete baseline survey and at least 1-weekly survey result. It is unclear what the ineligibility criteria were for the 30 missing (1327 minus 1297), but the explanation may be in the appendix.

Interventions

Alcohol-based hand sanitiser (62% ethyl alcohol in a gel base) in a squeeze bottle and TECNOL procedure masks with ear loops (KC Ltd) and educational material or masks and educational material or no intervention. Compliance was encouraged within halls and outside. Sleep wearing was optional.

All participants received basic video-linked instruction on cough etiquette and hand sanitation. At baseline and weekly during the study, participants were asked to fill in a web-based survey collecting demographic and ILI symptom data. This was supplemented by direct observation of compliance by staff.

Compliance with “optimal handwashing” (at least 20 seconds 5 or more times a day) was significantly higher in the sanitiser-and-mask arm.

Outcomes

Laboratory details are described in appendix.

Effectiveness: ILI, defined as cough and at least 1 constitutional symptom (fever/feverishness, chills, headache, myalgia). ILI cases were given contact nurses' phone numbers to record the illness and paid USD 25 to provide a throat swab. 368 participants had ILI, and 94 of these had a throat swab analysed by PCR. 10 of these were positive for influenza (7 for A and 3 for B).

Safety: N/A

Notes

The authors conclude that “These findings suggest that face masks and hand hygiene may reduce respiratory illnesses in shared living settings and mitigate the impact of the influenza A (H1N1) pandemic”. This conclusion is based on a significantly lower level of ILI incidence in the mask and hand sanitiser arm compared to the other 2 arms after adjustment for covariates (30% to 50% less in arm 1 compared to controls in the last 2 weeks of the study).

Comparison with the ILI rate of the control arm may not be a reflection of the underlying rate of ILI because the intervention arm received instruction on hand sanitation and hand etiquette.

The play of adjustments is unclear. The intracluster correlation coefficient is reported in the footer of Table 4. Its very small size suggests lack of clustering within halls.

The role of spring break is mentioned in the Discussion, as are the results of this study compared to other studies included in our review (Cowling 2008 and MacIntyre 2009).

The authors report that 147 of 1297 participants (11.3%) had ILI symptoms “at baseline” and were excluded from analysis. During the 6 weeks of the study, 368 of 1150 participants (32%) had ILI. This averages out at about 5% per week. It is unclear what the term “at baseline” means; presumably this means during the 2 to 3 weeks of participant enrolment. If this is so, the reason for the triggering of the interventions (tied to influenza isolation) are obscure, as the trial is supposedly about ILI, and an ILI outbreak was already under way “at baseline”.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Described as randomised, but sequence generation not reported
Allocation concealment (selection bias)	High risk	The residence hall units were randomised by blindly selecting a uniform ticket with the name of each hall out of a container (A.S.M. and A.A.) for randomisation assignment to each study arm.

Aiello 2010 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	<p>Attrition is reported as follows: 9, 11, and 19 ineligible and 26, 52, and 21 lost to follow-up (respectively by arm), for a total of 39 and 99 for each reason for attrition. In total, 1297 (97%) of 1331 participants completed a baseline and at least 1-weekly survey.</p> <p>The text reports an ITT analysis with only 1 ILI episode included by participant.</p> <p>No reasons for the attrition of participants and swab volunteers are reported (were the swabs taken from a random sample or not?).</p>
Selective reporting (reporting bias)	High risk	<p>There is no information on the causes of ILI other than the reporting on the 10 influenza PCR-positive swabs of 94 out of 368 students with ILI. This is a very low rate (and the Discussion confirms that the influenza season was mild), but investigation of the other known causes of ILI is not even mentioned in the text. This is especially important because stress, alcohol intake levels, and influenza vaccination were a significant predictor of ILI symptoms (Table 1). The reason for selective testing and/or reporting of influenza viruses tests over the other causes of ILI are unclear, especially as the study objective was focused on ILI. The text is also difficult to follow, weaving the reporting of ILI and influenza without a clear rationale.</p>

Aiello 2012
Study characteristics

Methods	<p>During the 2007 to 2008 influenza season, 1111 students residing in university residence halls were cluster-randomised by residence house (N = 37) to either face mask and hand hygiene, face mask only, or control arms. Discrete time survival analysis using generalised models estimated rate ratios according to study arm, each week and cumulatively over the 6-week intervention period, for clinically verified ILI and laboratory-confirmed influenza A or B.</p>
Participants	<p>A total of 1187 young adults living in 37 residence halls, randomly assigned to 1 of 3 groups for 6 weeks: face mask use (n = 392), face masks with hand hygiene (n = 349), control (n = 370)</p> <p>Inclusion criteria: aged 18 or more, willing to wear mask and use alcohol-based hand sanitiser, have a throat swab specimen collected when ill, and complete the baseline and weekly surveys over the 6-week study period</p> <p>Exclusion criteria: individuals reporting a skin allergy to alcohol were excluded</p>
Interventions	<p>Participants were assigned to face mask and hand hygiene, face mask only, or control group during the study. See Table 1 for details.</p>
Outcomes	<p>Clinically verified ILI: case definition (presence of cough and at least 1 or more of fever/feverishness, chills, or body aches)</p> <p>Laboratory-confirmed influenza A or B. Throat swab specimens were tested for influenza A or B using RT-PCR.</p>

Aiello 2012 (Continued)

No safety outcomes reported.

Notes This study has the same trial registration number as the [Aiello 2010](#) study; the study was funded by government and pharmaceutical industry.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer generation of sequence described.
Allocation concealment (selection bias)	Low risk	All residence houses in each of the residence halls were randomised prior to the intervention implementation.
Blinding of participants and personnel (performance bias) All outcomes	High risk	No blinding for study participants and personnel
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition low and similar in each group
Selective reporting (reporting bias)	Low risk	2 outcomes specified and reported.

Aiello 2010
Study characteristics

Methods	<p>Cluster-RCT assessing the effects of hand sanitiser and masks versus masks or no intervention on ILI symptoms. The trial was conducted in university halls of residence with more than 100 student residents in a US university during the 2006 to 2007 influenza "season". The study lasted 6 weeks.</p> <p>The units of randomisation were 7 of the 15 halls. 1 hall was very large (1240 residents), and the 6 remaining ones, which had between 110 and 830 residents, were combined into 2 clusters roughly equivalent in size. The 3 clusters were then randomised by random extraction of the clustered halls' names out of a container. The largest hall (single-cluster) was randomised to the mask and hand sanitiser arm; the 4-halls cluster received masks; and the remaining 2 halls were assigned as controls.</p>
Participants	<p>A total of 1297 with completed baseline survey and at least 1 weekly survey result were analysed (face mask and hand hygiene group = 367; face mask-only group = 378; control group = 552).</p> <p>Inclusion criteria: aged 18 or more, willing to wear mask and use alcohol-based hand sanitiser, have a throat swab specimen collected when ill, and complete the baseline and weekly surveys over the 6-week study period</p> <p>Exclusion criteria: individuals reporting a skin allergy to alcohol were excluded</p> <p>Recruitment of students began in 26 November, but the trial did not go "live" with distribution of intervention materials until 22 January 2007 when the first case of influenza was confirmed on campus by laboratory tests. Enrolment continued until 16 February 2007, and the study was completed on 16 March 2007. During the study period there was a 1-week break when the majority of residents left cam-</p>

Aiello 2010 (Continued)

pus. There were 1327 eligible participants, 1297 of which had a complete baseline survey and at least 1-weekly survey result. It is unclear what the ineligibility criteria were for the 30 missing (1327 minus 1297), but the explanation may be in the appendix.

Interventions	<p>Alcohol-based hand sanitiser (62% ethyl alcohol in a gel base) in a squeeze bottle and TECNOL procedure masks with ear loops (KC Ltd) and educational material or masks and educational material or no intervention. Compliance was encouraged within halls and outside. Sleep wearing was optional.</p> <p>All participants received basic video-linked instruction on cough etiquette and hand sanitation. At baseline and weekly during the study, participants were asked to fill in a web-based survey collecting demographic and ILI symptom data. This was supplemented by direct observation of compliance by staff.</p> <p>Compliance with “optimal handwashing” (at least 20 seconds 5 or more times a day) was significantly higher in the sanitiser-and-mask arm.</p>
Outcomes	<p>Laboratory details are described in appendix.</p> <p>Effectiveness: ILI, defined as cough and at least 1 constitutional symptom (fever/feverishness, chills, headache, myalgia). ILI cases were given contact nurses' phone numbers to record the illness and paid USD 25 to provide a throat swab. 368 participants had ILI, and 94 of these had a throat swab analysed by PCR. 10 of these were positive for influenza (7 for A and 3 for B).</p> <p>Safety: N/A</p>
Notes	<p>The authors conclude that “These findings suggest that face masks and hand hygiene may reduce respiratory illnesses in shared living settings and mitigate the impact of the influenza A (H1N1) pandemic”. This conclusion is based on a significantly lower level of ILI incidence in the mask and hand sanitiser arm compared to the other 2 arms after adjustment for covariates (30% to 50% less in arm 1 compared to controls in the last 2 weeks of the study).</p> <p>Comparison with the ILI rate of the control arm may not be a reflection of the underlying rate of ILI because the intervention arm received instruction on hand sanitation and hand etiquette.</p> <p>The play of adjustments is unclear. The intracluster correlation coefficient is reported in the footer of Table 4. Its very small size suggests lack of clustering within halls.</p> <p>The role of spring break is mentioned in the Discussion, as are the results of this study compared to other studies included in our review (Cowling 2008 and MacIntyre 2009).</p> <p>The authors report that 147 of 1297 participants (11.3%) had ILI symptoms “at baseline” and were excluded from analysis. During the 6 weeks of the study, 368 of 1150 participants (32%) had ILI. This averages out at about 5% per week. It is unclear what the term “at baseline” means; presumably this means during the 2 to 3 weeks of participant enrolment. If this is so, the reason for the triggering of the interventions (tied to influenza isolation) are obscure, as the trial is supposedly about ILI, and an ILI outbreak was already under way “at baseline”.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Described as randomised, but sequence generation not reported
Allocation concealment (selection bias)	High risk	The residence hall units were randomised by blindly selecting a uniform ticket with the name of each hall out of a container (A.S.M. and A.A.) for randomisation assignment to each study arm.
Blinding of participants and personnel (performance bias)	High risk	Unblinded study

Aiello 2010 (Continued)

All outcomes

Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	<p>Attrition is reported as follows: 9, 11, and 19 ineligible and 26, 52, and 21 lost to follow-up (respectively by arm), for a total of 39 and 99 for each reason for attrition. In total, 1297 (97%) of 1331 participants completed a baseline and at least 1-weekly survey.</p> <p>The text reports an ITT analysis with only 1 ILI episode included by participant.</p> <p>No reasons for the attrition of participants and swab volunteers are reported (were the swabs taken from a random sample or not?).</p>
Selective reporting (reporting bias)	High risk	<p>There is no information on the causes of ILI other than the reporting on the 10 influenza PCR-positive swabs of 94 out of 368 students with ILI. This is a very low rate (and the Discussion confirms that the influenza season was mild), but investigation of the other known causes of ILI is not even mentioned in the text. This is especially important because stress, alcohol intake levels, and influenza vaccination were a significant predictor of ILI symptoms (Table 1). The reason for selective testing and/or reporting of influenza viruses tests over the other causes of ILI are unclear, especially as the study objective was focused on ILI. The text is also difficult to follow, weaving the reporting of ILI and influenza without a clear rationale.</p>

Aiello 2012
Study characteristics

Methods	<p>During the 2007 to 2008 influenza season, 1111 students residing in university residence halls were cluster-randomised by residence house (N = 37) to either face mask and hand hygiene, face mask only, or control arms. Discrete time survival analysis using generalised models estimated rate ratios according to study arm, each week and cumulatively over the 6-week intervention period, for clinically verified ILI and laboratory-confirmed influenza A or B.</p>
Participants	<p>A total of 1187 young adults living in 37 residence halls, randomly assigned to 1 of 3 groups for 6 weeks: face mask use (n = 392), face masks with hand hygiene (n = 349), control (n = 370)</p> <p>Inclusion criteria: aged 18 or more, willing to wear mask and use alcohol-based hand sanitiser, have a throat swab specimen collected when ill, and complete the baseline and weekly surveys over the 6-week study period</p> <p>Exclusion criteria: individuals reporting a skin allergy to alcohol were excluded</p>
Interventions	<p>Participants were assigned to face mask and hand hygiene, face mask only, or control group during the study. See Table 1 for details.</p>
Outcomes	<p>Clinically verified ILI: case definition (presence of cough and at least 1 or more of fever/feverishness, chills, or body aches)</p> <p>Laboratory-confirmed influenza A or B. Throat swab specimens were tested for influenza A or B using RT-PCR.</p> <p>No safety outcomes reported.</p>

Aiello 2012 (Continued)

Notes This study has the same trial registration number as the [Aiello 2010](#) study; the study was funded by government and pharmaceutical industry.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer generation of sequence described.
Allocation concealment (selection bias)	Low risk	All residence houses in each of the residence halls were randomised prior to the intervention implementation.
Blinding of participants and personnel (performance bias) All outcomes	High risk	No blinding for study participants and personnel
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition low and similar in each group
Selective reporting (reporting bias)	Low risk	2 outcomes specified and reported.

Alzahr 2018
Study characteristics

Methods	A cluster-RCT conducted amongst girls attending 4 primary schools between January and March 2018. The participants attended a hand hygiene workshop. The schoolgirls' absences were followed up for 5 weeks. Incidence rate, percentage of absence days, and absence rate were calculated for total and upper respiratory infections absences.
Participants	A total of 496 schoolgirls aged of 6 to 12 years, attending 4 public primary girls' schools in the city of Riyadh, Saudi Arabia between January and March 2018. Students were randomised to education group (n = 234) or control group (n = 262). Exclusion criteria: not reported
Interventions	Hand hygiene workshop. See Table 1 for details.
Outcomes	Incidence rate, percentage of absence days, and absence rate were calculated for total and upper respiratory infections absences. The episode of URIs was defined as having 2 of the following symptoms for a day or 1 of the symptoms for 2 or more consecutive days: 1) a runny nose, 2) a stuffy or blocked nose or noisy breathing, 3) sneezing, 4) a cough, 5) a sore throat, and 6) feeling hot, having a fever or a chill. No safety outcomes reported.
Notes	Source of funding unclear

Alzahr 2018 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient detail provided.
Allocation concealment (selection bias)	Low risk	Schools allocated prior to all schoolgirls attending selected schools were invited to participate.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Unclear risk	No protocol available

Arbogast 2016
Study characteristics

Methods	A 13.5-month prospective cluster-RCT executed with alcohol-based hand sanitiser in strategic workplace locations and personal use (intervention group) and brief hand hygiene education (both groups). 4 years of retrospective data were collected for all participants.
Participants	Data for a total of 1183 participants were analysed (intervention group = 525, control group = 607). Inclusion criteria: all employees at 3 facilities who were 18 years of age or older, were enrolled in the company health insurance coverage, did not transfer between sites, and worked onsite full time (≥ 32 hours) were eligible for the study Exclusion criteria: not reported
Interventions	Alcohol-based hand sanitiser in strategic workplace locations and personal use (intervention group) and brief hand hygiene education (both groups). See Table 1 for details.
Outcomes	(1) The number of healthcare insurance claims, for a defined set of preventable illnesses, per participant per year, and (2) absenteeism, defined as the number of sick episodes per participant per year Claims based on ICD-9 codes No safety outcomes reported.
Notes	Industry funded; only 2 clusters (1 per group) included, hence study data not included in meta-analysis.

Risk of bias

Arbogast 2016 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No details provided.
Allocation concealment (selection bias)	Unclear risk	No details provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal and similar in 2 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Arbogast 2016
Study characteristics

Methods	A 13.5-month prospective cluster-RCT executed with alcohol-based hand sanitiser in strategic workplace locations and personal use (intervention group) and brief hand hygiene education (both groups). 4 years of retrospective data were collected for all participants.
Participants	Data for a total of 1183 participants were analysed (intervention group = 525, control group = 607). Inclusion criteria: all employees at 3 facilities who were 18 years of age or older, were enrolled in the company health insurance coverage, did not transfer between sites, and worked onsite full time (≥ 32 hours) were eligible for the study Exclusion criteria: not reported
Interventions	Alcohol-based hand sanitiser in strategic workplace locations and personal use (intervention group) and brief hand hygiene education (both groups). See Table 1 for details.
Outcomes	(1) The number of healthcare insurance claims, for a defined set of preventable illnesses, per participant per year, and (2) absenteeism, defined as the number of sick episodes per participant per year Claims based on ICD-9 codes No safety outcomes reported.
Notes	Industry funded; only 2 clusters (1 per group) included, hence study data not included in meta-analysis.

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Arbogast 2016 (Continued)

Random sequence generation (selection bias)	Unclear risk	No details provided.
Allocation concealment (selection bias)	Unclear risk	No details provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal and similar in 2 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Azor-Martinez 2016
Study characteristics

Methods	Randomised, controlled, and open study with an 8-month follow-up. The experimental group washed their hands with soap and water, together with using hand sanitiser, and the control group followed their usual handwashing procedures. Absenteeism rates due to URIs were compared between the 2 groups through a multivariate Poisson regression analysis. The per cent of days missed in both groups were compared with a z test.
Participants	<p>A sample of 1341 (intervention group = 621, control group = 720)</p> <p>Inclusion criteria: children 4 to 12 years old, attending 5 state schools in Almeria (Spain) whose parents/guardians had signed an informed consent document</p> <p>Exclusion criteria: children who had any of the following chronic illnesses that predisposed them to infection: neoplasia, primary and secondary immunodeficiencies, cystic fibrosis, chronic treatment with high doses of steroids or immunosuppressants</p>
Interventions	Hand-washing workshops of 2-hour duration. The experimental group washed their hands with soap and water together with using hand sanitiser, whilst the control group followed usual hand-washing procedures. See Table 1 for details.
Outcomes	<p>Absenteeism rates due to URIs</p> <p>Per cent of days missed</p> <p>Respiratory illness was defined by 2 of the following symptoms during 1 day, or 1 of the symptoms for 2 consecutive days: (1) runny nose; (2) stuffy or blocked nose or noisy breathing; (3) cough; (4) feeling hot or feverish or having chills; (5) sore throat; or (6) sneezing.</p> <p>A school absenteeism case (episode) was defined as when a child failed to attend school due to an URI. Common infectious illnesses, such as conjunctivitis, and skin infections were not included. Other causes for absenteeism, such as doctors' appointments, family vacations, and accident injuries, were also excluded.</p>

Azor-Martinez 2016 (Continued)

No safety outcomes reported.

Notes Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	A random number table was used.
Allocation concealment (selection bias)	Low risk	Schools/classes allocated prior to children recruited.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition levels high and different in the 2 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Azor-Martinez 2016
Study characteristics

Methods	Randomised, controlled, and open study with an 8-month follow-up. The experimental group washed their hands with soap and water, together with using hand sanitiser, and the control group followed their usual handwashing procedures. Absenteeism rates due to URIs were compared between the 2 groups through a multivariate Poisson regression analysis. The per cent of days missed in both groups were compared with a z test.
Participants	A sample of 1341 (intervention group = 621, control group = 720) Inclusion criteria: children 4 to 12 years old, attending 5 state schools in Almeria (Spain) whose parents/guardians had signed an informed consent document Exclusion criteria: children who had any of the following chronic illnesses that predisposed them to infection: neoplasia, primary and secondary immunodeficiencies, cystic fibrosis, chronic treatment with high doses of steroids or immunosuppressants
Interventions	Hand-washing workshops of 2-hour duration. The experimental group washed their hands with soap and water together with using hand sanitiser, whilst the control group followed usual hand-washing procedures. See Table 1 for details.
Outcomes	Absenteeism rates due to URIs Per cent of days missed

Azor-Martinez 2016 (Continued)

Respiratory illness was defined by 2 of the following symptoms during 1 day, or 1 of the symptoms for 2 consecutive days: (1) runny nose; (2) stuffy or blocked nose or noisy breathing; (3) cough; (4) feeling hot or feverish or having chills; (5) sore throat; or (6) sneezing.

A school absenteeism case (episode) was defined as when a child failed to attend school due to an URI. Common infectious illnesses, such as conjunctivitis, and skin infections were not included. Other causes for absenteeism, such as doctors' appointments, family vacations, and accident injuries, were also excluded.

No safety outcomes reported.

Notes	Government funded	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	A random number table was used.
Allocation concealment (selection bias)	Low risk	Schools/classes allocated prior to children recruited.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition levels high and different in the 2 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Azor-Martinez 2018
Study characteristics

Methods	A cluster-RCT, controlled, and open study of 911 children aged 0 to 3 years attending 24 DCCs in Almería, Spain, with an 8-month follow-up. 2 intervention groups of DCC families performed educational and hand hygiene measures, 1 with soap and water (n = 274), another with hand sanitiser (n = 339), and the control group followed usual hand-washing procedures (n = 298). Respiratory infection (RI) episode rates were compared through multilevel Poisson regression models. The percentage of days missed were compared with Poisson exact tests.
Participants	A total of 911 children attending 24 DCCs in Almería (Spain). Inclusion criteria: children between 0 and 3 years old enrolled in DCCs and attending for at least 15 hours per week whose parents or guardians had signed an informed consent Exclusion criteria: children with chronic illness or medication that could affect their likelihood of contracting an infection

Azor-Martinez 2018 (Continued)

Data were analysed for 911 participants: hand sanitiser group (n = 339), soap and water group (n = 274), and control group (n = 298).

Interventions	2 intervention groups. 1 group used soap and water, another used hand sanitiser, whilst the control group followed usual hand-washing procedures. Groups received 1-hour hand hygiene workshop. See Table 1 for details.
Outcomes	<p>Primary: RI incidence rate</p> <p>Secondary: (1) the presence or absence of at least 1 antibiotic prescription for each new RI episode during the study period (topical antibiotics were excluded), and (2) the percentage of RI absenteeism days in the 3 groups calculated as the ratio of RI absenteeism days to all possible days of attendance</p> <p>DCC absenteeism episode was defined as when a child failed to attend a DCC because of an RI.</p> <p>Respiratory illness was defined as the presence of 2 of the following symptoms during 1 day or the presence of 1 of the symptoms for 2 consecutive days: (1) runny nose, (2) stuffy or blocked nose or noisy breathing, (3) cough, (4) feeling hot or feverish or having chills, (5) sore throat, or (6) sneezing.</p> <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer randomisation using statistical software for the sequence
Allocation concealment (selection bias)	Low risk	Clusters assigned prior to recruitment.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal and similar in 3 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Azor-Martinez 2018
Study characteristics

Methods	A cluster-RCT, controlled, and open study of 911 children aged 0 to 3 years attending 24 DCCs in Almería, Spain, with an 8-month follow-up. 2 intervention groups of DCC families performed educational and hand hygiene measures, 1 with soap and water (n = 274), another with hand sanitiser (n = 339), and the control group followed usual hand-washing procedures (n = 298). Respiratory infection (RI)
---------	---

Azor-Martinez 2018 (Continued)

episode rates were compared through multilevel Poisson regression models. The percentage of days missed were compared with Poisson exact tests.

Participants	<p>A total of 911 children attending 24 DCCs in Almería (Spain).</p> <p>Inclusion criteria: children between 0 and 3 years old enrolled in DCCs and attending for at least 15 hours per week whose parents or guardians had signed an informed consent</p> <p>Exclusion criteria: children with chronic illness or medication that could affect their likelihood of contracting an infection</p> <p>Data were analysed for 911 participants: hand sanitiser group (n = 339), soap and water group (n = 274), and control group (n = 298).</p>
Interventions	<p>2 intervention groups. 1 group used soap and water, another used hand sanitiser, whilst the control group followed usual hand-washing procedures. Groups received 1-hour hand hygiene workshop. See Table 1 for details.</p>
Outcomes	<p>Primary: RI incidence rate</p> <p>Secondary: (1) the presence or absence of at least 1 antibiotic prescription for each new RI episode during the study period (topical antibiotics were excluded), and (2) the percentage of RI absenteeism days in the 3 groups calculated as the ratio of RI absenteeism days to all possible days of attendance</p> <p>DCC absenteeism episode was defined as when a child failed to attend a DCC because of an RI.</p> <p>Respiratory illness was defined as the presence of 2 of the following symptoms during 1 day or the presence of 1 of the symptoms for 2 consecutive days: (1) runny nose, (2) stuffy or blocked nose or noisy breathing, (3) cough, (4) feeling hot or feverish or having chills, (5) sore throat, or (6) sneezing.</p> <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer randomisation using statistical software for the sequence
Allocation concealment (selection bias)	Low risk	Clusters assigned prior to recruitment.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal and similar in 3 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Ban 2015
Study characteristics

Methods	"Group randomised" trial. Only 2 clusters, which were 2 kindergartens in Xiantao City, Hubei Province, China
Participants	Data for a total of 393 participants were analysed (intervention group = 194, control group = 199). 5 classes (221 children) randomly selected from 1 kindergarten in the intervention group and 6 classes (244 children) randomly selected from another kindergarten in the control group. Children were aged 5 or under. There were 72 exclusions from the analysis.
Interventions	Intervention group: hand hygiene and surface-cleaning education and provision of products for kindergarten and home use. Control group: usual practice. See Table 1 for details.
Outcomes	Respiratory illness, defined as: 2 or more of the following: fever, cough and expectoration, runny nose and nasal congestion, collected by parental questionnaire. Axillary temperature higher than 37.3 °C or the range of temperature fluctuation is more than 1 °C. 'Cough and expectoration' were defined as 3 or more coughs in a single hour and lasting for 4 or more hours in a single day, with or without expectoration. 'Runny nose and nasal congestion' were defined as a runny nose lasting for 4 or more hours in 1 day, with or without nasal congestion.

Notes

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Method not described, and only 2 clusters.
Allocation concealment (selection bias)	Unclear risk	Method not described.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Parental report, and parents were aware of treatment allocation
Selective reporting (reporting bias)	High risk	Attrition reported and balanced between groups, but high rate of attrition in a trial with small numbers of participants.

Ban 2015
Study characteristics

Ban 2015 (Continued)

Methods	"Group randomised" trial. Only 2 clusters, which were 2 kindergartens in Xiantao City, Hubei Province, China
Participants	Data for a total of 393 participants were analysed (intervention group = 194, control group = 199). 5 classes (221 children) randomly selected from 1 kindergarten in the intervention group and 6 classes (244 children) randomly selected from another kindergarten in the control group. Children were aged 5 or under. There were 72 exclusions from the analysis.
Interventions	Intervention group: hand hygiene and surface-cleaning education and provision of products for kindergarten and home use. Control group: usual practice. See Table 1 for details.
Outcomes	Respiratory illness, defined as: 2 or more of the following: fever, cough and expectoration, runny nose and nasal congestion, collected by parental questionnaire. Axillary temperature higher than 37.3 °C or the range of temperature fluctuation is more than 1 °C. 'Cough and expectoration' were defined as 3 or more coughs in a single hour and lasting for 4 or more hours in a single day, with or without expectoration. 'Runny nose and nasal congestion' were defined as a runny nose lasting for 4 or more hours in 1 day, with or without nasal congestion.

Notes

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Method not described, and only 2 clusters.
Allocation concealment (selection bias)	Unclear risk	Method not described.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Parental report, and parents were aware of treatment allocation
Selective reporting (reporting bias)	High risk	Attrition reported and balanced between groups, but high rate of attrition in a trial with small numbers of participants.

Barasheed 2014
Study characteristics

Methods	Pilot, non-blinded, parallel, cluster-RCT
Participants	22 tents were randomly selected from the Australian pilgrims camped in Mina, during Hajj in 2011; 12 tents were allocated to the mask group and 10 tents to the control group. A total of 164 Australian pil-

Barasheed 2014 (Continued)

grims were recruited: 75 in the mask group (39 'cases' and 36 'contacts') and 89 in the control group (36 'cases' and 53 'contacts').

Inclusion criteria for index case: 1) Australian pilgrims of any gender aged > 15 years who attend the Hajj 2011, and 2) have symptoms of respiratory infection for 3 days. For close tent contact: 1) Australian pilgrims of any gender aged 15 years or more who attend the Hajj 2011, and 2) pilgrims who share the same tent and sleep "immediately close" to the index case.

Exclusion criteria: for index case: 1) pilgrims who do not suffer from symptoms of respiratory infection, 2) pilgrims who present with symptoms of respiratory infection for > 3 days, and 3) children aged less than 15 years. For close tent contact: 1) pilgrims who are symptomatic at presentation, 2) pilgrims who are not close tent contacts of an index case, and 3) children aged less than 15 years. Only 10% to 15% of potential participants took part in the study.

Interventions	"supervised mask use" versus "no supervised mask use". See Table 1 for details.
Outcomes	<p>Laboratory: 2 nasal swabs from all ILI cases and contacts, 1 for influenza POCT using the QuickVue Influenza (A+B) assay (Quidel Corporation, San Diego, USA) and 1 for later nucleic acid testing for influenza and other respiratory viruses. However, there was a problem with getting POCT on time during Hajj.</p> <p>Effectiveness: to assess the effectiveness of face masks in the prevention of transmission of ILI. ILI was defined as subjective (or proven) fever plus 1 respiratory symptom (e.g. dry or productive cough, runny nose, sore throat, shortness of breath).</p> <p>Safety: none planned or reported</p>
Notes	<p>The study was conducted from 4 November 2011 to 10 November 2011.</p> <p>Funding: government (Qatar National Research Fund (QNRF))</p> <p>Compliance with face mask use by pilgrims was 56 of 75 (76%) in the mask group and 11 of 89 (12%) in the control group ($P < 0.001$). The proportion of face mask user in the 'mask' tents was 76% for both males (19/25) and females (38/50). The most often reported reason for not wearing face masks was discomfort (15%).</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information provided.
Allocation concealment (selection bias)	Unclear risk	"tents were randomised to either intervention group (supervised mask tent) or control group (no supervised mask tent) by an independent study coordinator who was not an investigator", but did not mention how
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Because advice from the Saudi Ministry of Hajj to all pilgrims included recommending the wearing of masks, all pilgrims, both cases and controls, were asked about mask-wearing"
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Self-reported outcomes (nasal swab was performed for those who reported ILI symptoms and was not intended as systematic detection). ILI was defined as subjective (or proven) fever plus 1 respiratory symptom.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up, all numbers were reported from enrolment to analysis

Barasheed 2014 (Continued)

Selective reporting (reporting bias)	Low risk	All planned outcomes were reported.
--------------------------------------	----------	-------------------------------------

Barasheed 2014
Study characteristics

Methods	Pilot, non-blinded, parallel, cluster-RCT
Participants	<p>22 tents were randomly selected from the Australian pilgrims camped in Mina, during Hajj in 2011; 12 tents were allocated to the mask group and 10 tents to the control group. A total of 164 Australian pilgrims were recruited: 75 in the mask group (39 'cases' and 36 'contacts') and 89 in the control group (36 'cases' and 53 'contacts').</p> <p>Inclusion criteria for index case: 1) Australian pilgrims of any gender aged > 15 years who attend the Hajj 2011, and 2) have symptoms of respiratory infection for 3 days. For close tent contact: 1) Australian pilgrims of any gender aged 15 years or more who attend the Hajj 2011, and 2) pilgrims who share the same tent and sleep "immediately close" to the index case.</p> <p>Exclusion criteria: for index case: 1) pilgrims who do not suffer from symptoms of respiratory infection, 2) pilgrims who present with symptoms of respiratory infection for > 3 days, and 3) children aged less than 15 years. For close tent contact: 1) pilgrims who are symptomatic at presentation, 2) pilgrims who are not close tent contacts of an index case, and 3) children aged less than 15 years. Only 10% to 15% of potential participants took part in the study.</p>
Interventions	"supervised mask use" versus "no supervised mask use". See Table 1 for details.
Outcomes	<p>Laboratory: 2 nasal swabs from all ILI cases and contacts, 1 for influenza POCT using the QuickVue Influenza (A+B) assay (Quidel Corporation, San Diego, USA) and 1 for later nucleic acid testing for influenza and other respiratory viruses. However, there was a problem with getting POCT on time during Hajj.</p> <p>Effectiveness: to assess the effectiveness of face masks in the prevention of transmission of ILI. ILI was defined as subjective (or proven) fever plus 1 respiratory symptom (e.g. dry or productive cough, runny nose, sore throat, shortness of breath).</p> <p>Safety: none planned or reported</p>
Notes	<p>The study was conducted from 4 November 2011 to 10 November 2011.</p> <p>Funding: government (Qatar National Research Fund (QNRF))</p> <p>Compliance with face mask use by pilgrims was 56 of 75 (76%) in the mask group and 11 of 89 (12%) in the control group ($P < 0.001$). The proportion of face mask user in the 'mask' tents was 76% for both males (19/25) and females (38/50). The most often reported reason for not wearing face masks was discomfort (15%).</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information provided.
Allocation concealment (selection bias)	Unclear risk	"tents were randomised to either intervention group (supervised mask tent) or control group (no supervised mask tent) by an independent study coordinator who was not an investigator", but did not mention how

Barasheed 2014 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Because advice from the Saudi Ministry of Hajj to all pilgrims included recommending the wearing of masks, all pilgrims, both cases and controls, were asked about mask-wearing"
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Self-reported outcomes (nasal swab was performed for those who reported ILI symptoms and was not intended as systematic detection). ILI was defined as subjective (or proven) fever plus 1 respiratory symptom.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up, all numbers were reported from enrolment to analysis
Selective reporting (reporting bias)	Low risk	All planned outcomes were reported.

Biswas 2019
Study characteristics

Methods	Cluster-RCT in 24 primary schools in Dhaka to assess the effectiveness of hand sanitiser and a respiratory hygiene education intervention in reducing ILI and laboratory-confirmed influenza during June to September 2015. 12 schools were randomly selected to receive hand sanitiser and respiratory hygiene education, and 12 schools received no intervention. Field staff actively followed children daily to monitor for new ILI episodes (cough with fever) through school visits and by phone if a child was absent. When an illness episode was identified, medical technologists collected nasal swabs to test for influenza viruses.
Participants	A total of 10,855 students were enrolled in the study (intervention schools = 5077 children; control schools = 5778 children). Children aged 5 to 10 years educated in 24 randomly selected primary schools in Dhaka, Bangladesh Exclusion: schools that offered education above grade 5 because of differences in student populations, as well as schools that had previously received a hand or respiratory hygiene intervention
Interventions	Hand sanitiser and respiratory hygiene education versus no intervention. See Table 1 for details.
Outcomes	Incidence of ILI Incidence of laboratory-confirmed influenza (RT-PCR) An ILI episode was defined as measured fever ≥ 38 °C or subjective fever and cough. If a child was absent, the field staff followed up by phone to identify the reason for absenteeism and to determine if the child met the ILI case definition. If a child in a participating school had an ILI episode, a trained medical technologist visited the child's household to obtain consent from the child's parent/guardian and collect a nasal swab from the child within 48 hours of symptom onset. If it was outside the 48-hour window, the sample was not collected. No safety outcomes reported.
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Biswas 2019 (Continued)

Random sequence generation (selection bias)	Low risk	Sequence generated using a computer-based random number generator.
Allocation concealment (selection bias)	Low risk	Allocation completed prior to individuals being recruited.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Information missing for 30 children (28 children in the control schools and 2 children in the intervention schools)
Selective reporting (reporting bias)	Unclear risk	No protocol available

Biswas 2019
Study characteristics

Methods	Cluster-RCT in 24 primary schools in Dhaka to assess the effectiveness of hand sanitiser and a respiratory hygiene education intervention in reducing ILI and laboratory-confirmed influenza during June to September 2015. 12 schools were randomly selected to receive hand sanitiser and respiratory hygiene education, and 12 schools received no intervention. Field staff actively followed children daily to monitor for new ILI episodes (cough with fever) through school visits and by phone if a child was absent. When an illness episode was identified, medical technologists collected nasal swabs to test for influenza viruses.
Participants	A total of 10,855 students were enrolled in the study (intervention schools = 5077 children; control schools = 5778 children). Children aged 5 to 10 years educated in 24 randomly selected primary schools in Dhaka, Bangladesh Exclusion: schools that offered education above grade 5 because of differences in student populations, as well as schools that had previously received a hand or respiratory hygiene intervention
Interventions	Hand sanitiser and respiratory hygiene education versus no intervention. See Table 1 for details.
Outcomes	Incidence of ILI Incidence of laboratory-confirmed influenza (RT-PCR) An ILI episode was defined as measured fever $\geq 38^{\circ}\text{C}$ or subjective fever and cough. If a child was absent, the field staff followed up by phone to identify the reason for absenteeism and to determine if the child met the ILI case definition. If a child in a participating school had an ILI episode, a trained medical technologist visited the child's household to obtain consent from the child's parent/guardian and collect a nasal swab from the child within 48 hours of symptom onset. If it was outside the 48-hour window, the sample was not collected. No safety outcomes reported.

Biswas 2019 (Continued)

Notes Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Sequence generated using a computer-based random number generator.
Allocation concealment (selection bias)	Low risk	Allocation completed prior to individuals being recruited.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	Information missing for 30 children (28 children in the control schools and 2 children in the intervention schools)
Selective reporting (reporting bias)	Unclear risk	No protocol available

Canini 2010
Study characteristics

Methods	A cluster-RCT conducted in France during the 2008 to 2009 influenza season. Households were recruited during a medical visit of a household member with a positive rapid influenza A test and symptoms lasting less than 48 hours. Households were randomised either to the mask or control group for 7 days. In the intervention arm, the index case had to wear a surgical mask from the medical visit and for a period of 5 days. The trial was initially intended to include 372 households, but was prematurely interrupted after the inclusion of 105 households (306 contacts) following the advice of an independent steering committee. Generalised estimating equations were used to test the association between the intervention and the proportion of household contacts who developed an ILI during the 7 days following the inclusion.
Participants	A total of 105 households were randomised, which represented 148 contacts in the intervention arm and 158 in the control arm. The study was conducted in 3 French regions (Ile de France, Aquitaine, and Franche-Comté) and included households of size 3 to 8. Exclusion criteria: if index patient was treated for asthma or chronic obstructive pulmonary disease or was hospitalised
Interventions	Surgical mask versus no mask. See Table 1 for details.
Outcomes	The primary endpoint was the proportion of household contacts who developed an ILI during the 7 days following inclusion. Exploratory cluster-level efficacy outcome, the proportion of households with 1 or more secondary illness in household contacts.

Canini 2010 (Continued)

A temperature over 37.8 °C with cough or sore throat was used as primary clinical case definition.

Adverse reactions due to mask-wearing

Notes Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation lists were generated by a computerised program.
Allocation concealment (selection bias)	Low risk	Randomisation was performed centrally by the GP after written consent on an interactive voice response system dedicated to the study.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	All households included in analysis.
Selective reporting (reporting bias)	Low risk	All specified outcomes reported.

Canini 2010
Study characteristics

Methods	A cluster-RCT conducted in France during the 2008 to 2009 influenza season. Households were recruited during a medical visit of a household member with a positive rapid influenza A test and symptoms lasting less than 48 hours. Households were randomised either to the mask or control group for 7 days. In the intervention arm, the index case had to wear a surgical mask from the medical visit and for a period of 5 days. The trial was initially intended to include 372 households, but was prematurely interrupted after the inclusion of 105 households (306 contacts) following the advice of an independent steering committee. Generalised estimating equations were used to test the association between the intervention and the proportion of household contacts who developed an ILI during the 7 days following the inclusion.
Participants	A total of 105 households were randomised, which represented 148 contacts in the intervention arm and 158 in the control arm. The study was conducted in 3 French regions (Ile de France, Aquitaine, and Franche-Comté) and included households of size 3 to 8. Exclusion criteria: if index patient was treated for asthma or chronic obstructive pulmonary disease or was hospitalised
Interventions	Surgical mask versus no mask. See Table 1 for details.

Canini 2010 (Continued)

Outcomes	<p>The primary endpoint was the proportion of household contacts who developed an ILI during the 7 days following inclusion. Exploratory cluster-level efficacy outcome, the proportion of households with 1 or more secondary illness in household contacts.</p> <p>A temperature over 37.8 °C with cough or sore throat was used as primary clinical case definition.</p> <p>Adverse reactions due to mask-wearing</p>
----------	--

Notes	Government funded
-------	-------------------

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation lists were generated by a computerised program.
Allocation concealment (selection bias)	Low risk	Randomisation was performed centrally by the GP after written consent on an interactive voice response system dedicated to the study.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	All households included in analysis.
Selective reporting (reporting bias)	Low risk	All specified outcomes reported.

Carabin 1999
Study characteristics

Methods	<p>Cluster-RCT carried out in DCCs in the Canadian province of Quebec between 1 September 1996 and 30 November 1997 (15 months). The aim was to test the effects of a hygiene programme on the incidence of diarrhoea and fecal contamination (data not extracted) and on colds and URTIs. The design included before and after periods analysed to assess the Hawthorne effect of study participation on control DCCs. The unit of randomisation was DCC, but analysis was also carried out at classroom and single-child level. This is a common mistake in cluster-RCT analysis. DCCs were stratified by URTI incidence preceding the trial and randomised by location. Cluster coefficients are not reported.</p>
Participants	<p>A total of 1729 children aged 18 to 36 months in 47 DCCs (83 toddler classrooms)</p> <p>Inclusion criteria: presence of at least 1 sandbox and 1 play area and of at least 12 available toddler places</p> <p>For the autumn of 1997 intervention group (24 DCCs, 43 classrooms, and 414 children), control group (23 DCCs, 23 classrooms, and 374 children). It is not clear what is the distribution and data for the autumn of 1996.</p>

Carabin 1999 (Continued)

Interventions	Training session (1 day) with washing of hands, toy cleaning, window opening, sand pit cleaning, and repeated exhortations to hand wash
Outcomes	Laboratory: N/A Effectiveness: diarrhoea and coliform contamination (data not extracted) Colds (nasal discharge with at least 1 of the following: fever, sneezing, cough, sore throat, earache, malaise, irritability) URTI (cold of at least 2 days' duration) Surveillance was carried out by educators, annotating absences or illness on calendars. Researchers also filled in a phone questionnaire with answers by DCC directors. Safety: N/A
Notes	Risk of bias: high (no description of randomisation; partial reporting of outcomes, numerators, and denominators) Notes: the authors conclude that the intervention reduced the incidence of colds (IRR 0.80, 95% CI 0.68 to 0.93). This was a confusingly written study with unclear interweaving of 2 study designs. For unclear reasons analysis was only carried out for the first autumn. Unclear why colds are not reported in the results. Cluster-coefficients and randomisation process were not described.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Block randomisation of DCC according to region, but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	Not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Blinding not possible (hygiene session plus educational material versus none)
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded
Incomplete outcome data (attrition bias) All outcomes	High risk	Originally 52 eligible DCCs with 89 classrooms agreed to take part, but 5 dropped out (2 closed, 1 was sold, 2 either did not provide data or the data were "unreliable", and 6 classrooms had insufficient data). 43 children failing to attend DCC for at least 5 days in the autumn were also excluded. ITT analysis was carried out including an additional DCC whose director refused to let staff attend the training session. No correction made for clustering.
Selective reporting (reporting bias)	High risk	Denominators unclear and not explained

Chard 2019
Study characteristics

Methods	Cluster-RCT conducted amongst 100 randomly selected primary schools lacking functional WASH facilities in Saravane Province, Lao People's Democratic Republic. Schools were randomly assigned to either
---------	---

Chard 2019 (Continued)

the intervention (n = 50) or comparison (n = 50) arm. Intervention schools received a school water supply, sanitation facilities, hand-washing facilities, drinking water filters, and behaviour change education and promotion. Comparison schools received the intervention after research activities had ended. At unannounced visits every 6 to 8 weeks, enumerators recorded pupils' roll-call absence, enrolment, attrition, progression to the next grade, and reported illness (diarrhoea, respiratory infection, conjunctivitis), and conducted structured observations to measure intervention fidelity and adherence. Stool samples were collected annually prior to de-worming and analysed for soil-transmitted helminth (STH) infection. In addition to our primary ITT analysis, we conducted secondary analyses to quantify the role of intervention fidelity and adherence on project impacts.

Participants	100 primary schools (50 intervention, 50 comparison) with a total of 3993 pupils were enrolled throughout the study period (intervention schools = 2021 pupils, control schools = 1972 pupils). Up to 40 pupils selected from grades 3 to 5 in each school using systematic stratified sampling, with grade and sex as the stratification variables. Pupils selected at baseline were followed throughout the entire study period; pupils who left the school due to abandonment or transfer were replaced at the beginning of the following academic year, maintaining equal grade and sex ratios when possible. Pupils who progressed from fifth to the sixth grade were replaced with pupils from grade 3 the following academic year.
Interventions	Water supply, sanitation facilities, hand-washing facilities, drinking water filters, and behaviour change education and promotion versus control. See Table 1 for details.
Outcomes	<p>Primary impact of interest was pupil absence, measured by school-wide roll-call at each visit.</p> <p>Secondary health impacts included diarrhoea, symptoms of respiratory infection, and conjunctivitis/non-vision-related eye illness collected through pupil interviews.</p> <p>Pupils were considered to have symptoms of respiratory infection if they reported cough, runny nose, stuffy nose, or sore throat.</p> <p>No safety outcomes reported.</p>
Notes	Funded by government and pharmaceutical industry

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient details provided.
Allocation concealment (selection bias)	Low risk	Schools allocated prior to recruitment of individuals.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded
Incomplete outcome data (attrition bias) All outcomes	Low risk	Exclusions were due to participants leaving school, hence unlikely to cause bias.
Selective reporting (reporting bias)	Low risk	All specified outcomes reported.

Carabin 1999
Study characteristics

Methods	Cluster-RCT carried out in DCCs in the Canadian province of Quebec between 1 September 1996 and 30 November 1997 (15 months). The aim was to test the effects of a hygiene programme on the incidence of diarrhoea and fecal contamination (data not extracted) and on colds and URTIs. The design included before and after periods analysed to assess the Hawthorne effect of study participation on control DCCs. The unit of randomisation was DCC, but analysis was also carried out at classroom and single-child level. This is a common mistake in cluster-RCT analysis. DCCs were stratified by URTI incidence preceding the trial and randomised by location. Cluster coefficients are not reported.
Participants	A total of 1729 children aged 18 to 36 months in 47 DCCs (83 toddler classrooms) Inclusion criteria: presence of at least 1 sandbox and 1 play area and of at least 12 available toddler places For the autumn of 1997 intervention group (24 DCCs, 43 classrooms, and 414 children), control group (23 DCCs, 23 classrooms, and 374 children). It is not clear what is the distribution and data for the autumn of 1996.
Interventions	Training session (1 day) with washing of hands, toy cleaning, window opening, sand pit cleaning, and repeated exhortations to hand wash
Outcomes	Laboratory: N/A Effectiveness: diarrhoea and coliform contamination (data not extracted) Colds (nasal discharge with at least 1 of the following: fever, sneezing, cough, sore throat, earache, malaise, irritability) URTI (cold of at least 2 days' duration) Surveillance was carried out by educators, annotating absences or illness on calendars. Researchers also filled in a phone questionnaire with answers by DCC directors. Safety: N/A
Notes	Risk of bias: high (no description of randomisation; partial reporting of outcomes, numerators, and denominators) Notes: the authors conclude that the intervention reduced the incidence of colds (IRR 0.80, 95% CI 0.68 to 0.93). This was a confusingly written study with unclear interweaving of 2 study designs. For unclear reasons analysis was only carried out for the first autumn. Unclear why colds are not reported in the results. Cluster-coefficients and randomisation process were not described.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Block randomisation of DCC according to region, but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	Not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Blinding not possible (hygiene session plus educational material versus none)
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded

Carabin 1999 (Continued)

Incomplete outcome data (attrition bias) All outcomes	High risk	Originally 52 eligible DCCs with 89 classrooms agreed to take part, but 5 dropped out (2 closed, 1 was sold, 2 either did not provide data or the data were "unreliable", and 6 classrooms had insufficient data). 43 children failing to attend DCC for at least 5 days in the autumn were also excluded. ITT analysis was carried out including an additional DCC whose director refused to let staff attend the training session. No correction made for clustering.
Selective reporting (reporting bias)	High risk	Denominators unclear and not explained

Chard 2019
Study characteristics

Methods	Cluster-RCT conducted amongst 100 randomly selected primary schools lacking functional WASH facilities in Saravane Province, Lao People's Democratic Republic. Schools were randomly assigned to either the intervention (n = 50) or comparison (n = 50) arm. Intervention schools received a school water supply, sanitation facilities, hand-washing facilities, drinking water filters, and behaviour change education and promotion. Comparison schools received the intervention after research activities had ended. At unannounced visits every 6 to 8 weeks, enumerators recorded pupils' roll-call absence, enrolment, attrition, progression to the next grade, and reported illness (diarrhoea, respiratory infection, conjunctivitis), and conducted structured observations to measure intervention fidelity and adherence. Stool samples were collected annually prior to de-worming and analysed for soil-transmitted helminth (STH) infection. In addition to our primary ITT analysis, we conducted secondary analyses to quantify the role of intervention fidelity and adherence on project impacts.
Participants	100 primary schools (50 intervention, 50 comparison) with a total of 3993 pupils were enrolled throughout the study period (intervention schools = 2021 pupils, control schools = 1972 pupils). Up to 40 pupils selected from grades 3 to 5 in each school using systematic stratified sampling, with grade and sex as the stratification variables. Pupils selected at baseline were followed throughout the entire study period; pupils who left the school due to abandonment or transfer were replaced at the beginning of the following academic year, maintaining equal grade and sex ratios when possible. Pupils who progressed from fifth to the sixth grade were replaced with pupils from grade 3 the following academic year.
Interventions	Water supply, sanitation facilities, hand-washing facilities, drinking water filters, and behaviour change education and promotion versus control. See Table 1 for details.
Outcomes	Primary impact of interest was pupil absence, measured by school-wide roll-call at each visit. Secondary health impacts included diarrhoea, symptoms of respiratory infection, and conjunctivitis/non-vision-related eye illness collected through pupil interviews. Pupils were considered to have symptoms of respiratory infection if they reported cough, runny nose, stuffy nose, or sore throat. No safety outcomes reported.
Notes	Funded by government and pharmaceutical industry

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient details provided.

Chard 2019 (Continued)

Allocation concealment (selection bias)	Low risk	Schools allocated prior to recruitment of individuals.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded
Incomplete outcome data (attrition bias) All outcomes	Low risk	Exclusions were due to participants leaving school, hence unlikely to cause bias.
Selective reporting (reporting bias)	Low risk	All specified outcomes reported.

Correa 2012
Study characteristics

Methods	Cluster-RCT in childcare facilities in Colombia from 16 April to 18 December 2008 (3 school terms) testing the effects of hand hygiene using an alcohol-based hand rub versus standard practice
Participants	42 childcare facilities in 6 towns in Colombia. A total of 1727 were enrolled (intervention group = 794 from 21 centres, control group = 933 from 21 centres). Inclusion criteria: licensed to care for 12 or more children aged 1 to 5 years for 8 hours a day, 5 times per week, and where availability of tap water was limited
Interventions	Intervention: alcohol-based hand wash as an addition to hand-washing Control: usual hand-washing practice See Table 1 for details.
Outcomes	ARI defined as: 2 or more of the following symptoms for at least 24 hours, lasting at least 2 days: runny, stuffy, or blocked nose or noisy breathing; cough; fever, hot sensation, or chills; and/or sore throat. Ear pain alone was considered an ARI.
Notes	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"...using the random function in Microsoft Excel™ (Microsoft Corp., Redmond, Washington, United States), random numbers (1 or 2) were generated and allotted 1:1 within each group. Finally, a researcher flipped a coin to decide which number would correspond to either arm (heads = 1, intervention; tails = 2, control)."
Allocation concealment (selection bias)	Unclear risk	Method not described

Correa 2012 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Lost to follow-up similar in each group and not substantial
Selective reporting (reporting bias)	Unclear risk	No protocol available

Cowling 2008
Study characteristics

Methods	Cluster-RCT carried out in Hong Kong SARS between February and September 2007. The study assessed the effects of non-pharmaceutical interventions on the household transmission of influenza over a 9-day period. ILI cases whose family contacts had been symptom-free for at least 2 weeks rapid-tested for influenza A and B were used and randomised to 3 interventions. Randomisation was carried out in 2 different schedules (2:1:1 for the first 100 households, and subsequently 8:1:1), but it is unclear why and how this was done.
Participants	<p>A total of 350 of 944 originally enrolled participants representing 122 households were analysed (control group = 71 households with 205 household contacts, face mask = 21 households with 61 household contacts, HH = 30 households with 84 household contacts).</p> <p>Inclusion criteria: residents of Hong Kong aged at least 2 years, reporting at least 2 symptoms of ILI (such as fever \geq 38 degrees, cough, headache, coryza, sore throat, muscle aches and pains) and positive influenza A+B rapid test and living in a household with at least 2 other individuals, none of whom had ILI in the preceding 14 days</p> <p>Households were excluded because subsequent laboratory testing (culture) was negative.</p> <p>Attrition was not explained.</p>
Interventions	Households were randomised to either wearing face masks with education (as the control group plus education about face mask use) or hand-washing with special medicated soap (with alcohol sanitiser) with education (as the control group plus education about hand-washing) or education about general healthy lifestyle and diet (control group). The soap was distributed in special containers that were weighed at the start and end of the study. Interventions visits to the households were done on average 1 day after randomisation of index case household.
Outcomes	<p>Laboratory: QuickVue RTI MDCK culture Samples were harvested using NTS, but the text refers to a second procedure from June 2007 onwards testing for non-influenza viruses, with no data reported.</p> <p>Effectiveness: secondary attack ratios (SAR): SAR is the proportion of household contacts of an index case who were subsequently ill with influenza (symptomatic contact individuals with at least 1 NTS positive for influenza by viral culture or PCR)</p>

Cowling 2008 (Continued)

3 clinical definitions were used for secondary analysis:

1. Fever \geq 38 degrees, or at least 2 of following symptoms: headache, coryza, sore throat, muscle aches and pains
2. At least 2 of the following S/S: fever \geq 37.8 degrees, cough, headache, sore throat, muscle aches and pains
3. Fever \geq 37.8 degrees plus cough or sore throat

Safety: no harms were reported in any of the arms

Notes

The trial authors conclude that "The secondary attack ratios were lower than anticipated, and lower than reported in other countries, perhaps due to differing patterns of susceptibility, lack of significant antigenic drift in circulating influenza virus strains recently, and/or issues related to the symptomatic recruitment design. Lessons learnt from this pilot have informed changes for the main study in 2008". Although billed as a pilot study, the text is highly confusing and at times contradictory. The intervention was delivered at a home visit up to 36 hours after the index case was seen in the outpatients. This is a long time and perhaps the reason for failure of the intervention. Practically, the intervention will have to be organised before even seeking medical care, i.e. people know to do it when the child gets sick at home.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation was computer generated by a biostatistician. "A pre-specified table of random numbers will be used to assign one of the three interventions to the household of the index case."
Allocation concealment (selection bias)	Low risk	The households of eligible study index patients were allocated to 3 groups in a 1:1:1 ratio under a block randomisation structure with randomly permuted block sizes of 18, 24, and 30 using a random-number generator. Allocation was concealed from treating physicians and clinics and implemented by study nurses at the time of the initial household visit.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Participants and people who administered the interventions were not blinded to the interventions, but participants were not informed of the specific nature of the interventions applied to other participating households.
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded
Incomplete outcome data (attrition bias) All outcomes	High risk	Dropout was accounted for. Dropout from the randomised population was high: 32% in control group, 37.5% in hand hygiene group, and 39.4% in face mask and hand hygiene group. Reasons for dropout were distributed evenly across the 3 groups. Authors report follow-up as proportion of patients remaining in the study after initial dropout.
Selective reporting (reporting bias)	High risk	The choice of season, change in randomisation schedules, and unexplained dropouts amongst contacts; the use of QuickVue, which proved unreliable, reporting bias on non-influenza isolates resulted in a judgement of high risk of bias.

Cowling 2009
Study characteristics

Methods	Cluster-RCT
Participants	<p>A total of 407 index cases and 794 household contacts were analysed.</p> <p>Of 407 enrolled households, 322 received the allocated interventions as follows:</p> <ol style="list-style-type: none"> 1. control group = 112 households with 346 contacts (only 91 households analysed with 279 contacts); 2. hand hygiene = 106 households with 329 contacts (only 85 households analysed with 257 contacts); 3. face mask + hand hygiene = 104 households with 340 contacts (only 83 households analysed with 258 contacts). <p>Inclusion criteria: households in Hong Kong. Index cases from 45 outpatient clinics in both the private and public sectors across Hong Kong. They enrolled individuals who reported at least 2 symptoms of ARI (temperature 37.8 °C, cough, headache, sore throat, or myalgia); had symptom onset within 48 hours; and lived in a household with at least 2 other people, none of whom had reported ARI in the preceding 14 days. After giving informed consent, participants provided nasal and throat swab specimens.</p> <p>2750 patients were eligible and tested between 2 January and 30 September 2008.</p>
Interventions	<p>Participants with a positive rapid-test result and their household contacts were randomly assigned to 1 of 3 study groups: control (lifestyle measures - 134 households), control plus enhanced hand hygiene only (136 households), and control plus face masks and enhanced hand hygiene (137 households) for all household members. No detailed description of the instructions was given to participants.</p>
Outcomes	<p>Influenza virus infection in household contacts, as confirmed by RT-PCR or diagnosed clinically after 7 days</p> <p>"The primary outcome measure was the secondary attack ratio at the individual level: that is, the proportion of household contacts infected with influenza virus. We evaluated the secondary attack ratio using a laboratory definition (a household contact with a nose and throat swab specimen positive for influenza by RT-PCR) as the primary analysis and 2 secondary clinical definitions of influenza based on self-reported data from the symptom diaries as secondary analyses."</p> <p>Statistical analysis: adjusted for clustering Results: no statistically significant difference in secondary attack ratio between groups in total population. Statistically significant reduction in RT-PCR confirmed influenza virus infections in the household contacts in 154 households in which the intervention was applied within 36 hours of symptom onset in the index patient. Adherence to hand hygiene was between 44% and 62%. Adherence of index patient to wearing a face mask between 15% and 49%.</p>
Notes	<p>"In an unintentional deviation from that protocol, 49 of the 407 randomly allocated persons had a household contact with influenza symptoms at recruitment (a potential co-index patient). We also randomly assigned 6 of 407 persons who had symptoms for slightly more than 48 hours."</p> <p>The trial authors conclude that "Hand hygiene and face masks seemed to prevent household transmission of influenza virus when implemented within 36 hours of index patient symptom onset. These findings suggest that non-pharmaceutical interventions are important for mitigation of pandemic and inter-pandemic influenza".</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	<p>Randomisation was computer generated by a biostatistician.</p> <p>"A pre-specified table of random numbers will be used to assign one of the three interventions to the household of the index case."</p>

Cowling 2009 (Continued)

Allocation concealment (selection bias)	Low risk	The households of eligible study index patients were allocated to 3 groups in a 1:1:1 ratio under a block randomisation structure with randomly permuted block sizes of 18, 24, and 30 using a random-number generator. Allocation was concealed from treating physicians and clinics and implemented by study nurses at the time of the initial household visit.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Quote: "Participants and personnel administering the interventions were not blinded to group assignment."
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	It is not stated if the outcome assessor was blinded.
Incomplete outcome data (attrition bias) All outcomes	High risk	Dropout was accounted for. Dropout from the randomised population was high: 32% in control group, 37.5% in hand hygiene group, and 39.4% in face mask and hand hygiene group. Reasons for dropout were distributed evenly across the 3 groups. Trial authors report follow-up as proportion of patients remaining in the study after initial dropout.
Selective reporting (reporting bias)	Unclear risk	In general good reporting

DiVita 2011
Study characteristics

Methods	The impact of hand-washing promotion on the risk of household transmission of influenza, ILI, and fever was tested in rural Bangladesh. ILI was defined as fever in children < 5 years old and fever with cough or sore throat in individuals > 5 years old. Households were randomised to intervention or control. The intervention group received hand-washing stations with soap and daily hand-washing motivation at critical times for pathogen transmission, such as after coughing or sneezing. Daily surveillance was conducted, and household members with fever were tested for influenza viruses by PCR. Secondary attack ratios (SAR) were calculated for influenza, ILI, and fever in each arm. Logistic regression with generalised estimating equations was used to estimate the significance of the SAR comparison whilst controlling for clustering by household.
Participants	The study included 233 patient index cases (intervention group = 100, control group 133) with 2540 household contacts (intervention group = 134, control group = 1226). Inclusion criteria: index case patients (individuals who developed ILI within the previous 2 days and were the only symptomatic person in their household) as well as their household contacts
Interventions	Hand-washing stations with soap and daily hand-washing motivation versus control. See Table 1 for details.
Outcomes	SAR were calculated for influenza, ILI, and fever. ILI was defined as fever in children < 5 years old and fever with cough or sore throat in individuals > 5 years old. No safety outcomes reported.
Notes	Funding source unknown

DiVita 2011 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient details provided
Allocation concealment (selection bias)	Unclear risk	Insufficient details provided
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Insufficient details provided
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Insufficient details provided
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient details provided
Selective reporting (reporting bias)	Unclear risk	Insufficient details provided

Cowling 2008
Study characteristics

Methods	<p>Cluster-RCT carried out in Hong Kong SARS between February and September 2007. The study assessed the effects of non-pharmaceutical interventions on the household transmission of influenza over a 9-day period. ILI cases whose family contacts had been symptom-free for at least 2 weeks rapid-tested for influenza A and B were used and randomised to 3 interventions. Randomisation was carried out in 2 different schedules (2:1:1 for the first 100 households, and subsequently 8:1:1), but it is unclear why and how this was done.</p>
Participants	<p>A total of 350 of 944 originally enrolled participants representing 122 households were analysed (control group = 71 households with 205 household contacts, face mask = 21 households with 61 household contacts, HH = 30 households with 84 household contacts).</p> <p>Inclusion criteria: residents of Hong Kong aged at least 2 years, reporting at least 2 symptoms of ILI (such as fever ≥ 38 degrees, cough, headache, coryza, sore throat, muscle aches and pains) and positive influenza A+B rapid test and living in a household with at least 2 other individuals, none of whom had ILI in the preceding 14 days</p> <p>Households were excluded because subsequent laboratory testing (culture) was negative.</p> <p>Attrition was not explained.</p>
Interventions	<p>Households were randomised to either wearing face masks with education (as the control group plus education about face mask use) or hand-washing with special medicated soap (with alcohol sanitiser) with education (as the control group plus education about hand-washing) or education about general healthy lifestyle and diet (control group). The soap was distributed in special containers that were</p>

Cowling 2008 (Continued)

weighed at the start and end of the study. Interventions visits to the households were done on average 1 day after randomisation of index case household.

Outcomes	<p>Laboratory: QuickVue RTI MDCK culture Samples were harvested using NTS, but the text refers to a second procedure from June 2007 onwards testing for non-influenza viruses, with no data reported.</p> <p>Effectiveness: secondary attack ratios (SAR): SAR is the proportion of household contacts of an index case who were subsequently ill with influenza (symptomatic contact individuals with at least 1 NTS positive for influenza by viral culture or PCR)</p> <p>3 clinical definitions were used for secondary analysis:</p> <ol style="list-style-type: none"> 1. Fever \geq 38 degrees, or at least 2 of following symptoms: headache, coryza, sore throat, muscle aches and pains 2. At least 2 of the following S/S: fever \geq 37.8 degrees, cough, headache, sore throat, muscle aches and pains 3. Fever \geq 37.8 degrees plus cough or sore throat <p>Safety: no harms were reported in any of the arms</p>
Notes	<p>The trial authors conclude that "The secondary attack ratios were lower than anticipated, and lower than reported in other countries, perhaps due to differing patterns of susceptibility, lack of significant antigenic drift in circulating influenza virus strains recently, and/or issues related to the symptomatic recruitment design. Lessons learnt from this pilot have informed changes for the main study in 2008". Although billed as a pilot study, the text is highly confusing and at times contradictory. The intervention was delivered at a home visit up to 36 hours after the index case was seen in the outpatients. This is a long time and perhaps the reason for failure of the intervention. Practically, the intervention will have to be organised before even seeking medical care, i.e. people know to do it when the child gets sick at home.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation was computer generated by a biostatistician. "A pre-specified table of random numbers will be used to assign one of the three interventions to the household of the index case."
Allocation concealment (selection bias)	Low risk	The households of eligible study index patients were allocated to 3 groups in a 1:1:1 ratio under a block randomisation structure with randomly permuted block sizes of 18, 24, and 30 using a random-number generator. Allocation was concealed from treating physicians and clinics and implemented by study nurses at the time of the initial household visit.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Participants and people who administered the interventions were not blinded to the interventions, but participants were not informed of the specific nature of the interventions applied to other participating households.
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded
Incomplete outcome data (attrition bias) All outcomes	High risk	Dropout was accounted for. Dropout from the randomised population was high: 32% in control group, 37.5% in hand hygiene group, and 39.4% in face

Cowling 2008 (Continued)

mask and hand hygiene group. Reasons for dropout were distributed evenly across the 3 groups.

Authors report follow-up as proportion of patients remaining in the study after initial dropout.

Selective reporting (reporting bias)	High risk	The choice of season, change in randomisation schedules, and unexplained dropouts amongst contacts; the use of QuickVue, which proved unreliable, reporting bias on non-influenza isolates resulted in a judgement of high risk of bias.
--------------------------------------	-----------	--

Cowling 2009
Study characteristics

Methods	Cluster-RCT
Participants	<p>A total of 407 index cases and 794 household contacts were analysed.</p> <p>Of 407 enrolled households, 322 received the allocated interventions as follows:</p> <ol style="list-style-type: none"> 1. control group = 112 households with 346 contacts (only 91 households analysed with 279 contacts); 2. hand hygiene = 106 households with 329 contacts (only 85 households analysed with 257 contacts); 3. face mask + hand hygiene = 104 households with 340 contacts (only 83 households analysed with 258 contacts). <p>Inclusion criteria: households in Hong Kong. Index cases from 45 outpatient clinics in both the private and public sectors across Hong Kong. They enrolled individuals who reported at least 2 symptoms of ARI (temperature 37.8 °C, cough, headache, sore throat, or myalgia); had symptom onset within 48 hours; and lived in a household with at least 2 other people, none of whom had reported ARI in the preceding 14 days. After giving informed consent, participants provided nasal and throat swab specimens.</p> <p>2750 patients were eligible and tested between 2 January and 30 September 2008.</p>
Interventions	Participants with a positive rapid-test result and their household contacts were randomly assigned to 1 of 3 study groups: control (lifestyle measures - 134 households), control plus enhanced hand hygiene only (136 households), and control plus face masks and enhanced hand hygiene (137 households) for all household members. No detailed description of the instructions was given to participants.
Outcomes	<p>Influenza virus infection in household contacts, as confirmed by RT-PCR or diagnosed clinically after 7 days</p> <p>"The primary outcome measure was the secondary attack ratio at the individual level: that is, the proportion of household contacts infected with influenza virus. We evaluated the secondary attack ratio using a laboratory definition (a household contact with a nose and throat swab specimen positive for influenza by RT-PCR) as the primary analysis and 2 secondary clinical definitions of influenza based on self-reported data from the symptom diaries as secondary analyses."</p> <p>Statistical analysis: adjusted for clustering</p> <p>Results: no statistically significant difference in secondary attack ratio between groups in total population. Statistically significant reduction in RT-PCR confirmed influenza virus infections in the household contacts in 154 households in which the intervention was applied within 36 hours of symptom onset in the index patient. Adherence to hand hygiene was between 44% and 62%. Adherence of index patient to wearing a face mask between 15% and 49%.</p>
Notes	"In an unintentional deviation from that protocol, 49 of the 407 randomly allocated persons had a household contact with influenza symptoms at recruitment (a potential co-index patient). We also randomly assigned 6 of 407 persons who had symptoms for slightly more than 48 hours."

Cowling 2009 (Continued)

The trial authors conclude that "Hand hygiene and face masks seemed to prevent household transmission of influenza virus when implemented within 36 hours of index patient symptom onset. These findings suggest that non-pharmaceutical interventions are important for mitigation of pandemic and inter-pandemic influenza".

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation was computer generated by a biostatistician. "A pre-specified table of random numbers will be used to assign one of the three interventions to the household of the index case."
Allocation concealment (selection bias)	Low risk	The households of eligible study index patients were allocated to 3 groups in a 1:1:1 ratio under a block randomisation structure with randomly permuted block sizes of 18, 24, and 30 using a random-number generator. Allocation was concealed from treating physicians and clinics and implemented by study nurses at the time of the initial household visit.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Quote: "Participants and personnel administering the interventions were not blinded to group assignment."
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	It is not stated if the outcome assessor was blinded.
Incomplete outcome data (attrition bias) All outcomes	High risk	Dropout was accounted for. Dropout from the randomised population was high: 32% in control group, 37.5% in hand hygiene group, and 39.4% in face mask and hand hygiene group. Reasons for dropout were distributed evenly across the 3 groups. Trial authors report follow-up as proportion of patients remaining in the study after initial dropout.
Selective reporting (reporting bias)	Unclear risk	In general good reporting

DiVita 2011
Study characteristics

Methods	The impact of hand-washing promotion on the risk of household transmission of influenza, ILI, and fever was tested in rural Bangladesh. ILI was defined as fever in children < 5 years old and fever with cough or sore throat in individuals > 5 years old. Households were randomised to intervention or control. The intervention group received hand-washing stations with soap and daily hand-washing motivation at critical times for pathogen transmission, such as after coughing or sneezing. Daily surveillance was conducted, and household members with fever were tested for influenza viruses by PCR. Secondary attack ratios (SAR) were calculated for influenza, ILI, and fever in each arm. Logistic regression with generalised estimating equations was used to estimate the significance of the SAR comparison whilst controlling for clustering by household.
Participants	The study included 233 patient index cases (intervention group = 100, control group 133) with 2540 household contacts (intervention group = 134, control group = 1226).

DiVita 2011 (Continued)

Inclusion criteria: index case patients (individuals who developed ILI within the previous 2 days and were the only symptomatic person in their household) as well as their household contacts

Interventions Hand-washing stations with soap and daily hand-washing motivation versus control. See [Table 1](#) for details.

Outcomes SAR were calculated for influenza, ILI, and fever.

ILI was defined as fever in children < 5 years old and fever with cough or sore throat in individuals > 5 years old.

No safety outcomes reported.

Notes Funding source unknown

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient details provided
Allocation concealment (selection bias)	Unclear risk	Insufficient details provided
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Insufficient details provided
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Insufficient details provided
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient details provided
Selective reporting (reporting bias)	Unclear risk	Insufficient details provided

Farr 1988a
Study characteristics

Methods 6-month cluster-RCT, controlled, double-blind of the efficacy of virucidal nasal tissues in the prevention of natural cold, conducted in Charlottesville, Virginia, USA. Many of the families were enrolled because 1 or more family members worked at the State Farm Insurance Company; the remaining families were recruited from the Charlottesville community by advertisement in a local newspaper. Families were randomly assigned by the sponsoring company to receive boxes of treated tissues, placebo tissues, or no tissues. The randomisation was performed by computer. Study participants and investigators were unaware of the type of tissues each family was randomised to receive. Blinding efficacy was tested using a questionnaire: the mothers in each family were asked twice if she believed her family was using virucidal or placebo tissues.

Participants in the treated and placebo groups were instructed to use only tissues received through the study, whilst families in the additional control group without tissues were allowed to continue their

Farr 1988a (Continued)

usual practice of personal hygiene. Each family member kept a daily listing of respiratory symptoms on a record card. A nurse epidemiologist visited each family monthly to encourage recording.

Participants	186 families, 58 in the active group, 59 in the placebo group, and 69 in the no-tissues group. A total of 302 families were originally recruited; 116 families who did not comply with the study protocol, lost their surveillance cards, could not complete the protocol were excluded from the analysis.
Interventions	Use of virucidal tissues versus placebo tissues versus no tissues. The treated tissues were impregnated with malic and citric acids and sodium lauryl sulphate, whilst placebo tissues contained saccharin.
Outcomes	Laboratory: serological evidence: no Effectiveness: respiratory illness Safety: N/A
Notes	The authors concluded that virucidal tissues have only a small impact on the overall rate of natural acute respiratory illnesses. The total illness rate was lower in families using virucidal tissues than in both of the other study groups, but only the difference between active and placebo groups was statistically significant (3.4 illness per person versus 3.9 for placebo group, $P = 0.04$, and 3.6 for the no-tissue control group, $P = 0.2$, and overall 14% to 5% reduction). The questionnaire results suggest that some bias may have been present since a majority of mothers in the virucide group believed they were receiving the 'active' tissues. Another possible explanation of the low effectiveness of virucidal tissues is poor compliance by children in use of the virucidal tissues. A well-designed and honestly reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"The randomisation was performed by computer in each trial." However, method of sequence generation is not stated.
Allocation concealment (selection bias)	Unclear risk	"In trial I, families were randomly assigned by the sponsoring company to receive boxes of treated tissues, placebo tissues or no tissues." "Families with one or two children were randomised in one stratum, and families with three or more children were randomised in a second stratum in trial I." Concealment of allocation not described
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Study participants and investigators were unaware of the type of tissues which each family was randomised to receive in both trials. In trial I, the mother in each family was asked twice if she believed her family was using active or placebo tissues, first after three months and then at the end of the study."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Study participants and investigators were unaware of the type of tissues which each family was randomised to receive in both trials. In trial I, the mother in each family was asked twice if she believed her family was using active or placebo tissues, first after three months and then at the end of the study."
Incomplete outcome data (attrition bias) All outcomes	High risk	"A total of 116 of the 302 families were excluded from the analysis. Families were excluded if they lost their surveillance cards or did not conscientiously record data, did not comply with the study protocol, or simply could not complete the protocol for family reasons. It was discovered that families with five or more members had so many colds that it was not possible to distinguish primary and secondary illnesses. These large families were therefore excluded from the analysis in trial I and were excluded from enrolment in trial II."

Farr 1988a (Continued)

Selective reporting (reporting bias)	Low risk	All indicated outcomes are reported.
--------------------------------------	----------	--------------------------------------

Farr 1988b
Study characteristics

Methods	6-month randomised, controlled, double-blind trial of the efficacy of virucidal nasal tissues in the prevention of natural cold, conducted in Charlottesville, Virginia, USA. Families were recruited from the Charlottesville community by advertisement in a local newspaper. Families were randomly assigned by the sponsoring company to receive either virucidal tissues or placebo-treated tissues. Stratified randomisation was performed by computer, and the strata were defined by total number in the family. Study participants and investigators were unaware of the type of tissues each family was randomised to receive. Each family member kept a daily listing of respiratory symptoms on a record card. A nurse epidemiologist visited each family monthly to encourage recording. In addition, a study monitor visited each family bimonthly to further encourage compliance and reporting of symptoms.
Participants	98 families, 58 in the active group and 40 in the placebo group. 231 families were initially recruited, 222 completed the trial, data of 98 families were analysed. The other families were excluded from the analysis because they complained of side effects (sneezing, etc.) or reported not using the tissues regularly.
Interventions	Use of virucidal tissues versus placebo tissues. The treated tissues were impregnated with malic and citric acids and sodium lauryl sulphate, whilst the placebo tissues contained succinic acid. Participants in the treated and placebo groups were instructed to only use tissues received through the study.
Outcomes	Laboratory: serological evidence: no Effectiveness: respiratory illness Safety: N/A
Notes	The study suggests that virucidal tissues have only a small impact on the overall rate of natural acute respiratory illnesses. The total illness rate was lower in families using virucidal tissues than in the other study group, but the difference between active and placebo groups was not statistically significant. There was a small, non-significant drop in illness rates across families (5%). The tissues appeared to be ineffective as the drop was confined to primary illness unaffected by tissue use. The placebo (succinic acid) was not inert, and was associated with cough and nasal burning. This impacted on allocation concealment. A well-designed and honestly reported study marred by transparent allocation

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"The randomisation was performed by computer in each trial." However, method of sequence generation is not stated.
Allocation concealment (selection bias)	Unclear risk	"In trial II, families were randomly assigned by the sponsor to receive either virucidal tissues or placebo treated tissues." "In trial II, stratified randomisation was again used, but this time the strata were defined by total number in the family (i.e., one stratum for two-member families, another stratum for three-member families, and a final one for four-member families)." Concealment of allocation not described

Farr 1988b (Continued)

Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Study participants and investigators were unaware of the type of tissues which each family was randomised to receive in both trials."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Study participants and investigators were unaware of the type of tissues which each family was randomised to receive in both trials."
Incomplete outcome data (attrition bias) All outcomes	High risk	"A total of 222 (of 231) families completed trial II; 9 families were terminated early (table 1). In 124 families, one or more family members reported not using the tissues regularly and/or reported having significant side effects. The data from these families were not analysed, leaving 58 families (177 persons) and 40 families (114 persons) for analysis in the virucide and placebo groups, respectively."
Selective reporting (reporting bias)	Low risk	All indicated outcomes are reported.

Feldman 2016

Study characteristics

Methods	Prospective cluster-RCT. Ships from a single, central naval base. Ships were stratified by vessel classes (corvette, fast missile boat, and patrol boat).
Participants	All people participating in security operations, routine exercises, and patrol at a single, central naval base were eligible. The actual number of participants in the groups is not reported.
Interventions	Chlorhexidine gluconate (CHG) dispensers in addition to soap-and-water hand-washing versus soap-and-water hand-washing. See Table 1 for details.
Outcomes	Laboratory: bacterial palm cultures from 30 sailors from each group using a modified bag broth technique with sterile brain-heart broth, at 0 and 4 months (sample participants) Effectiveness: Primary outcome: incidence of infectious diseases reported by the computerised patient records system using ICD-9 diagnoses and grouped into diarrhoeal, respiratory, and skin infections; the number of sick call visits; and the number of sick leave and light-duty days incurred by the sailors Secondary outcome: subclinical morbidity (i.e. symptoms of self-reported infectious diseases) Safety: not reported
Notes	No report on adherence Funding: governmental (Israeli Defense Force Medical Corps) Study was conducted between May and September 2014 (4 months follow-up). CHG availability onboard the ships did not reduce the transmission of infectious diseases or colonisation.

Risk of bias

Feldman 2016 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No description of randomisation
Allocation concealment (selection bias)	Unclear risk	No description of allocation
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded. Self-reported outcomes
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No information if personnel collecting data for ICD-9 diagnosis were blinded
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	No participants flow chart, no attrition data
Selective reporting (reporting bias)	Unclear risk	No protocol to compare

Goodall 2014
Study characteristics

Methods	A 2X2 factorial RCT with 4 treatment arms: <ol style="list-style-type: none"> 1. Vitamin D₃ and gargling 2. Placebo and gargling 3. Vitamin D₃ and no gargling 4. Placebo and no gargling
Participants	600 students from McMaster University, Hamilton, Ontario, Canada, randomised to the following: <ol style="list-style-type: none"> 1. Vitamin D and gargling (N = 150, analysed 135) 2. Vitamin D and no gargling (N = 150, 123 outcomes included in analysis) 3. Placebo and gargling (N = 150, 121 known outcomes included in analysis) 4. Placebo and no gargling (N = 150, 113 known outcomes included in analysis) Inclusion criteria: aged ≥ 17 years and lived with at least 1 student housemate. Exclusion criteria: students with contraindicated medical conditions (hypercalcaemia, parathyroid disorder, chronic kidney disease, use of anticonvulsants, malabsorption syndromes, sarcoidosis), who were currently or planning to become pregnant, who were taking ≥ 1000 international units (IU)/day vitamin D, or who were unable to swallow capsules
Interventions	See TIDieR Table (Table 9).
Outcomes	Laboratory (influenza assessed via weekly self-collected nasal swabs; only swabs for symptomatic participants were assessed). Lab-confirmed influenza was determined by testing the Day 1 nasal swabs using an in-house enterovirus/rhinovirus PCR and, if negative, a commercial multiplex PCR able to detect 16 respiratory viruses and viral subtypes (xTAG RVP FAST, Luminex, Austin TX).

Goodall 2014 (Continued)

Clinical URTI assessed via weekly online surveys.

Clinical URTI is defined as the participant's perception of cold in conjunction with 1 or more symptoms (runny/stuffy nose, congestion, cough, sneezing, sore throat, muscle aches, or fever). When participants reported symptoms but were uncertain if they were ill, adjudication was applied by 2 clinicians.

Safety:

None assessed/reported by the investigators.

Notes

Study was conducted during 2 periods: September to October in 2010 and 2011.

Partial governmental funding

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No description on how the randomisation sequence was generated
Allocation concealment (selection bias)	Low risk	Study used opaque, sealed, serially numbered envelopes. Envelopes were only accessed when both personnel were present.
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Due to the nature of gargling with tap water, this intervention was not blinded. However, all other aspects of the study were blinded. Self-reported symptoms were adjudicated by 2 clinicians.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Except for gargling, all other participants and study personnel were blinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Study flow chart and reasons for lost to follow-up are provided, imputation used for missing outcomes.
Selective reporting (reporting bias)	Low risk	All planned study outcomes were reported and match the published study protocol.

Gwaltney 1980
Study characteristics

Methods

Study assessed the effectiveness of aqueous iodine applied to the fingers in blocking hand transmission of experimental infection with rhinovirus from 1 volunteer to another. Healthy, young adult volunteers were recruited from the general population at the University of Virginia, Charlottesville. Volunteers were not informed about the contents of the hand preparation until after the study. 2 experiments were conducted to evaluate the virucidal activity of aqueous iodine applied to the fingers immediately before viral contamination. Another 2 experiments were conducted to determine whether there was sufficient residual activity of aqueous iodine after 2 hours to interrupt viral spread by the hand route. Volunteers who were donors of virus for the hand exposures were challenged intranasally on 3 consecutive days with the rhinovirus strain HH. Recipients were randomly assigned to receive iodine or placebo. The donors contaminated their hands with nasal secretions by finger to nose contact before the exposure. Hand contact was made between a donor and a recipient by stroking of the fingers for 10 seconds. Donors and recipients wore masks during the exposure period.

Gwaltney 1980 (Continued)

Participants	15 and 20 volunteers in 2 experiments
Interventions	Treatment of fingers with iodine versus placebo. The virucidal preparation used was aqueous iodine (2% iodine and 4% potassium iodide). The placebo was an aqueous solution of food colours.
Outcomes	Experimental rhinovirus infection reduced (P = 0.06) Laboratory: serological evidence Effectiveness: rhinovirus infection (based on serology, isolation, and clinical symptoms) with high-score clinical illness. Score was published elsewhere. Safety: N/A
Notes	Risk of bias: high (poor description of randomisation process, concealment, or allocation) Notes: the study suggests that aqueous iodine applied to the fingers was effective in blocking transmission by hand contact of experimental infection with rhinovirus for up to 2 hours after application (1 out of 10 volunteers were infected compared to 6 out of 10 in the placebo preparation arm, P = 0.06 with Fisher's exact test). The effectiveness of iodine treatment of the fingers in interrupting viral transmission in volunteers recommends its use for attempting to block transmission of rhinovirus under natural conditions. Although the cosmetic properties of 2% aqueous iodine make it impractical for routine use, it can be used as an epidemiologic tool to study the importance of the hand transmission route and to develop an effective cosmetically acceptable hand preparation. A summarily reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Unclear risk	insufficient information
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"The virucidal preparation used was aqueous iodine... . The placebo was an aqueous solution of food colors... mixed to resemble the color of iodine. An odor of iodine was given to the placebo... . Volunteers were not informed about the contents of the hand preparation until after the study."
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	It is not stated whether the outcome assessor was blinded or not.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information
Selective reporting (reporting bias)	Unclear risk	Insufficient information

Hartinger 2016
Study characteristics

Methods	Communities were randomised to a comprehensive intervention was an improved solid-fuel stove, installation of a kitchen sink with running water, solar drinking water disinfection, education on hand-washing, and separating animals from the kitchen environment.
---------	---

Hartinger 2016 (Continued)

Participants	534 children (267 in each group) in 51 communities (25 in intervention, 26 in control group). 250 children/households in the intervention group and 253 children/households in the control group were available for follow-up. Conducted in a rural farming area
Interventions	Environmental home-based intervention package consisting of improved solid-fuel stoves, kitchen sinks, solar disinfection of drinking water, and hygiene promotion. See Table 1 for details.
Outcomes	<p>Laboratory: <i>Escherichia coli</i> (not relevant to this review)</p> <p>Effectiveness: weekly collection of daily diary data on illness. ARI was defined as child presenting cough or difficulty breathing, or both. ALRI was defined as child presenting cough or difficulty breathing, with a raised respiratory rate (> 50 per min in children aged 6 to 11 months and > 40 per min in children aged 12 months) on 2 consecutive measurements.</p> <p>Safety: none described in methods and none reported</p>
Notes	<p>The authors conclude that “combined home-based environmental interventions slightly reduced childhood diarrhoea, but the confidence interval included unity. Effects on growth and respiratory outcomes were not observed, despite high user compliance of the interventions. The absent effect on respiratory health might be due to insufficient household air quality improvements of the improved stoves and additional time needed to achieve attitudinal and behaviour change when providing composite interventions”.</p> <p>Well-reported trial. Age of children not reported.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Covariate-constrained randomisation is mentioned, but method not described.
Allocation concealment (selection bias)	Unclear risk	Method not mentioned
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data collected by field worker and recorded by parent. All would be aware of allocation.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Low attrition rate, reasons stated, balanced between groups.
Selective reporting (reporting bias)	Low risk	It is unlikely that other outcomes were measured but not reported.

Gwaltney 1980
Study characteristics

Methods	Study assessed the effectiveness of aqueous iodine applied to the fingers in blocking hand transmission of experimental infection with rhinovirus from 1 volunteer to another. Healthy, young adult vol-
---------	--

Gwaltney 1980 (Continued)

unteers were recruited from the general population at the University of Virginia, Charlottesville. Volunteers were not informed about the contents of the hand preparation until after the study. 2 experiments were conducted to evaluate the virucidal activity of aqueous iodine applied to the fingers immediately before viral contamination. Another 2 experiments were conducted to determine whether there was sufficient residual activity of aqueous iodine after 2 hours to interrupt viral spread by the hand route. Volunteers who were donors of virus for the hand exposures were challenged intranasally on 3 consecutive days with the rhinovirus strain HH. Recipients were randomly assigned to receive iodine or placebo. The donors contaminated their hands with nasal secretions by finger to nose contact before the exposure. Hand contact was made between a donor and a recipient by stroking of the fingers for 10 seconds. Donors and recipients wore masks during the exposure period.

Participants	15 and 20 volunteers in 2 experiments
Interventions	Treatment of fingers with iodine versus placebo. The virucidal preparation used was aqueous iodine (2% iodine and 4% potassium iodide). The placebo was an aqueous solution of food colours.
Outcomes	Experimental rhinovirus infection reduced ($P = 0.06$) Laboratory: serological evidence Effectiveness: rhinovirus infection (based on serology, isolation, and clinical symptoms) with high-score clinical illness. Score was published elsewhere. Safety: N/A
Notes	Risk of bias: high (poor description of randomisation process, concealment, or allocation) Notes: the study suggests that aqueous iodine applied to the fingers was effective in blocking transmission by hand contact of experimental infection with rhinovirus for up to 2 hours after application (1 out of 10 volunteers were infected compared to 6 out of 10 in the placebo preparation arm, $P = 0.06$ with Fisher's exact test). The effectiveness of iodine treatment of the fingers in interrupting viral transmission in volunteers recommends its use for attempting to block transmission of rhinovirus under natural conditions. Although the cosmetic properties of 2% aqueous iodine make it impractical for routine use, it can be used as an epidemiologic tool to study the importance of the hand transmission route and to develop an effective cosmetically acceptable hand preparation. A summarily reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Unclear risk	insufficient information
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"The virucidal preparation used was aqueous iodine... . The placebo was an aqueous solution of food colors... mixed to resemble the color of iodine. An odor of iodine was given to the placebo... . Volunteers were not informed about the contents of the hand preparation until after the study."
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	It is not stated whether the outcome assessor was blinded or not.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information
Selective reporting (reporting bias)	Unclear risk	Insufficient information

Hartinger 2016
Study characteristics

Methods	Communities were randomised to a comprehensive intervention was an improved solid-fuel stove, installation of a kitchen sink with running water, solar drinking water disinfection, education on hand-washing, and separating animals from the kitchen environment.
Participants	534 children (267 in each group) in 51 communities (25 in intervention, 26 in control group). 250 children/households in the intervention group and 253 children/households in the control group were available for follow-up. Conducted in a rural farming area
Interventions	Environmental home-based intervention package consisting of improved solid-fuel stoves, kitchen sinks, solar disinfection of drinking water, and hygiene promotion. See Table 1 for details.
Outcomes	<p>Laboratory: <i>Escherichia coli</i> (not relevant to this review)</p> <p>Effectiveness: weekly collection of daily diary data on illness. ARI was defined as child presenting cough or difficulty breathing, or both. ALRI was defined as child presenting cough or difficulty breathing, with a raised respiratory rate (> 50 per min in children aged 6 to 11 months and > 40 per min in children aged 12 months) on 2 consecutive measurements.</p> <p>Safety: none described in methods and none reported</p>
Notes	<p>The authors conclude that “combined home-based environmental interventions slightly reduced childhood diarrhoea, but the confidence interval included unity. Effects on growth and respiratory outcomes were not observed, despite high user compliance of the interventions. The absent effect on respiratory health might be due to insufficient household air quality improvements of the improved stoves and additional time needed to achieve attitudinal and behaviour change when providing composite interventions”.</p> <p>Well-reported trial. Age of children not reported.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Covariate-constrained randomisation is mentioned, but method not described.
Allocation concealment (selection bias)	Unclear risk	Method not mentioned
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data collected by field worker and recorded by parent. All would be aware of allocation.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Low attrition rate, reasons stated, balanced between groups.
Selective reporting (reporting bias)	Low risk	It is unlikely that other outcomes were measured but not reported.

Hubner 2010
Study characteristics

Methods	A prospective, controlled, intervention-control group design to assess the epidemiological and economical impact of alcohol-based hand disinfectants use at workplace. Volunteers in public administrations in the municipality of the city of Greifswald were randomised into 2 groups. Participants in the intervention group were provided with alcoholic hand disinfection, the control group was unchanged. In all, 1230 person-months were evaluated.
Participants	<p>Employees (n = 134) from the administration of the Ernst-Moritz-Arndt University Greifswald, the municipality of Greifswald and the state of Mecklenburg-Pomerania, were recruited for the study and randomised to intervention (N = 67) or control (N = 67). Final analysis was performed on 64 from the intervention and 65 from the control group.</p> <p>Inclusion criteria: all administrative officers, who did not already apply hand disinfection at work, were considered for participation and were invited by email or mail (n = 850). The 134 participants declared their written consent to participate and completed a pre-study survey with demographic, social, health, and work-related questions to provide data for randomisation.</p> <p>Exclusion criteria: employees that were already using hand disinfectants at work</p>
Interventions	Alcohol-based hand disinfectants use at workplace versus usual hygiene. See Table 1 for details.
Outcomes	Respiratory and gastrointestinal symptoms and days of work were recorded based on a monthly questionnaire over 1 year.

Notes

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No details provided.
Allocation concealment (selection bias)	Unclear risk	No details provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Self-reported outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	Lost to follow-up minimal and similar in 2 groups
Selective reporting (reporting bias)	Unclear risk	No protocol available

Huda 2012
Study characteristics

Methods	Poorly described cluster-RCT. Partial report of the SHEWA-B trial focused on changing 11 targeted behaviours in villages to measure the impact on diarrhoea and respiratory illness amongst children. Unit of randomisation is not clear, but was probably a village. A group of 10 to 17 households within a village were the participants, based on the household having at least 1 child under the age of 5.
Participants	A total of 1692 participants (intervention = 848, control = 844) at baseline and 1699 participants at 18 months (intervention = 849, control = 850) Households were eligible if they have a child < 5 years of age and a guardian agreed to participate.
Interventions	SHEWA-B programme targeting improved latrine coverage and usage, access to and use of arsenic-free water, and improved hygiene practices using soaps. See Table 1 for details.
Outcomes	Laboratory: none described in methods and none reported Effectiveness: ARI and diarrhoea. ARI defined as cough and fever or difficulty breathing and fever within 48 h prior to interview. Safety: none described in methods and none reported
Notes	The authors conclude that “The prevalence of childhood diarrhea and respiratory illness was similar in the intervention and control communities”. Poorly reported trial

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Mentions random-number tables, but not clear if this was for random selection or randomisation
Allocation concealment (selection bias)	Unclear risk	Method not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data on illness were collected by a resident of the village, who was likely to know treatment allocation.
Incomplete outcome data (attrition bias) All outcomes	High risk	Not reported. No flow diagram
Selective reporting (reporting bias)	Unclear risk	Unlikely that other outcomes were measured and not reported

Huda 2012
Study characteristics

Huda 2012 (Continued)

Methods	Poorly described cluster-RCT. Partial report of the SHEWA-B trial focused on changing 11 targeted behaviours in villages to measure the impact on diarrhoea and respiratory illness amongst children. Unit of randomisation is not clear, but was probably a village. A group of 10 to 17 households within a village were the participants, based on the household having at least 1 child under the age of 5.
Participants	A total of 1692 participants (intervention = 848, control = 844) at baseline and 1699 participants at 18 months (intervention = 849, control = 850) Households were eligible if they have a child < 5 years of age and a guardian agreed to participate.
Interventions	SHEWA-B programme targeting improved latrine coverage and usage, access to and use of arsenic-free water, and improved hygiene practices using soaps. See Table 1 for details.
Outcomes	Laboratory: none described in methods and none reported Effectiveness: ARI and diarrhoea. ARI defined as cough and fever or difficulty breathing and fever within 48 h prior to interview. Safety: none described in methods and none reported
Notes	The authors conclude that “The prevalence of childhood diarrhea and respiratory illness was similar in the intervention and control communities”. Poorly reported trial

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Mentions random-number tables, but not clear if this was for random selection or randomisation
Allocation concealment (selection bias)	Unclear risk	Method not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data on illness were collected by a resident of the village, who was likely to know treatment allocation.
Incomplete outcome data (attrition bias) All outcomes	High risk	Not reported. No flow diagram
Selective reporting (reporting bias)	Unclear risk	Unlikely that other outcomes were measured and not reported

Ibfelt 2015
Study characteristics

Methods	Cluster-RCT in 12 daycare nurseries in Denmark. Centres in the intervention group had their linen and children's toys commercially cleaned and disinfected every 2 weeks. Control group centres had usual
---------	---

ibfelt 2015 (Continued)

practice. Swabbing for bacteria and respiratory viruses was conducted at baseline and the end of the intervention period.

Participants	<p>12 nurseries in Copenhagen (intervention = 6, control = 6) with a total of 587 children aged 6 months to 3 years</p> <p>Not clear how many children were in each group. Data on illness collected at the individual level, and on presence of bacteria and viruses at the cluster level.</p>
Interventions	<p>Washing and disinfection of toys and linen every 2 weeks for 3 months. See Table 1 for details.</p>
Outcomes	<p>Laboratory: counts of bacteria (not relevant to this review) and 11 respiratory viruses at baseline and end of intervention period, taken from swabs of 10 predefined locations in playroom (7 locations) and toilet area (3 locations). Viruses were influenza A and B; coronavirus NL63229E, OC43, and HKU1; parainfluenza virus 1, 2, 3, and 4; rhinovirus; RSV A/B; adenovirus; enterovirus; parechovirus; metapneumovirus; and bocavirus. Testing by PCR</p> <p>Effectiveness: illness counts in the children. Absence due to sickness recorded daily with reason categorised, but no definitions of illness provided.</p> <p>Safety: none mentioned in methods and none reported</p>
Notes	<p>The authors conclude that “Although cleaning and disinfection of toys every two weeks can decrease the microbial load in nurseries, it does not appear to reduce sickness absence among nursery children”.</p> <p>The results of the disinfection are reported as follows: “The most prevalent virus was coronavirus (97% positive samples), followed by bocavirus (96%), adenovirus (73%) and rhinovirus (46%). The intervention reduced the presence of adenovirus, rhinovirus and RSV approximately two- to five-fold [odds ratio (OR) 2.4, 95% confidence interval (CI) 1.1-5.0 for adenovirus; OR 5.3, 95% CI 2.3-12.4 for rhinovirus; OR 4.1, 95% CI 1.5-11.2 for RSV] compared with the control group. On the other hand, metapneumovirus was found significantly less often in the control group than in the intervention group. The intervention had no effect on the detection of other viruses. The fomites with the highest presence of respiratory virus were pillows and sofas, followed by toys and playroom tables. When looking at the samples from the toys alone, there was a significant decrease following the intervention in the intervention group compared with the control group for rhinovirus (OR 3.8, 95% CI 1.3-10.5; P = 0.01) and RSV (OR 5.2, 95% CI 1.1-23.8; P = 0.04), but not adenovirus”.</p> <p>This a poorly reported cluster-RCT. Its importance lies in the surface viral prevalence data (which could have been overestimated by PCR) and the finding that even in the presence of high viral prevalence, sickness was lower in the control (no surface disinfection) arm. This suggests the absence of other factors that could activate surface respiratory viruses.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Method not mentioned
Allocation concealment (selection bias)	Unclear risk	Method not mentioned
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias)	Low risk	Objective measure of bacterial and viral counts. However, illness reporting is unclear.

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

ibfelt 2015 (Continued)

All outcomes

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	No attrition or denominators given for results.
Selective reporting (reporting bias)	Low risk	Unlikely that other outcomes were measured but not reported

ibfelt 2015
Study characteristics

Methods	Cluster-RCT in 12 daycare nurseries in Denmark. Centres in the intervention group had their linen and children's toys commercially cleaned and disinfected every 2 weeks. Control group centres had usual practice. Swabbing for bacteria and respiratory viruses was conducted at baseline and the end of the intervention period.
Participants	12 nurseries in Copenhagen (intervention = 6, control = 6) with a total of 587 children aged 6 months to 3 years Not clear how many children were in each group. Data on illness collected at the individual level, and on presence of bacteria and viruses at the cluster level.
Interventions	Washing and disinfection of toys and linen every 2 weeks for 3 months. See Table 1 for details.
Outcomes	Laboratory: counts of bacteria (not relevant to this review) and 11 respiratory viruses at baseline and end of intervention period, taken from swabs of 10 predefined locations in playroom (7 locations) and toilet area (3 locations). Viruses were influenza A and B; coronavirus NL63229E, OC43, and HKU1; parainfluenza virus 1, 2, 3, and 4; rhinovirus; RSV A/B; adenovirus; enterovirus; parechovirus; metapneumovirus; and bocavirus. Testing by PCR Effectiveness: illness counts in the children. Absence due to sickness recorded daily with reason categorised, but no definitions of illness provided. Safety: none mentioned in methods and none reported
Notes	The authors conclude that "Although cleaning and disinfection of toys every two weeks can decrease the microbial load in nurseries, it does not appear to reduce sickness absence among nursery children". The results of the disinfection are reported as follows: "The most prevalent virus was coronavirus (97% positive samples), followed by bocavirus (96%), adenovirus (73%) and rhinovirus (46%). The intervention reduced the presence of adenovirus, rhinovirus and RSV approximately two- to five-fold [odds ratio (OR) 2.4, 95% confidence interval (CI) 1.1-5.0 for adenovirus; OR 5.3, 95% CI 2.3-12.4 for rhinovirus; OR 4.1, 95% CI 1.5-11.2 for RSV] compared with the control group. On the other hand, metapneumovirus was found significantly less often in the control group than in the intervention group. The intervention had no effect on the detection of other viruses. The fomites with the highest presence of respiratory virus were pillows and sofas, followed by toys and playroom tables. When looking at the samples from the toys alone, there was a significant decrease following the intervention in the intervention group compared with the control group for rhinovirus (OR 3.8, 95% CI 1.3-10.5; P = 0.01) and RSV (OR 5.2, 95% CI 1.1-23.8; P = 0.04), but not adenovirus". This a poorly reported cluster-RCT. Its importance lies in the surface viral prevalence data (which could have been overestimated by PCR) and the finding that even in the presence of high viral prevalence, sickness was lower in the control (no surface disinfection) arm. This suggests the absence of other factors that could activate surface respiratory viruses.

ibfelt 2015 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Method not mentioned
Allocation concealment (selection bias)	Unclear risk	Method not mentioned
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Objective measure of bacterial and viral counts. However, illness reporting is unclear.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	No attrition or denominators given for results.
Selective reporting (reporting bias)	Low risk	Unlikely that other outcomes were measured but not reported

Ide 2014
Study characteristics

Methods	Randomised, open-label, 2-group parallel study of 757 high school students (15 to 17 years of age) conducted for 90 days during the influenza epidemic season from 1 December 2011 to 28 February 2012, in 6 high schools in Shizuoka Prefecture, Japan. The green tea gargling group gargled 3 times a day with bottled green tea, and the water gargling group did the same with tap water. The water group was restricted from gargling with green tea.
Participants	A total of 747 students were enrolled (green tea gargling group = 384, water gargling group = 363) High school students (15 to 17 years of age) who attended 6 high schools in the Kakegawa and Ogasa districts of Shizuoka Prefecture, Japan
Interventions	See TIDieR Table (Table 1).
Outcomes	Incidence of laboratory-confirmed influenza Incidence of clinically defined influenza infection Time for which the participant was free from clinically defined influenza infection Clinically defined influenza infection, specified as fever ($\geq 37.8^\circ\text{C}$) plus any 2 of the following additional symptoms: cough, sore throat, headache, or myalgia. Influenza infection with viral antigen was detected by immunochromatographic assay. No safety data reported.
Notes	

Ide 2014 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated permuted block randomised schema
Allocation concealment (selection bias)	Low risk	Randomised at the Data Management Center of Shizuoka General Hospital in Japan
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Minimal attrition
Selective reporting (reporting bias)	Unclear risk	Protocol not available

Ide 2014
Study characteristics

Methods	Randomised, open-label, 2-group parallel study of 757 high school students (15 to 17 years of age) conducted for 90 days during the influenza epidemic season from 1 December 2011 to 28 February 2012, in 6 high schools in Shizuoka Prefecture, Japan. The green tea gargling group gargled 3 times a day with bottled green tea, and the water gargling group did the same with tap water. The water group was restricted from gargling with green tea.
Participants	A total of 747 students were enrolled (green tea gargling group = 384, water gargling group = 363) High school students (15 to 17 years of age) who attended 6 high schools in the Kakegawa and Ogasa districts of Shizuoka Prefecture, Japan
Interventions	See TIDieR Table (Table 1).
Outcomes	Incidence of laboratory-confirmed influenza Incidence of clinically defined influenza infection Time for which the participant was free from clinically defined influenza infection Clinically defined influenza infection, specified as fever ($\geq 37.8^\circ\text{C}$) plus any 2 of the following additional symptoms: cough, sore throat, headache, or myalgia. Influenza infection with viral antigen was detected by immunochromatographic assay. No safety data reported.
Notes	

Idc 2014 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated permuted block randomised schema
Allocation concealment (selection bias)	Low risk	Randomised at the Data Management Center of Shizuoka General Hospital in Japan
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	Minimal attrition
Selective reporting (reporting bias)	Unclear risk	Protocol not available

Idc 2016
Study characteristics

Methods	Randomised controlled study in Japan. Participants were randomly allocated into the catechin-treated (epigallocatechin gallate-treated) or non-treated face mask groups for 60 days from January to March 2016. Incidence of laboratory-confirmed influenza infection was measured and compared between groups using Fisher's exact test. Multivariate analysis was performed to calculate adjusted ORs and associated 95% CIs.
Participants	Participants included workers in a nursing home, a rehabilitation facility, and a hospital. A total of 234 participants were eligible for the study (catechin group, n = 118; control group, n = 116).
Interventions	Catechin-treated mask versus non-treated face mask. See Table 1 for details.
Outcomes	Incidence of laboratory-confirmed influenza infection Laboratory-confirmed influenza infection with viral antigen detected by immunochromatographic assay performed when participants reported ILI. No safety outcomes reported.
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Idc 2016 (Continued)

Random sequence generation (selection bias)	Unclear risk	Computer-generated randomisation, but method not stated
Allocation concealment (selection bias)	Low risk	Central randomisation service at Data Management Centre of Shizouka General Hospital
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Double-blinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Attrition minimal
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

Idc 2016
Study characteristics

Methods	Randomised controlled study in Japan. Participants were randomly allocated into the catechin-treated (epigallocatechin gallate-treated) or non-treated face mask groups for 60 days from January to March 2016. Incidence of laboratory-confirmed influenza infection was measured and compared between groups using Fisher's exact test. Multivariate analysis was performed to calculate adjusted ORs and associated 95% CIs.
Participants	Participants included workers in a nursing home, a rehabilitation facility, and a hospital. A total of 234 participants were eligible for the study (catechin group, n = 118; control group, n = 116).
Interventions	Catechin-treated mask versus non-treated face mask. See Table 1 for details.
Outcomes	Incidence of laboratory-confirmed influenza infection Laboratory-confirmed influenza infection with viral antigen detected by immunochromatographic assay performed when participants reported ILI. No safety outcomes reported.
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Computer-generated randomisation, but method not stated
Allocation concealment (selection bias)	Low risk	Central randomisation service at Data Management Centre of Shizouka General Hospital

Ide 2016 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	Low risk	Double-blinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Attrition minimal
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition minimal
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

Jacobs 2009
Study characteristics

Methods	Open RCT lasting 77 days from January 2008 to test “superiority” of face masks in preventing "URTI". This term appears as an acronym in the introduction and is not explained. It is assumed that it stands for 'upper respiratory infections', but it is preceded in the text by the term 'common cold', which is also lacking a definition. Randomisation was carried out in blocks within each of 3 professional figures (physicians, nurses, and “co-medical” personnel).
Participants	33 HCWs mainly females aged around 34 to 37 in a tertiary healthcare hospital in Tokyo, Japan. HCW with “predisposing conditions” (undefined) to “URTI” and those taking antibiotics were excluded. A baseline descriptive survey was carried out including “quality of life”. 1 participant dropped out at end of week 1, but no reason is reported nor the allocation arm. Analysis was performed on 32 participants (mask = 17, no mask = 15).
Interventions	Surgical mask MA-3 (Osu Sangyo, Japan) during all phases of hospital work (n = 17) or no mask (n = 15) (except when specifically required by hospital SOPs)
Outcomes	Laboratory: N/A Effectiveness: URTI is defined on the basis of a symptoms score, with a score > 14 being a URTI according to Jackson’s 1958 criteria (“Jackson score”). These are not explained in text, although the symptoms are listed in Table 3 (any, sore throat, runny nose, stuffy nose, sneeze, cough, headache, ear ache, feel bad) together with their mean and scores SD by intervention arm. Safety: the text does not mention or report harms. These appear to be indistinguishable from URTI symptoms (e.g. headache which is reported as of significantly longer duration in the intervention arm). Compliance is self-reported as high (84.3% of participants).
Notes	The authors conclude that “Face mask use in healthcare workers has not been demonstrated to provide benefit in terms of cold symptoms or getting colds. A larger study is needed to definitively establish non-inferiority of no mask use”. This is a small, badly reported trial. The purpose of trials is to test hypotheses not to prove or disprove 'superiority' of interventions. There is no power calculation, and CIs are not reported (although there is a mention in Discussion). No accurate definitions of a series of important variables (e.g. URTI, runny nose, etc.) are reported, and the Jackson scores are not explained, nor their use in Japanese personnel or language validated.

Jacobs 2009 (Continued)

Intervention arm data not extracted due to the uncertainty of its meaning.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Open RCT, but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	"Mask and no mask groups were formed using block randomisation of subjects within their respective job categories: nurses, doctors, and co-medical personnel." Concealment of allocation not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study. Blinding not possible, as 1 group wore face masks
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	1 dropout in each group accounted for. "Analyses were performed following the principles of intention-to-treat."
Selective reporting (reporting bias)	High risk	NB: influenza vaccine coverage was 100% in mask group and only 81% in the non-mask-wearing group.

Kotch 1994
Study characteristics

Methods	Pair-matched, cluster-RCT conducted from 19 October 1988 to 23 May 1989 in 24 childcare centres in North Carolina, USA The trial tested the effects of a hand-washing and environment sterilising programme on diarrhoea (data not extracted) and ARIs. Child daycare centres had to care for 30 children or less, at least 5 of whom had to be in nappies, and intending to stay open for at least another 2 years. Randomisation is not described, nor are cluster coefficients reported.
Participants	389 children aged 3 years or less in daycare for at least 20 hours a week. There were some withdrawals, but attrition of participants is not stated, only that in the end data for 31 intervention classrooms and 36 control classrooms were available. 291 children aged up to 24 months and 80 over 24 months took part. The text is very confusing, as 371 seems to be the total of the number of families that took part. No denominator breakdown by arm is reported, and numerators are only reported as new episodes per child-year.
Interventions	Structured hand-washing and environment (including surfaces, sinks, toilets, and toys) disinfecting programme with waterless disinfectant scrub
Outcomes	Laboratory: N/A Effectiveness: ARI (coughing, runny nose, wheezing, sore throat, or earache) Safety: N/A
Notes	Risk of bias: high (poor reporting of randomisation, outcomes, numerators and denominators)

Kotch 1994 (Continued)

Note: the authors conclude that the fully adjusted RR for prevention of ARIs was 0.94 (–2.43 to 0.66). A poorly reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Pair-matched cluster-randomised, controlled trial", but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	Centres were matched in pairs and then randomly allocated to either intervention or control programmes. Allocation concealment was not reported.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible (intervention was training session)
Blinding of outcome assessment (detection bias) All outcomes	High risk	"The same staff who conducted the training unobtrusively recorded observations at 5-week intervals"
Incomplete outcome data (attrition bias) All outcomes	High risk	18 families were dropped, denominator not clear.
Selective reporting (reporting bias)	High risk	Denominators not clearly reported

Ladegaard 1999
Study characteristics

Methods	RCT with cluster-randomisation to intervention or control. Of 10 institutions, 2 were excluded because they wanted institutions to be comparable in uptake area (i.e. housing and income). Interventions were administered to children, parents, and teachers at the institutions.
Participants	Children 0 to 6 years old
Interventions	Multifaceted: information, t-shirts to the children with: "Clean hands - yes, thank you", performance of a fairytale "The princess who did not want to wash her hands", exercise in hand-washing, importance of clean and fresh air. The aims of the intervention were to: <ol style="list-style-type: none"> 1. increase the hygiene education of the daycare teachers; 2. motivate the children by practical learning to have better hand hygiene; and 3. inform the parents about better hand hygiene.
Outcomes	34% decrease in "sickness" (probably mostly gastroenteritis)
Notes	Risk of bias: only limited data available Note: the authors conclude that there was a 34% decrease in sickness in the intervention arm; this is probably overall sickness, as gastroenteritis is part of the outcomes (data not extracted). Only limited data available from translation by Jørgen Lous.

Risk of bias

Ladegaard 1999 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not described
Allocation concealment (selection bias)	Unclear risk	Randomisation by "lottery", the same as "flip the coin". Concealment not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible
Blinding of outcome assessment (detection bias) All outcomes	High risk	Not possible
Incomplete outcome data (attrition bias) All outcomes	High risk	Total numbers of children included in each arm not reported.
Selective reporting (reporting bias)	High risk	Limited data reported, in particular denominators missing.

Larson 2010
Study characteristics

Methods	<p>Cluster block-randomised, controlled trial carried out between 20 November 2006 and 20 June 2008 in an upper Manhattan immigrant Latino neighbourhood ("19 month data collection period"). The study aimed at assessing the effects of education versus education and hand sanitiser use versus education and hand sanitiser use and common mask use against upper respiratory infections over a period of under 2 years. Follow-up was through an automated telephone system with a small financial incentive (USD 20) for those with 75% or more compliance. Those reporting an ILI received a visit within 48 hours for swabbing.</p> <p>An index case was someone who at the "onset day of illness nobody else in the household had been symptomatic within the previous five days".</p> <p>A secondary case for each episode "was any member of the household who developed symptoms within five days following the index case"; "The secondary attack rate was defined as the number of secondary cases recorded within 5 days of the onset of symptoms in the index case divided by the number of household members minus one".</p> <p>The text implies that the unit of observation was the episode ("study subjects contributed more than one episode in which they were considered to be the index case").</p>
Participants	<p>617 households were randomised to the education group (n = 211), the hand sanitiser group (n = 205), and the hand sanitiser and mask group (n = 201). There were 2708 participants, mostly adult Latino immigrants to the USA.</p> <p>Recruitment and allocation were carried out by household. There had to be at least 3 people living in the household, with at least 1 being a preschool or elementary school child, speaking English or Spanish, having a telephone, willingness to complete symptom assessments and have bimonthly home visits, and not using alcohol-based hand sanitiser routinely.</p> <p>Intracluster correlation coefficients are reported on page 179 of the manuscript.</p>

Larson 2010 (Continued)

Interventions

Written Spanish or English language educational materials regarding the prevention and treatment of URTIs and influenza or the same educational materials and hand sanitiser (Purell, J&J), in large (8- and 4-ounce) and small (1-ounce) containers to be carried by individual household members to work or school, or the same interventions as well as regular surgical face masks (Procedure Face Masks for adults and children, Kimberly-Clark) with instructions for both the caretaker and the ill person to wear them when an ILI occurred in any household member. Replenishment of intervention stocks was done at the bimonthly home visit.

Caretakers had to wear a mask for 7 days when within 3 feet of a symptomatic case. They were also encouraged to wear masks within 3 feet of any household member. Reinforcing phone calls were made 3 times in 6 days.

The text clearly reports active influenza vaccine promotion during the bimonthly visits. (“The home visit to each household was made every 2 months to minimise study dropout, reinforce adherence to the assigned intervention, replenish product supplies and record use of supplies, answer questions, and correct ongoing misconceptions. At each visit, new educational materials regarding URTI prevention and treatment and influenza vaccination were distributed.” (PDF page 3). Also just before the Discussion as follows: “Influenza vaccination rates: There was an increase between the baseline and exit interview in all three groups that reported 50% or more of members receiving influenza vaccine (pre- versus post-intervention for each group: 21.1% and 40.8% in the Education group, 19.0% and 57.1% in the hand sanitiser group, and 22.4% and 43.5% in the hand sanitiser and face mask group (P = 0.001). Additionally, those in the hand sanitiser group reported a significantly greater increase than the other 2 groups, controlling for baseline rates (P = 0.002)”)

Coverage was unequal across groups, no information on the progressive impact of the vaccine, or indeed the nature of the vaccine(s) is reported. Apparently the first season was mild and the vaccine mismatched, compliance with the trial interventions was low in Arm 3, and a local epidemic of *Staphylococcus aureus* meant that the control group started washing hands.

The trial authors report no effect on reporting rates of vaccine coverage by arms, but with so many confounders who knows?

Outcomes

Laboratory: PCR carried out on samples from deep nasal swabs for influenza and the most common other pathogens (RSV, rhinovirus, enterovirus, parainfluenza viruses, etc.). The text describing the results of the swabbing is confusing, but in general appears to be non-random “Households reported 669 episodes of ILI (0 to 5 per individual)”. Of the 234 deep nasal swabs obtained, 33.3% (n = 78) tested positive for influenza: 43.6% (n = 34) were influenza A and 56.4% (n = 44) were influenza B. Amongst the 66.7% who tested negative for influenza, 30.8% (48/156) tested positive for other viruses: 7 for respiratory syncytial virus, 9 for parainfluenza, 11 for enterovirus, 10 for rhinovirus, 6 for adenovirus, and 5 for metapneumovirus. Swabs were not obtained from the remaining 435 reported ILI episodes for the following reasons: 72.0% (n = 313) did not meet the CDC definition of an ILI and were therefore included in the URTI symptom count; 21.4% of episodes (n = 93) were reported after 48 hours of ILI onset or the participant refused to be swabbed; and the research staff were unable to reach the participant in 6.7% of episodes (n = 29).

As no definition of URTI is given, it is unclear what kind of biases were introduced by the non-swabbing of the 313/435 “not meeting CDC definition”.

Effectiveness: ILI (CDC definition): “temperature of 37.8°C or more and cough and/or sore throat in the absence of a known cause other than influenza”

URTI only referred to as “Viral upper respiratory infections (URTIs)”.

Safety: N/A

Notes

The authors conclude that “the Hand Sanitizer group was significantly more likely to report that no household member had symptoms (P,0.01), but there were no significant differences in rates of infection by intervention group in multivariate analyses. Knowledge improved significantly more in the Hand Sanitizer group (P,0.0001). The proportion of households that reported >50% of members receiving influenza vaccine increased during the study (P,0.001). Despite the fact that compliance with mask wearing was poor, mask wearing as well as increased crowding, lower education levels of caretakers, and index cases 0–5 years of age (compared with adults) were associated with significantly lower sec-

Larson 2010 (Continued)

ondary transmission rates (all $P < 0.02$). In this population, there was no detectable additional benefit of hand sanitiser or face masks over targeted education on overall rates of URTIs, but mask wearing was associated with reduced secondary transmission and should be encouraged during outbreak situations. During the study period, community concern about methicillin-resistant *Staphylococcus aureus* was occurring, perhaps contributing to the use of hand sanitiser in the Education control group, and diluting the intervention's measurable impact".

The study is at high risk of bias. Randomisation and reasons for dropout are not described. Differentials in cluster characteristics across arms point to randomisation not having worked, and the confounding effects of a postrandomisation staphylococcal scare are difficult to judge. Symptom-driven follow-up gives no idea of the effects on asymptomatic ILI/influenza. Poor definitions (URTI?). There are unexplained dropouts, and the analysis plan is unclear. Finally, the very small number of cases of influenza and an unclear swabbing attrition may introduce further elements of confounding.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Cluster block randomised, controlled trial", but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	"Households were block randomised into one of three groups" Allocation concealment not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	Blinding of participants and personnel was not possible.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Blinding of outcome assessment is not stated.
Incomplete outcome data (attrition bias) All outcomes	High risk	In control group households (n = 211), 26 dropped out and 37 did not consent. In hand sanitiser group households (n = 205), 21 dropped out and 36 did not consent. In hand sanitiser and face mask group households (n = 201), 19 dropped out and 35 did not consent. Reasons for dropout were not described.
Selective reporting (reporting bias)	Unclear risk	617 of 772 eligible households were randomised.

Little 2015
Study characteristics

Methods	Individuals sharing a household by mailed invitation through general practices in England were recruited. After consent, participants were randomised online by an automated computer-generated random-number program to receive either no access or access to a bespoke automated web-based intervention that maximised hand-washing intention, monitored hand-washing behaviour, provided tailored feedback, reinforced helpful attitudes and norms, and addressed negative beliefs. Participants were enrolled into an additional cohort (randomised to receive intervention or no intervention) to assess whether the baseline questionnaire on hand-washing would affect hand-washing behavior.
---------	--

Little 2015 (Continued)

our. Participants were not masked to intervention allocation, but statistical analysis commands were constructed masked to group. The primary outcome was number of episodes of RTIs in index participants in a modified intention-to-treat population of randomly assigned participants who completed follow-up at 16 weeks.

Participants	<p>344 physician offices were recruited over a wide area of England, and 20,066 participants were enrolled and randomised to intervention (N = 16,086) and control (N = 10,026).</p> <p>Modified ITT was performed on 16,908 participants who completed the follow-up questionnaire at 16 weeks (intervention = 8241 and control = 8667).</p> <p>Inclusion criteria: adult patients (aged 18 years or older) identified from computerised lists in general practitioner (GP) practices in England, for whom there was at least 1 other individual living in the household who was willing to report illness to the index person</p> <p>Exclusion criteria: patients with severe mental problems (e.g. major uncontrolled depression or schizophrenia, dementia, or severe mental impairment) or who were terminally ill, and those reporting a skin complaint that would restrict hand-washing</p>
Interventions	<p>Automated web-based intervention that maximised hand-washing intention, monitored hand-washing behaviour, provided tailored feedback, reinforced helpful attitudes and norms, and addressed negative beliefs. Control no access to intervention web pages. See Table 1 for details.</p>
Outcomes	<p>The primary outcome was the number of index individuals that reported 1 or more RTIs (including ILI) at 16 weeks.</p> <p>Secondary: duration of symptoms, transmission of respiratory infections, gastrointestinal infections, attendance at the practice, and use of health service resources</p> <p>Infections self-reported by participants. RTI defined as 2 symptoms of an RTI for at least 1 day or 1 symptom for 2 consecutive days. Definition of ILI was a high temperature (feeling very hot or very cold; or measured temperature > 37.5 °C), a respiratory symptom (sore throat, cough, or runny nose), and a systemic symptom (headache, severe fatigue, severe muscle aches, or severe malaise).</p> <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Participants were automatically randomly assigned by the intervention software, but sequence generation not described.
Allocation concealment (selection bias)	Low risk	Participants were automatically randomly assigned by the intervention software.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	High attrition that was different in the 2 groups

Little 2015 (Continued)

Selective reporting (reporting bias)	Low risk	Specified outcomes reported.
--------------------------------------	----------	------------------------------

Jacobs 2009
Study characteristics

Methods	Open RCT lasting 77 days from January 2008 to test “superiority” of face masks in preventing "URTI". This term appears as an acronym in the introduction and is not explained. It is assumed that it stands for 'upper respiratory infections', but it is preceded in the text by the term 'common cold', which is also lacking a definition. Randomisation was carried out in blocks within each of 3 professional figures (physicians, nurses, and “co-medical” personnel).	
Participants	33 HCWs mainly females aged around 34 to 37 in a tertiary healthcare hospital in Tokyo, Japan. HCW with “predisposing conditions” (undefined) to “URTI” and those taking antibiotics were excluded. A baseline descriptive survey was carried out including “quality of life”. 1 participant dropped out at end of week 1, but no reason is reported nor the allocation arm. Analysis was performed on 32 participants (mask = 17, no mask = 15).	
Interventions	Surgical mask MA-3 (Osu Sangyo, Japan) during all phases of hospital work (n = 17) or no mask (n = 15) (except when specifically required by hospital SOPs)	
Outcomes	Laboratory: N/A Effectiveness: URTI is defined on the basis of a symptoms score, with a score > 14 being a URTI according to Jackson’s 1958 criteria (“Jackson score”). These are not explained in text, although the symptoms are listed in Table 3 (any, sore throat, runny nose, stuffy nose, sneeze, cough, headache, ear ache, feel bad) together with their mean and scores SD by intervention arm. Safety: the text does not mention or report harms. These appear to be indistinguishable from URTI symptoms (e.g. headache which is reported as of significantly longer duration in the intervention arm). Compliance is self-reported as high (84.3% of participants).	
Notes	The authors conclude that “Face mask use in healthcare workers has not been demonstrated to provide benefit in terms of cold symptoms or getting colds. A larger study is needed to definitively establish non-inferiority of no mask use”. This is a small, badly reported trial. The purpose of trials is to test hypotheses not to prove or disprove 'superiority' of interventions. There is no power calculation, and CIs are not reported (although there is a mention in Discussion). No accurate definitions of a series of important variables (e.g. URTI, runny nose, etc.) are reported, and the Jackson scores are not explained, nor their use in Japanese personnel or language validated. Intervention arm data not extracted due to the uncertainty of its meaning.	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Open RCT, but sequence generation not reported

Jacobs 2009 (Continued)

Allocation concealment (selection bias)	Unclear risk	"Mask and no mask groups were formed using block randomisation of subjects within their respective job categories: nurses, doctors, and co-medical personnel." Concealment of allocation not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study. Blinding not possible, as 1 group wore face masks
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	1 dropout in each group accounted for. "Analyses were performed following the principles of intention-to-treat."
Selective reporting (reporting bias)	High risk	NB: influenza vaccine coverage was 100% in mask group and only 81% in the non-mask-wearing group.

Kotch 1994
Study characteristics

Methods	Pair-matched, cluster-RCT conducted from 19 October 1988 to 23 May 1989 in 24 childcare centres in North Carolina, USA The trial tested the effects of a hand-washing and environment sterilising programme on diarrhoea (data not extracted) and ARIs. Child daycare centres had to care for 30 children or less, at least 5 of whom had to be in nappies, and intending to stay open for at least another 2 years. Randomisation is not described, nor are cluster coefficients reported.
Participants	389 children aged 3 years or less in daycare for at least 20 hours a week. There were some withdrawals, but attrition of participants is not stated, only that in the end data for 31 intervention classrooms and 36 control classrooms were available. 291 children aged up to 24 months and 80 over 24 months took part. The text is very confusing, as 371 seems to be the total of the number of families that took part. No denominator breakdown by arm is reported, and numerators are only reported as new episodes per child-year.
Interventions	Structured hand-washing and environment (including surfaces, sinks, toilets, and toys) disinfecting programme with waterless disinfectant scrub
Outcomes	Laboratory: N/A Effectiveness: ARI (coughing, runny nose, wheezing, sore throat, or earache) Safety: N/A
Notes	Risk of bias: high (poor reporting of randomisation, outcomes, numerators and denominators) Note: the authors conclude that the fully adjusted RR for prevention of ARIs was 0.94 (-2.43 to 0.66). A poorly reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Pair-matched cluster-randomised, controlled trial", but sequence generation not reported

Kotch 1994 (Continued)

Allocation concealment (selection bias)	Unclear risk	Centres were matched in pairs and then randomly allocated to either intervention or control programmes. Allocation concealment was not reported.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible (intervention was training session)
Blinding of outcome assessment (detection bias) All outcomes	High risk	"The same staff who conducted the training unobtrusively recorded observations at 5-week intervals"
Incomplete outcome data (attrition bias) All outcomes	High risk	18 families were dropped, denominator not clear.
Selective reporting (reporting bias)	High risk	Denominators not clearly reported

Ladegaard 1999
Study characteristics

Methods	RCT with cluster-randomisation to intervention or control. Of 10 institutions, 2 were excluded because they wanted institutions to be comparable in uptake area (i.e. housing and income). Interventions were administered to children, parents, and teachers at the institutions.
Participants	Children 0 to 6 years old
Interventions	Multifaceted: information, t-shirts to the children with: "Clean hands - yes, thank you", performance of a fairytale "The princess who did not want to wash her hands", exercise in hand-washing, importance of clean and fresh air. The aims of the intervention were to: <ol style="list-style-type: none"> 1. increase the hygiene education of the daycare teachers; 2. motivate the children by practical learning to have better hand hygiene; and 3. inform the parents about better hand hygiene.
Outcomes	34% decrease in "sickness" (probably mostly gastroenteritis)
Notes	Risk of bias: only limited data available Note: the authors conclude that there was a 34% decrease in sickness in the intervention arm; this is probably overall sickness, as gastroenteritis is part of the outcomes (data not extracted). Only limited data available from translation by Jørgen Lous.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not described
Allocation concealment (selection bias)	Unclear risk	Randomisation by "lottery", the same as "flip the coin". Concealment not reported

Ladegaard 1999 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible
Blinding of outcome assessment (detection bias) All outcomes	High risk	Not possible
Incomplete outcome data (attrition bias) All outcomes	High risk	Total numbers of children included in each arm not reported.
Selective reporting (reporting bias)	High risk	Limited data reported, in particular denominators missing.

Larson 2010
Study characteristics

Methods	<p>Cluster block-randomised, controlled trial carried out between 20 November 2006 and 20 June 2008 in an upper Manhattan immigrant Latino neighbourhood (“19 month data collection period”). The study aimed at assessing the effects of education versus education and hand sanitiser use and common mask use against upper respiratory infections over a period of under 2 years. Follow-up was through an automated telephone system with a small financial incentive (USD 20) for those with 75% or more compliance. Those reporting an ILI received a visit within 48 hours for swabbing.</p> <p>An index case was someone who at the “onset day of illness nobody else in the household had been symptomatic within the previous five days”.</p> <p>A secondary case for each episode “was any member of the household who developed symptoms within five days following the index case”; “The secondary attack rate was defined as the number of secondary cases recorded within 5 days of the onset of symptoms in the index case divided by the number of household members minus one”.</p> <p>The text implies that the unit of observation was the episode (“study subjects contributed more than one episode in which they were considered to be the index case”).</p>
Participants	<p>617 households were randomised to the education group (n = 211), the hand sanitiser group (n = 205), and the hand sanitiser and mask group (n = 201). There were 2708 participants, mostly adult Latino immigrants to the USA.</p> <p>Recruitment and allocation were carried out by household. There had to be at least 3 people living in the household, with at least 1 being a preschool or elementary school child, speaking English or Spanish, having a telephone, willingness to complete symptom assessments and have bimonthly home visits, and not using alcohol-based hand sanitiser routinely.</p> <p>Intracluster correlation coefficients are reported on page 179 of the manuscript.</p>
Interventions	<p>Written Spanish or English language educational materials regarding the prevention and treatment of URIs and influenza or the same educational materials and hand sanitiser (Purell, J&J), in large (8- and 4-ounce) and small (1-ounce) containers to be carried by individual household members to work or school, or the same interventions as well as regular surgical face masks (Procedure Face Masks for adults and children, Kimberly-Clark) with instructions for both the caretaker and the ill person to wear them when an ILI occurred in any household member. Replenishment of intervention stocks was done at the bimonthly home visit.</p>

Larson 2010 (Continued)

Caretakers had to wear a mask for 7 days when within 3 feet of a symptomatic case. They were also encouraged to wear masks within 3 feet of any household member. Reinforcing phone calls were made 3 times in 6 days.

The text clearly reports active influenza vaccine promotion during the bimonthly visits. (“The home visit to each household was made every 2 months to minimise study dropout, reinforce adherence to the assigned intervention, replenish product supplies and record use of supplies, answer questions, and correct ongoing misconceptions. At each visit, new educational materials regarding URTI prevention and treatment and influenza vaccination were distributed.” (PDF page 3). Also just before the Discussion as follows: “Influenza vaccination rates: There was an increase between the baseline and exit interview in all three groups that reported 50% or more of members receiving influenza vaccine (pre- versus post-intervention for each group: 21.1% and 40.8% in the Education group, 19.0% and 57.1% in the hand sanitiser group, and 22.4% and 43.5% in the hand sanitiser and face mask group ($P = 0.001$). Additionally, those in the hand sanitiser group reported a significantly greater increase than the other 2 groups, controlling for baseline rates ($P = 0.002$)”)

Coverage was unequal across groups, no information on the progressive impact of the vaccine, or indeed the nature of the vaccine(s) is reported. Apparently the first season was mild and the vaccine mismatched, compliance with the trial interventions was low in Arm 3, and a local epidemic of *Staphylococcus aureus* meant that the control group started washing hands.

The trial authors report no effect on reporting rates of vaccine coverage by arms, but with so many confounders who knows?

Outcomes

Laboratory: PCR carried out on samples from deep nasal swabs for influenza and the most common other pathogens (RSV, rhinovirus, enterovirus, parainfluenza viruses, etc.). The text describing the results of the swabbing is confusing, but in general appears to be non-random “Households reported 669 episodes of ILI (0 to 5 per individual)”. Of the 234 deep nasal swabs obtained, 33.3% ($n = 78$) tested positive for influenza: 43.6% ($n = 34$) were influenza A and 56.4% ($n = 44$) were influenza B. Amongst the 66.7% who tested negative for influenza, 30.8% (48/156) tested positive for other viruses: 7 for respiratory syncytial virus, 9 for parainfluenza, 11 for enterovirus, 10 for rhinovirus, 6 for adenovirus, and 5 for metapneumovirus. Swabs were not obtained from the remaining 435 reported ILI episodes for the following reasons: 72.0% ($n = 313$) did not meet the CDC definition of an ILI and were therefore included in the URTI symptom count; 21.4% of episodes ($n = 93$) were reported after 48 hours of ILI onset or the participant refused to be swabbed; and the research staff were unable to reach the participant in 6.7% of episodes ($n = 29$).

As no definition of URTI is given, it is unclear what kind of biases were introduced by the non-swabbing of the 313/435 “not meeting CDC definition”.

Effectiveness: ILI (CDC definition): “temperature of 37.8°C or more and cough and/or sore throat in the absence of a known cause other than influenza”

URTI only referred to as “Viral upper respiratory infections (URTIs)”.

Safety: N/A

Notes

The authors conclude that “the Hand Sanitizer group was significantly more likely to report that no household member had symptoms ($P,0.01$), but there were no significant differences in rates of infection by intervention group in multivariate analyses. Knowledge improved significantly more in the Hand Sanitizer group ($P,0.0001$). The proportion of households that reported >50% of members receiving influenza vaccine increased during the study ($P,0.001$). Despite the fact that compliance with mask wearing was poor, mask wearing as well as increased crowding, lower education levels of caretakers, and index cases 0–5 years of age (compared with adults) were associated with significantly lower secondary transmission rates (all $P,0.02$). In this population, there was no detectable additional benefit of hand sanitiser or face masks over targeted education on overall rates of URTIs, but mask wearing was associated with reduced secondary transmission and should be encouraged during outbreak situations. During the study period, community concern about methicillin-resistant *Staphylococcus aureus* was occurring, perhaps contributing to the use of hand sanitiser in the Education control group, and diluting the intervention’s measurable impact”.

The study is at high risk of bias. Randomisation and reasons for dropout are not described. Differentials in cluster characteristics across arms point to randomisation not having worked, and the confounding

Larson 2010 (Continued)

effects of a postrandomisation staphylococcal scare are difficult to judge. Symptom-driven follow-up gives no idea of the effects on asymptomatic ILI/influenza. Poor definitions (URTI?). There are unexplained dropouts, and the analysis plan is unclear. Finally, the very small number of cases of influenza and an unclear swabbing attrition may introduce further elements of confounding.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Cluster block randomised, controlled trial", but sequence generation not reported
Allocation concealment (selection bias)	Unclear risk	"Households were block randomised into one of three groups" Allocation concealment not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	Blinding of participants and personnel was not possible.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Blinding of outcome assessment is not stated.
Incomplete outcome data (attrition bias) All outcomes	High risk	In control group households (n = 211), 26 dropped out and 37 did not consent. In hand sanitiser group households (n = 205), 21 dropped out and 36 did not consent. In hand sanitiser and face mask group households (n = 201), 19 dropped out and 35 did not consent. Reasons for dropout were not described.
Selective reporting (reporting bias)	Unclear risk	617 of 772 eligible households were randomised.

Little 2015
Study characteristics

Methods	Individuals sharing a household by mailed invitation through general practices in England were recruited. After consent, participants were randomised online by an automated computer-generated random-number program to receive either no access or access to a bespoke automated web-based intervention that maximised hand-washing intention, monitored hand-washing behaviour, provided tailored feedback, reinforced helpful attitudes and norms, and addressed negative beliefs. Participants were enrolled into an additional cohort (randomised to receive intervention or no intervention) to assess whether the baseline questionnaire on hand-washing would affect hand-washing behaviour. Participants were not masked to intervention allocation, but statistical analysis commands were constructed masked to group. The primary outcome was number of episodes of RTIs in index participants in a modified intention-to-treat population of randomly assigned participants who completed follow-up at 16 weeks.
Participants	344 physician offices were recruited over a wide area of England, and 20,066 participants were enrolled and randomised to intervention (N = 16,086) and control (N = 10,026).

Little 2015 (Continued)

Modified ITT was performed on 16,908 participants who completed the follow-up questionnaire at 16 weeks (intervention = 8241 and control = 8667).

Inclusion criteria: adult patients (aged 18 years or older) identified from computerised lists in general practitioner (GP) practices in England, for whom there was at least 1 other individual living in the household who was willing to report illness to the index person

Exclusion criteria: patients with severe mental problems (e.g. major uncontrolled depression or schizophrenia, dementia, or severe mental impairment) or who were terminally ill, and those reporting a skin complaint that would restrict hand-washing

Interventions	Automated web-based intervention that maximised hand-washing intention, monitored hand-washing behaviour, provided tailored feedback, reinforced helpful attitudes and norms, and addressed negative beliefs. Control no access to intervention web pages. See Table 1 for details.
Outcomes	<p>The primary outcome was the number of index individuals that reported 1 or more RTIs (including ILI) at 16 weeks.</p> <p>Secondary: duration of symptoms, transmission of respiratory infections, gastrointestinal infections, attendance at the practice, and use of health service resources</p> <p>Infections self-reported by participants. RTI defined as 2 symptoms of an RTI for at least 1 day or 1 symptom for 2 consecutive days. Definition of ILI was a high temperature (feeling very hot or very cold; or measured temperature > 37.5 °C), a respiratory symptom (sore throat, cough, or runny nose), and a systemic symptom (headache, severe fatigue, severe muscle aches, or severe malaise).</p> <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Participants were automatically randomly assigned by the intervention software, but sequence generation not described.
Allocation concealment (selection bias)	Low risk	Participants were automatically randomly assigned by the intervention software.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	High risk	High attrition that was different in the 2 groups
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

Loeb 2009

Study characteristics

Methods	<p>Open non-inferiority RCT carried out to compare the surgical mask with the N95 respirator in protecting healthcare workers against influenza. The trial was carried out between 2008 (enrolment started in September and follow-up on 12 January 2009) and 23 April 2009 (when all HCWs caring for febrile patients were told to wear an N95 respirator) because of the appearance of novel A/H1N1). The trial trigger was the beginning of the influenza season, defined as isolation of 2 or more viruses in a district in the same week. Following the 2003 SARS outbreak, all Ontario nurses caring for febrile patients (38 °C or more and new onset cough or SOB) had to wear surgical masks. The randomisation (carried out in blocks of 4 by centre) then consisted of either confirmation to same-maker surgical mask wear or N95 respirator wear. Investigators and laboratory staff were blind to allocation status, but for obvious reasons (the visible difference in interventions), participants were unblinded. "The criterion for non-inferiority was met if the lower limit of the 95% confidence interval (CI) for the reduction in incidence (N95 respirator minus surgical group) was greater than -9%". So this is the non-inferiority margin. It is assumed that the "minus surgical group" means minus surgical mask group.</p>
Participants	<p>Consenting nurses (n = 446 randomised) aged a mean of 36.2 years working full time (≥ 37 hours/week) in 23 acute units (a mix of paediatric, A&E, and acute medical units) in 8 hospitals in Ontario, Canada. 225 were randomised to the surgical mask and 221 to the N95 respirator. There were 13 and 11 dropouts, respectively from each arm (all accounted for), plus 21 and 19 lost to follow-up; 11 in each arm gave no reason, the others are accounted for. There were no deaths. The final total of 212 and 210 was included in the analysis. Table 1 reports the demographic data of participants by arm, which appear comparable.</p>
Interventions	<p>Surgical masks (as standard wear by the standard distributor) or fit-tested N95 respirator. All nurses wore gloves or gowns in the presence of a febrile patient.</p>
Outcomes	<p>Laboratory RT-PCR paired sera with 4-fold antibody rise from baseline (only for unvaccinated) nurses</p> <p>Effectiveness: follow-up (lasting a mean of around 97 days for both arms) was carried out twice-weekly on a web-based instrument. Nurses with new symptoms were asked to swab a nostril if any of the following signs or symptoms had developed: fever (temperature ≥ 38 °C), cough, nasal congestion, sore throat, headache, sinus problems, muscle aches, fatigue, earache, ear infection, or chills.</p> <p>The text defines influenza with laboratory confirmation, and separately reports criteria for swab triggering and a definition of ILI ("Influenza-like illness was defined as the presence of cough and fever: a temperature ≥ 38°C"). But this is not formally linked to influenza in the text, as it appears that primary focus was the detection of laboratory-confirmed influenza (either by RT-PCR or serology).</p> <p>Additional outcome data sought were work-related absenteeism and physician visits for respiratory illness.</p> <p>Secondary outcomes included detection of the following non-influenza viruses by PCR: parainfluenza virus types 1, 2, 3, and 4; respiratory syncytial virus types A and B; adenovirus; metapneumovirus; rhinovirus-enterovirus; and coronaviruses OC43, 229E, SARS, NL63, and HKU1.</p> <p>Audits to assess nurse compliance with the interventions were carried out in the room of each patient cared for. The text reports that 50 and 48 nurses in the surgical mask and N95 groups, respectively, had laboratory confirmation of influenza infection, indicating non-inferiority. Interestingly, non-inferiority seemed to be applicable both to seasonal viruses and nH1N1 viruses (as 8% and 11.9% were serologically positive to nH1N1). This finding is explained either by seeding or cross reaction with seasonal H1N1. Equivalent conclusions could be drawn for nurses with complete follow-up. Non-inferiority was applicable also to other ILI agents identified. None of the 52 individuals with positive isolates met the criteria for ILI.</p> <p>All cases of ILI were confirmed as having influenza (9 and 2 respectively). This means that all the 11 cases of ILI had influenza, but that most of those with a laboratory diagnosis of influenza did not have cough and fever. For example, the text reports that "Of the 44 nurses in each group who had influenza diagnosed by serology, 29 (65.9%) in the surgical mask group and 31 (70.5%) in the N95 respirator group had no symptoms". By implication, of the 88 nurses with antibody rises, 28 had symptoms of some kind, i.e. two-thirds were asymptomatic. Absenteeism was 1 versus 39 episodes in the mask ver-</p>

Loeb 2009 (Continued)

sus respirator arms. No episodes of LRTI were recorded. The number of family contacts with ILI were the same for each arm (45 versus 47). Physician visits were similar in both groups.

Safety: no AEs are reported

Notes

The authors conclude that "Among nurses in Ontario tertiary care hospitals, use of a surgical mask compared with a N95 respirator resulted in non-inferior rates of laboratory-confirmed influenza".

This a well-designed and conducted trial with credible conclusions. The only comment is that the focus in the analysis on influenza (symptomatic and asymptomatic) is not well-described, although the rationale is clear (interruption of transmission).

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Randomisation was performed centrally", but method of sequence generation not described.
Allocation concealment (selection bias)	Low risk	"...by an independent clinical trials coordinating group such that investigators were blind to the randomisation procedure and group assignment and was stratified by centre in permuted blocks of 4 participants."
Blinding of participants and personnel (performance bias) All outcomes	High risk	"It was not possible to conceal the identity of the N95 respirator or the surgical mask since manipulating these devices would interfere with their function"
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessment blinded: "Laboratory personnel conducting hemagglutinin inhibition assays, polymerase chain reaction (PCR), and viral culture for influenza were blinded to allocation."
Incomplete outcome data (attrition bias) All outcomes	Low risk	21 of 225 randomised to mask group and 19 of 221 randomised to N95 group were lost to follow-up, reasons reported. Study stopped early: "We had planned to stop the study at the end of influenza season. However, because of the 2009 influenza A(H1N1) pandemic, the study was stopped on April 23, 2009, when the Ontario Ministry of Health and Long-Term Care recommended N95 respirators for all healthcare workers taking care of patients with febrile respiratory illness."
Selective reporting (reporting bias)	Low risk	All outcomes reported.

Longini 1988
Study characteristics

Methods	Cluster-controlled, double-blind, randomised trial to assess the efficacy of virucidal tissues in interrupting family transmission of rhinovirus and influenza virus. The study was carried out in the community of Tecumseh, Michigan, USA during the period of 25 November 1984 to 28 April 1985. However, the authors only report results for the period of 13 January to 23 March 1985, when a high circulation of influenza A H3N2 and rhinovirus was detected.
Participants	296 households were enrolled, but 5 households were eliminated from the analysis for "technical reasons". The analysis was carried out in households with 3 to 5 members. The authors report data on 143 households randomised to virucidal tissues and 148 to placebo tissue. The average age in households

Longini 1988 (Continued)

was around 22, and the difference between arms was not significant. Randomisation was carried out by the sponsor, and tissues were pre-packed in coded boxes with no other identifying features and delivered to households at the beginning of the study period.

Interventions	Disposable 3-layered virucidal tissues (citric and malic acids with sodium lauryl sulphate in the middle layer) or placebo (succinic acid in the middle layer) tissues. They were used to blow the nose and for coughing or sneezing into. Households were also stratified by level of tissue use. Tissue use was significantly higher in the intervention arm (82% versus 71%).
Outcomes	Laboratory: yes - viral culture from nasal and throat swabs from symptomatic participants Effectiveness: ARI (with a proportion of laboratory-confirmed diagnosis in non-randomly chosen participants with symptoms lasting 2 days or more) Follow-up and surveillance was carried out using a telephone questionnaire. Safety: N/A
Notes	Risk of bias: high (inappropriate choice of placebo) Note: the authors conclude that virucidal tissues were up to 36.9% effective in preventing transmission of ARIs as measured by secondary attack rates (18.7% versus 11.8%). This finding was not statistically significant, but may well have been affected by the lack of do-nothing community controls. This a well-designed, well-written study despite the unexplained attrition of 5 families, the lack of reporting of cluster coefficients, and the differential in tissue use between the 2 arms, which raises questions about the robustness of double-blinding. Particularly notable is the discussion on the low generalisability of results from the study from the placebo arm given that even the inert barrier of the tissues is likely to have limited spread. Also, the lengths to which the authors went to obtain allocation concealment and maintenance of double-blind conditions

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Treated and placebo tissues were randomly assigned ..." Sequence generation not reported
Allocation concealment (selection bias)	Low risk	"Treated and placebo tissues were randomly assigned by the sponsor to 296 participating households stratified by household size, such that roughly half the households would receive treated tissues. Thus, the investigators were unaware of the assignment of treated tissues."
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Treated and placebo tissues were randomly assigned by the sponsor to the randomly assigned 296 households stratified by household size... The type of tissue was identified by code, and the boxes in which tissues were contained were not marked with any specific identifiers. Therefore, the study was double-blinded."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"The investigators were unaware of the assignment of the treated tissues"
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	296 households eligible. "The final sample used for analysis consisted of 143 households in the treatment group and 148 households in the placebo group."
Selective reporting (reporting bias)	High risk	"The analysis of secondary spread was restricted to households of three to five members for technical reasons, which eliminated five households." "The two groups were almost identical in composition."

Luby 2005

Study characteristics

Methods	<p>Partly double-blind, cluster-RCT carried out during 15 April 2002 to 5 April 2003 in Karachi, Pakistan. The trial assessed the effects of mother and child hand-washing on the incidence of respiratory infections, impetigo (data not extracted), and diarrhoea (data not extracted).</p> <p>Randomisation took place by computer-generated random numbers in 3 phases.</p> <ol style="list-style-type: none"> 25 neighbourhoods were assigned to hand-washing and 11 to standard practice. 300 households were assigned to using antiseptic soap. 300 households were assigned to using plain soap. 306 households were assigned to standard practice. 1523 children younger than 15 years were assigned to using antiseptic soap. 1640 children younger than 15 years were assigned to using plain soap. 1528 children younger than 15 years were assigned to standard practice. <p>Soaps were of identical weight, colour, and smell and were packed centrally with a coded packing case matched to households containing 96 bars. Neither fieldworkers nor participants were aware of the content. Control arm households were visited with the same frequency as intervention household but were given books and pens. Codes were held centrally by the manufacturer and broken after the end of the trial to allow analysis.</p>
Participants	<p>Householders of slums in Karachi.</p> <p>Of the 1523 children younger than 15 years assigned to using antiseptic soap, 117 dropped out (1 died, 51 were born in, and 65 aged out) = 1406; 504 were aged less than 5. Of 1640 children younger than 15 years assigned to using plain soap, 117 dropped out (3 died, 44 were born in, and 70 aged out) = 1523; 517 were aged less than 5. Of 1528 children younger than 15 years assigned to standard practice, 125 dropped out (3 died, 40 were born in, and 82 aged out) = 1403; 489 were aged less than 5.</p>
Interventions	<p>Instruction programme and antibacterial soap containing 1.2% triclocarban, or ordinary soap to be used throughout the day by householders, or standard procedure</p>
Outcomes	<p>Laboratory: N/A</p> <p>Effectiveness:</p> <ol style="list-style-type: none"> Number of new respiratory illness per person per week Pneumonia (cough or difficulty in breathing with a respiratory rate of > 60 min in children less than 60 days old, > 50 min in those less than 1 year old, and > 40 min for those aged 1 to 5 years) <p>Follow-up was weekly with household interview and direct observation. Children aged less than 5 were weighed, and the report presents stratification of results by child weight. Safety: N/A</p>
Notes	<p>Risk of bias: low (cluster coefficients and analysis by unit of randomisation provided)</p> <p>Note: the authors conclude that "handwashing" neighbourhoods has significantly fewer episodes of respiratory disease than controls (e.g. 50% less cough). "Handwashing" children aged less than 5 had 50% fewer episodes of pneumonia than controls (-65% to -35%). However, there was no difference in respiratory illness between types of soap. The report is confusing, with a shifting focus between children age groups. The impression reading is of an often rewritten manuscript. There is some loss of data (e.g. in the results by weight, i.e. risk group) because of lack of clarity on denominators. Despite this, the trial is a landmark.</p>

Risk of bias

Luby 2005 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation took place by computer-generated random numbers in 3 phases.
Allocation concealment (selection bias)	Low risk	"One of the investigators (SL) who did not participate in recruiting neighbourhoods or households programmed a spreadsheet to randomly generate the integers of a 1 or a 2. He applied the random numbers sequentially to the list of neighbourhoods. Neighbourhoods with a 1 were assigned to control, and those with a 2 were assigned to handwashing promotion. Random assignment continued until neighbourhoods consisted of at least 600 handwashing promotion households and 300 control households were assigned."
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"The antibacterial soap ... contained 1-2% triclocarban as an antibacterial substance. The plain soap was identical to the antibacterial soap except that it did not contain triclocarban... . Neither the fieldworkers nor the families knew whether soaps were antibacterial or plain."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Neither the fieldworkers nor the families knew whether soaps were antibacterial or plain."
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	89% of the study population followed up, but no data on the clusters.
Selective reporting (reporting bias)	Low risk	"At baseline, households in the three intervention groups were similar."

MacIntyre 2009
Study characteristics

Methods	Prospective cluster-RCT carried out in Sydney, Australia, to assess the use of surgical masks, P2 masks, and no masks in preventing ILI in households. The study was carried out during the 2 winter seasons of 2006 and 2007 (August to the end of October 2006 and June to the end of October 2007). "Gaussian random effects were incorporated in the model to account for the natural clustering of persons in households"
Participants	290 adults from 145 families. 47 households (94 enrolled adults and 180 children) were randomised to the surgical mask group, 46 (92 enrolled adults and 172 children) to the P2 mask group, and 52 (104 enrolled adults and 192 children) to the no-mask (control) group.
Interventions	Use of surgical masks and P2 mask versus no mask. The P2 mask is described as very cumbersome.
Outcomes	Laboratory: serological evidence Effectiveness: ILI (described as fever, history of fever or feeling feverish in the past week, myalgia, arthralgia, sore throat, cough, sneezing, runny nose, nasal congestion, headache) However, a positive laboratory finding for influenza converts the ILI definition into one of influenza. Safety: N/A
Notes	The study authors conclude that adherence to mask use significantly reduced the risk for ILI-associated infection, but < 50% of participants wore masks most of the time. They concluded that household use of face masks is associated with low adherence and is ineffective for controlling seasonal respiratory disease. Compliance was by self-report, therefore likely to be an underestimate.

MacIntyre 2009 (Continued)

The primary outcome was ILI or lab-positive illness. This showed no effect. Sensitivity analysis by adherence showed that under the assumption that the incubation period is equal to 1 day (the most probable value for the 2 most common viruses isolated, influenza (21) and rhinovirus (26)), adherent use of P2 or surgical masks significantly reduces the risk for ILI infection, with a hazard ratio = 0.26 (95% CI 0.09 to 0.77; P = 0.015). No other covariate was significant. Under the less likely assumption that the incubation period is equal to 2 days, the quantified effect of complying with P2 or surgical mask use remains strong, although borderline significant; hazard ratio was 0.32 (95% CI 0.11 to 0.98; P = 0.046). The study was underpowered to determine if there was a difference in efficacy between P2 and surgical masks (Table 5). The study conclusion appears to be a post hoc data exploration. Regardless of this, the study message is that respirator use in a family setting is unlikely to be effective as compliance is difficult unless there is a situation of real impending risk.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Participating households were randomised to 1 of 3 arms by a secure computerised randomisation process", but sequence generation not described.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	"Study participants and trial staff were not blinded, as it is not technically possible to blind the mask type to which participants were randomised."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"However, laboratory staff were blinded to the arm of randomisation."
Incomplete outcome data (attrition bias) All outcomes	Low risk	143 of 145 randomised families were analysed; 2 families in the control group were lost to follow-up during the study, for which no reasons were given.
Selective reporting (reporting bias)	Low risk	No differences between groups at baseline

MacIntyre 2011
Study characteristics

Methods	A cluster-RCT of 1441 HCWs in 15 Beijing hospitals was performed during the 2008 to 2009 winter. Participants wore masks or respirators during the entire work shift for 4 weeks. Outcomes included CRI, ILI, laboratory-confirmed respiratory virus infection, and influenza. A convenience no-mask/respirator group of 481 health workers from 9 hospitals was compared.
Participants	Participants (N = 1441) were hospital HCWs aged > 18 years from the emergency departments and respiratory wards of 15 hospitals. These wards were selected as high-risk settings in which repeated and multiple exposures to respiratory infections are expected. Participants were randomised to medical mask (N = 492 staff from 5 hospitals), N95 fit-tested masks (N = 461 staff from 5 hospitals), and N95 non-fit-tested mask (N = 488 staff from 5 hospitals).
Interventions	Fit-tested N95 respirators versus non-fit-tested N95 respirators versus medical masks. See Table 1 for details.

MacIntyre 2011 (Continued)

Outcomes	<p>Clinical respiratory illness, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom</p> <p>Influenza-like illness, defined as fever $\geq 38^{\circ}\text{C}$ plus 1 respiratory symptom (i.e. cough, runny nose, etc.)</p> <p>Laboratory-confirmed viral respiratory infection (detection of adenoviruses, human metapneumovirus, coronavirus 229E/NL63, parainfluenza viruses 1, 2, and 3, influenza viruses A and B, respiratory syncytial virus A and B, rhinovirus A or B, and coronavirus OC43/HKU1 by multiplex PCR)</p> <p>Laboratory-confirmed influenza A or B</p> <p>Adherence with mask or respirator use. Reported problems associated with using the masks or respirators</p>
Notes	Funding source unknown; control arm not randomised so has been ignored

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Randomisation process (using a secure computerised randomisation program), but sequence generation not described
Allocation concealment (selection bias)	Low risk	Hospitals randomised prior to inclusion of participants.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

MacIntyre 2013
Study characteristics

Methods	A cluster-RCT
Participants	<p>A total of 1669 nurses and doctors from 68 emergency departments and respiratory wards of 19 Beijing hospitals were included. Inclusion criteria: any nurse or doctor aged 18 years or older who worked full time in the emergency or respiratory wards was eligible. Exclusion: HCWs if they (1) were unable or refused to consent; (2) had beards, long moustaches, or long facial hair stubble; (3) had a current respiratory illness, rhinitis, and/or allergy; or (4) worked part time or did not work in the aforementioned wards or departments</p> <p>Final analysis was performed on 572 staff and 24 wards in medical mask group, 516 staff and 20 wards in the targeted N95 mask group, and 581 staff and 24 wards in the N95 mask group.</p>

MacIntyre 2013 (Continued)

Interventions	<p>Quote: "Masks used in the study were the 3M Standard Tie-On Surgical Mask (catalog number mask 1817; 3M, St. Paul, MN) and the 3M Health Care N95 Particulate Respirator (catalog number 1860; 3M)... . Participants wore the mask or respirator on every shift after being shown how to fit and wear it. Participants were supplied daily with either three masks for the medical mask arm or two N95 respirators. Participants using N95 respirators underwent a fit testing procedure using a 3M FT-30 Bitrex Fit Test Kit according to the manufacturer's instructions (3M)." See Table 1 for details.</p>	
Outcomes	<p>Laboratory:</p> <ol style="list-style-type: none"> 1. Laboratory-confirmed viral respiratory infection in symptomatic participants, defined as detection of adenoviruses; human metapneumovirus; coronaviruses 229E/NL63 and OC43/HKU1; parainfluenza viruses 1, 2, and 3; influenza viruses A and B; respiratory syncytial viruses A and B; or rhinoviruses A/B by nucleic acid testing (NAT) using a commercial multiplex polymerase chain reaction (Seegen, Inc., Seoul, Korea). 2. Laboratory-confirmed influenza A or B in symptomatic participants. 3. Laboratory-confirmed bacterial colonisation in symptomatic participants, defined as detection of <i>Streptococcus pneumoniae</i>, <i>Legionella</i>, <i>Bordetella pertussis</i>, chlamydia, <i>Mycoplasma pneumoniae</i>, or <i>Haemophilus influenzae</i> type B by multiplex polymerase chain reaction (Seegen, Inc.). <p>Effectiveness: CRI, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom. ILI, defined as fever (38 °C) plus 1 respiratory symptom</p> <p>Safety: adverse effects measured using a semi-structured questionnaire. Investigators stated that there was higher reported adverse effects and discomfort of N95 respirators compared with the other 2 arms. In terms of comfort, 52% (297 of 571) of the medical mask arm reported no problems, compared with 62% (317 of 512) of the targeted arm and 38% (217 of 574) of the N95 arm ($P < 0.001$).</p>	
Notes	<p>Compliance with the product was highest in the targeted N95 arm (82%; 422 of 516), then the medical mask arm (66%; 380 of 572), and the N95 arm (57%; 333 of 581); these differences were statistically significant ($P < 0.001$).</p> <p>The period study conducted: 28 December 2009 to 7 February 2010</p> <p>Funding: unclear</p>	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"using a secure computerized randomization program", but sequence generation not described
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Outcome was objectively assessed with lab confirmation in addition to clinical illness.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Laboratory outcomes are reported for all subjects (with at least one respiratory symptom or fever) tested, and then for the subset meeting the CRI definition"
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up. Flow chart and text match, investigators conducted ITT and PP analysis. All the outcomes were accounted for amongst all participants.

MacIntyre 2013 (Continued)

Selective reporting (reporting bias)	Low risk	All outcomes were reported as planned.
--------------------------------------	----------	--

MacIntyre 2015
Study characteristics

Methods	A cluster-RCT of cloth masks compared with medical masks in healthcare workers in 14 secondary-/tertiary-level hospitals in Hanoi, Vietnam. Hospital wards were randomised to: medical masks, cloth masks, or a control group (usual practice, which included mask wearing). Participants used the mask on every shift for 4 consecutive weeks.	
Participants	1607 hospital HCWs aged ≥ 18 years working full time in selected high-risk wards. Medical mask group (n = 580 HCWs), cloth mask group (n = 569 HCWs), control group (n = 458 HCWs)	
Interventions	Medical masks, cloth masks, or a control group. See Table 1 for details.	
Outcomes	Clinical respiratory illness, influenza-like illness, and laboratory-confirmed respiratory virus infection <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom 2. Influenza-like illness, defined as fever ≥ 38 °C plus 1 respiratory symptom 3. Laboratory-confirmed viral respiratory infection. Laboratory confirmation was by nucleic acid detection using multiplex reverse transcriptase PCR (RT-PCR) for 17 respiratory viruses. Adverse events associated with mask use	
Notes	Government funded	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Epi info V.6 was used to generate a randomisation allocation.
Allocation concealment (selection bias)	Low risk	74 wards randomised prior to recruitment of individuals.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Low risk	Specified endpoints reported.

MacIntyre 2015

Study characteristics

Methods	A cluster-RCT of cloth masks compared with medical masks in healthcare workers in 14 secondary-/tertiary-level hospitals in Hanoi, Vietnam. Hospital wards were randomised to: medical masks, cloth masks, or a control group (usual practice, which included mask wearing). Participants used the mask on every shift for 4 consecutive weeks.
Participants	1607 hospital HCWs aged ≥ 18 years working full time in selected high-risk wards. Medical mask group (n = 580 HCWs), cloth mask group (n = 569 HCWs), control group (n = 458 HCWs)
Interventions	Medical masks, cloth masks, or a control group. See Table 1 for details.
Outcomes	Clinical respiratory illness, influenza-like illness, and laboratory-confirmed respiratory virus infection <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom 2. Influenza-like illness, defined as fever ≥ 38 °C plus 1 respiratory symptom 3. Laboratory-confirmed viral respiratory infection. Laboratory confirmation was by nucleic acid detection using multiplex reverse transcriptase PCR (RT-PCR) for 17 respiratory viruses. Adverse events associated with mask use
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Epi info V.6 was used to generate a randomisation allocation.
Allocation concealment (selection bias)	Low risk	74 wards randomised prior to recruitment of individuals.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Low risk	Specified endpoints reported.

MacIntyre 2016

Study characteristics

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Copyright © 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

MacIntyre 2016 (Continued)

Methods	Cluster-RCT to examine medical mask use as source control for people with respiratory illness in 6 major hospitals in 2 districts of Beijing, China. Index cases with ILI were randomly allocated to medical mask (n = 123) and control arms (n = 122). Since 43 index cases in the control arm also used a mask during the study period, an as-treated post hoc analysis was performed by comparing outcomes amongst household members of index cases who used a mask (mask group) with household members of index cases who did not use a mask (no mask group).
Participants	245 index cases with ILI (medical mask = 123, control group = 122) and 597 household contacts (medical mask = 302, control group = 295)
Interventions	Medical mask versus no mask (control). See Table 1 for details.
Outcomes	<p>Clinical respiratory illness, ILI, and laboratory-confirmed viral respiratory infection</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms (cough, nasal congestion, runny nose, sore throat, or sneezes) or 1 respiratory symptom and a systemic symptom (chill, lethargy, loss of appetite, abdominal pain, muscle or joint aches). 2. ILI, defined as fever $\geq 38^{\circ}\text{C}$ plus 1 respiratory symptom. 3. Laboratory-confirmed viral respiratory infection, defined as detection of adenoviruses, human metapneumovirus, coronaviruses 229E/NL63 and OC43/HKU1, parainfluenza viruses 1, 2, and 3, influenza viruses A and B, respiratory syncytial virus A and B, or rhinovirus A/B by nucleic acid testing using a commercial multiplex PCR. <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Random allocation sequence using Microsoft Excel
Allocation concealment (selection bias)	High risk	Doctors enrolled the participants randomly to intervention and control arms.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Clinical endpoints assessed unblinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

MacIntyre 2016
Study characteristics
Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

MacIntyre 2016 (Continued)

Methods	Cluster-RCT to examine medical mask use as source control for people with respiratory illness in 6 major hospitals in 2 districts of Beijing, China. Index cases with ILI were randomly allocated to medical mask (n = 123) and control arms (n = 122). Since 43 index cases in the control arm also used a mask during the study period, an as-treated post hoc analysis was performed by comparing outcomes amongst household members of index cases who used a mask (mask group) with household members of index cases who did not use a mask (no mask group).
Participants	245 index cases with ILI (medical mask = 123, control group = 122) and 597 household contacts (medical mask = 302, control group = 295)
Interventions	Medical mask versus no mask (control). See Table 1 for details.
Outcomes	<p>Clinical respiratory illness, ILI, and laboratory-confirmed viral respiratory infection</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms (cough, nasal congestion, runny nose, sore throat, or sneezes) or 1 respiratory symptom and a systemic symptom (chill, lethargy, loss of appetite, abdominal pain, muscle or joint aches). 2. ILI, defined as fever $\geq 38^\circ\text{C}$ plus 1 respiratory symptom. 3. Laboratory-confirmed viral respiratory infection, defined as detection of adenoviruses, human metapneumovirus, coronaviruses 229E/NL63 and OC43/HKU1, parainfluenza viruses 1, 2, and 3, influenza viruses A and B, respiratory syncytial virus A and B, or rhinovirus A/B by nucleic acid testing using a commercial multiplex PCR. <p>No safety outcomes reported.</p>
Notes	Government funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Random allocation sequence using Microsoft Excel
Allocation concealment (selection bias)	High risk	Doctors enrolled the participants randomly to intervention and control arms.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Clinical endpoints assessed unblinded.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Low risk	Specified outcomes reported.

McConeghy 2017
Study characteristics
Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

McConeghy 2017 (Continued)

Methods	Pilot study of comprehensive intervention (education, cleaning of surfaces, audit and feedback) to staff of nursing homes versus usual care. Pair-matched cluster-randomised design with only 5 clusters (nursing homes) in each group
Participants	10 nursing homes in Colorado, USA Intervention group = 481 long-stay residents and control group = 380 'Long-stay' defined as resident at least 90 days prior to baseline, or recently readmitted after previous long stay.
Interventions	A multifaceted hand-washing/surface-cleaning intervention comprised of 1) 1-hour online educational module focused on how to prevent infections; 2) provided with an "essential bundle" of 7 products, ranging from hand sanitiser gel and foam to antiviral facial tissues, disinfecting spray, and hand and face wipe and recommendation to use 4 skin cream and wipe products; 3) audit and feedback system. See Table 1 for details.
Outcomes	Laboratory: surface cultures mentioned in Methods, but no results given Effectiveness: LRTI, all infections, hospitalisation, use of antibiotics (not relevant to this review) Safety: none mentioned in Methods and no results given
Notes	The authors conclude that "This multifaceted hand-washing and surface cleaning intervention was designed to reduce infection rates among nursing homes residents. In our 10-facility randomized, matched pair pilot study, we observed program compliance and satisfaction along with reductions in surface bacterial counts, but did not observe a statistically significant reduction in infection rates, antimicrobial use, or hospitalizations". Very poorly reported study with results not explained, summarised in Table 3 as RDs. Denominators and attrition are unclear.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Method not described
Allocation concealment (selection bias)	Unclear risk	Method not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Illness and absenteeism reported by treating staff.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	No attrition given. Data were collected from e-medical record at baseline, but not clear whether illness data during the study were collected by the same method.
Selective reporting (reporting bias)	High risk	Upper respiratory tract infection was mentioned in the Methods (intervention presumably would target these), but only LRTI and overall infection reported.

McConeghy 2017
Study characteristics

Methods	Pilot study of comprehensive intervention (education, cleaning of surfaces, audit and feedback) to staff of nursing homes versus usual care. Pair-matched cluster-randomised design with only 5 clusters (nursing homes) in each group
Participants	10 nursing homes in Colorado, USA Intervention group = 481 long-stay residents and control group = 380 'Long-stay' defined as resident at least 90 days prior to baseline, or recently readmitted after previous long stay.
Interventions	A multifaceted hand-washing/surface-cleaning intervention comprised of 1) 1-hour online educational module focused on how to prevent infections; 2) provided with an "essential bundle" of 7 products, ranging from hand sanitiser gel and foam to antiviral facial tissues, disinfecting spray, and hand and face wipe and recommendation to use 4 skin cream and wipe products; 3) audit and feedback system. See Table 1 for details.
Outcomes	Laboratory: surface cultures mentioned in Methods, but no results given Effectiveness: LRTI, all infections, hospitalisation, use of antibiotics (not relevant to this review) Safety: none mentioned in Methods and no results given
Notes	The authors conclude that "This multifaceted hand-washing and surface cleaning intervention was designed to reduce infection rates among nursing homes residents. In our 10-facility randomized, matched pair pilot study, we observed program compliance and satisfaction along with reductions in surface bacterial counts, but did not observe a statistically significant reduction in infection rates, antimicrobial use, or hospitalizations". Very poorly reported study with results not explained, summarised in Table 3 as RDs. Denominators and attrition are unclear.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Method not described
Allocation concealment (selection bias)	Unclear risk	Method not described
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	High risk	Illness and absenteeism reported by treating staff.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	No attrition given. Data were collected from e-medical record at baseline, but not clear whether illness data during the study were collected by the same method.

McConeghy 2017 (Continued)

Selective reporting (reporting bias)	High risk	Upper respiratory tract infection was mentioned in the Methods (intervention presumably would target these), but only LRTI and overall infection reported.
--------------------------------------	-----------	--

Millar 2016
Study characteristics

Methods	Cluster-RCT, open-label study, factorial design
Participants	Around 30,000 healthy, male army trainees aged 18 to 42 years at Fort Benning, Georgia were included. Inclusion criteria: trainees assigned to 1 of the 6 selected training battalions, trainees who present with an SSTI at the clinic or the hospital, provide informed consent. Exclusion criteria: fails to meet inclusion criteria. No denominator breakdown by arm is reported.
Interventions	Promotion of hand-washing in addition to a once-weekly application of chlorhexidine-based body wash. See Table 1 for details.
Outcomes	This study was nested in a large field-based RCT and utilised clinic-based medical records. Laboratory: none Effectiveness: incidence of ARI at 20 months. The case definition was any occurrence of the following ICD-9 symptom or disease-specific codes: 460 to 466, 480 to 488, and specifically 465.9, 482.9, 486, and 487.1. Safety: adverse effects neither planned nor reported by the investigators
Notes	The period study conducted: May 2010 to January 2012 Funding: government

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"computer-generated random numbers to 1 of the 3 study groups"
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	The study was open-label and self-reporting of ARI. It is planned as secondary objective of an original trial. Data abstractors were blinded to group assignment.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Data abstractors were blinded to group assignment.
Incomplete outcome data (attrition bias) All outcomes	High risk	There is a statistically significant difference between attrition rates in the 3 groups. The reasons for attrition are briefly reported in Table 1 of the original study (Ellis and colleagues 2014), but are unlikely to be related to the outcomes of this study. ARI cases were captured utilising clinic-based medical records, but this outcome is not prespecified in the protocol.

Millar 2016 (Continued)

Selective reporting (reporting bias)	High risk	The study was conducted for another purpose. According to the study protocol, the outcomes of interest in the current report were not mentioned as outcomes when the study was planned. ARI is not prespecified as an outcome in the protocol published on ClinicalTrials.gov.
--------------------------------------	-----------	--

Millar 2016
Study characteristics

Methods	Cluster-RCT, open-label study, factorial design
Participants	Around 30,000 healthy, male army trainees aged 18 to 42 years at Fort Benning, Georgia were included. Inclusion criteria: trainees assigned to 1 of the 6 selected training battalions, trainees who present with an SSTI at the clinic or the hospital, provide informed consent. Exclusion criteria: fails to meet inclusion criteria. No denominator breakdown by arm is reported.
Interventions	Promotion of hand-washing in addition to a once-weekly application of chlorhexidine-based body wash. See Table 1 for details.
Outcomes	This study was nested in a large field-based RCT and utilised clinic-based medical records. Laboratory: none Effectiveness: incidence of ARI at 20 months. The case definition was any occurrence of the following ICD-9 symptom or disease-specific codes: 460 to 466, 480 to 488, and specifically 465.9, 482.9, 486, and 487.1. Safety: adverse effects neither planned nor reported by the investigators
Notes	The period study conducted: May 2010 to January 2012 Funding: government

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"computer-generated random numbers to 1 of the 3 study groups"
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	The study was open-label and self-reporting of ARI. It is planned as secondary objective of an original trial. Data abstractors were blinded to group assignment.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Data abstractors were blinded to group assignment.
Incomplete outcome data (attrition bias) All outcomes	High risk	There is a statistically significant difference between attrition rates in the 3 groups. The reasons for attrition are briefly reported in Table 1 of the original study (Ellis and colleagues 2014), but are unlikely to be related to the out-

Millar 2016 (Continued)

comes of this study. ARI cases were captured utilising clinic-based medical records, but this outcome is not prespecified in the protocol.

Selective reporting (reporting bias)	High risk	The study was conducted for another purpose. According to the study protocol, the outcomes of interest in the current report were not mentioned as outcomes when the study was planned. ARI is not prespecified as an outcome in the protocol published on ClinicalTrials.gov.
--------------------------------------	-----------	--

Miyaki 2011
Study characteristics

Methods	A quasi-cluster-RCT	
Participants	<p>A total of 15,134 assigned to intervention (N = 6634 workers) and control (N = 8500 workers)</p> <p>Inclusion criteria: all general employees (aged 19 to 72 years in 2009) of 2 sibling companies of a major car industry in Kanagawa Prefecture, Japan. All workers who regularly reported to the workplace were included, regardless of treatment for chronic diseases.</p> <p>All employees have the same health insurance plan and were followed up in the same way.</p>	
Interventions	<p>"The intervention involved asking workers whose family members developed an influenza-like illness (ILI) to stay at home. If any co-habiting family members showed signs of influenza-like illness (ILI), employees ... were asked to stay at home voluntarily until 5 days has passed since the resolution of the ILS symptoms or 2 days after alleviation of fever." See Table 1 for details.</p>	
Outcomes	<p>Workroom: influenza A test kit (rapid test)</p> <p>Effectiveness: assess the effectiveness of household quarantine in reducing the incidence of influenza A H1N1. ILI was defined as a body temperature greater than 38 °C or more than 1 °C above the normal temperature accompanied with more than 2 of these symptoms: nasal mucus, pharyngeal pain, cough, chills or heat sensation</p> <p>Safety: the incidence of influenza A H1N1 amongst workers who were told to stay home if a family member developed ILI was higher (relative risk of 2.17; P < 0.001) compared to control group. No other safety measures/harms reported.</p> <p>Compliance: quote: "our intervention was not compulsory; we only asked the employees to leave the workplace for a while on full pay, and we succeeded in getting all workers' agreement. In our case, explaining that the home waiting policy might be beneficial to the whole workers and help to avoid stopping the manufacturing lines (explaining it is for the benefit of the public) and guaranteeing payment during the leave (financial support) helped them to obey our request."</p>	
Notes	<p>Period study conducted: 1 July 2009 to 19 February 2010</p> <p>Unfunded</p>	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No information given.
Allocation concealment (selection bias)	Unclear risk	No information given.

Miyaki 2011 (Continued)

Blinding of participants and personnel (performance bias) All outcomes	Low risk	The nature of the intervention (stay at home) was confirmed in the intervention group, where all workers agree as they were financially supported during absences due to ILI.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Company doctors diagnosed the disease through a positive result of an influenza A test or clinical symptoms", but not clear if they were blinded to assignment; however, the diagnostic process is meticulous and objectively confirmed.
Incomplete outcome data (attrition bias) All outcomes	Low risk	All cases are included in the analysis, and none were lost to follow-up.
Selective reporting (reporting bias)	Unclear risk	Although all outcomes of interest are clearly specified, described, and followed up, and text and numbers checked out well and based on the outcome stated for the study, there is no published protocol to match the planned vs the reported outcomes.

Miyaki 2011

Study characteristics

Methods	A quasi-cluster-RCT
Participants	A total of 15,134 assigned to intervention (N = 6634 workers) and control (N = 8500 workers) Inclusion criteria: all general employees (aged 19 to 72 years in 2009) of 2 sibling companies of a major car industry in Kanagawa Prefecture, Japan. All workers who regularly reported to the workplace were included, regardless of treatment for chronic diseases. All employees have the same health insurance plan and were followed up in the same way.
Interventions	"The intervention involved asking workers whose family members developed an influenza-like illness (ILI) to stay at home. If any co-habiting family members showed signs of influenza-like illness (ILI), employees ... were asked to stay at home voluntarily until 5 days has passed since the resolution of the ILS symptoms or 2 days after alleviation of fever." See Table 1 for details.
Outcomes	Workroom: influenza A test kit (rapid test) Effectiveness: assess the effectiveness of household quarantine in reducing the incidence of influenza A H1N1. ILI was defined as a body temperature greater than 38 °C or more than 1 °C above the normal temperature accompanied with more than 2 of these symptoms: nasal mucus, pharyngeal pain, cough, chills or heat sensation Safety: the incidence of influenza A H1N1 amongst workers who were told to stay home if a family member developed ILI was higher (relative risk of 2.17; P < 0.001) compared to control group. No other safety measures/harms reported. Compliance: quote: "our intervention was not compulsory; we only asked the employees to leave the workplace for a while on full pay, and we succeeded in getting all workers' agreement. In our case, explaining that the home waiting policy might be beneficial to the whole workers and help to avoid stopping the manufacturing lines (explaining it is for the benefit of the public) and guaranteeing payment during the leave (financial support) helped them to obey our request."
Notes	Period study conducted: 1 July 2009 to 19 February 2010

Miyaki 2011 (Continued)

Unfunded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No information given.
Allocation concealment (selection bias)	Unclear risk	No information given.
Blinding of participants and personnel (performance bias) All outcomes	Low risk	The nature of the intervention (stay at home) was confirmed in the intervention group, where all workers agree as they were financially supported during absences due to ILI.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Company doctors diagnosed the disease through a positive result of an influenza A test or clinical symptoms", but not clear if they were blinded to assignment; however, the diagnostic process is meticulous and objectively confirmed.
Incomplete outcome data (attrition bias) All outcomes	Low risk	All cases are included in the analysis, and none were lost to follow-up.
Selective reporting (reporting bias)	Unclear risk	Although all outcomes of interest are clearly specified, described, and followed up, and text and numbers checked out well and based on the outcome stated for the study, there is no published protocol to match the planned vs the reported outcomes.

Morton 2004
Study characteristics

Methods	Cross-over study to evaluate the effectiveness of an alcohol gel as an adjunct to regular hand-washing for decreasing absenteeism amongst elementary children by reducing specific communicable diseases such cold, flu, and conjunctivitis. The study was conducted in an elementary school in New England, USA. In the cross-over design, classrooms in each grade level were randomised to begin as the experimental group (alcohol gel) or the control group (regular hand-washing). A study protocol for hand hygiene was introduced following the germ unit education. The hand-washing product was a soap-and-water alternative that is approximately 60% ethyl alcohol. In phase 1 (46 days) children in 9 classrooms were in the experimental group, and children in 8 classrooms were in the control group. After a 1-week washout period when no children had access to the alcohol gel, phase 2 (47 days) started, and the classroom that had participated before as experimental group passed into the control group and vice versa. Data were collected by the parents, who informed the secretary or the school nurse of the reasons for a child's absence, including symptoms of any illness. Respiratory illnesses were defined by symptoms of URTI.
Participants	253 children, 120 girls and 133 boys, from kindergarten to 3rd grade. Of the eligible 285 students, 32 children dropped out (10 due to skin irritation and 22 because of lack of parental consent). No denominator breakdown by arm is reported because the study used a cross-over design.
Interventions	Use of an alcohol gel as an adjunct to regular hand-washing and educational programme versus regular hand-washing and educational programme
Outcomes	Laboratory: no

Morton 2004 (Continued)

 Effectiveness: days of absences from school for respiratory illness
 Safety: N/A

Notes

Risk of bias: high (no description of randomisation; partial reporting of outcomes, numerators and denominators)

Note: the authors conclude that significantly fewer children became ill whilst using the alcohol gel as an adjunct to regular hand-washing than when using regular hand-washing only (decreased school absenteeism of 43% with the use of alcohol gel on top of hand-washing). The authors also described, as a limitation of the study, the fact that the school nurse served as the data collector, which could be perceived as bias in measurement of the outcome variable.

Randomisation and allocation are not described; no cluster coefficients were reported; and attrition was not taken into consideration during the analysis. Unit of randomisation and analysis are different. No reporting by arm. No ORs, no CIs reported.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Unclear risk	Insufficient information
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	"A cross-over design was used. In the crossover design, classrooms in each grade level were randomized to begin as the experimental group (regular hand washing)."
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"The school nurse served as the data collector for the duration of the study. This could be perceived as bias in the measurement of the outcome variable, absenteeism related to infectious illness."
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information
Selective reporting (reporting bias)	Unclear risk	Insufficient information

Najnin 2019
Study characteristics

Methods	Cluster-RCT, parallel assignment
Participants	<p>Residents of the high-risk, cholera-prone study areas. Low-income communities in Mirpur area of urban Dhaka defined by low per capita income, poor sanitation, unsafe water use, sharing of water source, and poor living conditions. 90 geographic clusters were included, with 30-metre buffer zones.</p> <p>A total of 7842 households, with 52,237 individuals analysed</p> <p>Vaccine-only area: data were analysed for 1965 households consisting of 13,148 individuals</p> <p>Vaccine-plus-behaviour-change area: data were analysed for 3886 households consisting of 25,566 individuals</p> <p>Control area: data were analysed for 1991 households consisting of 13,523 individuals</p>

Najnin 2019 (Continued)

Study criteria from published protocol:

Inclusion criteria: apparently healthy residents of selected vaccination sites, aged 1 year and above, non-pregnant women, written informed consent

Exclusion criteria: age less than 1 year and pregnant women

Interventions	Hand-washing and water treatment promotion. See Table 1 for details.
Outcomes	<p>Laboratory: none used</p> <p>Effectiveness: prevalence of respiratory illness. People were classified as having respiratory illness if they reported having fever plus either cough or nasal congestion or fever plus breathing difficulty in the past 2 days of unannounced home visits: in each intervention group and amongst those who had soap/soapy water with water present in the hand-washing station (35% of all groups combined) versus those without this (regardless of the intervention group). Planned secondary outcome: prevalence of reported respiratory illness during 2-year intervention period</p> <p>Safety: no adverse effects planned or reported</p>
Notes	<p>The period study conducted: 2011 to 2013</p> <p>Funding: government and private Bill & Melinda Gates Foundation</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated randomisation sequence was used to allocate 90 geographical clusters to 1 of 3 groups. Before randomisation, clusters were stratified blocked into 2 categories according to the distance to the hospital. (parent article: Lancet. 2015 Oct 3;386(10001):1362-1371)
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	All trial participants and investigators were aware of group assignment. Several in and out migrations across all groups before, after, and during outcome monitoring, and large number of changes in intervention areas
Blinding of outcome assessment (detection bias) All outcomes	High risk	Several in and out migrations across all groups before, after, and during outcome monitoring, and large number of changes in intervention areas
Incomplete outcome data (attrition bias) All outcomes	High risk	High migration movement. This could have distorted the baseline characteristics even more. Very hard to assess because the numbers in the index paper are different from the parent paper (Qadri 2015). In addition to that, for each intervention, data were analysed for 15% to 30% of those allocated on start date. Each group started with approximately 80,000 people; the number analysed is much lower (237,216 people were in the study area on start date of outcome monitoring, the total number analysed across all groups was 52,237). No info about data on migrated individuals or on those who changed intervention areas was dealt with? Also data for prevalence of ARI adjusted for age and wealth were not shown. The outcome is addressed in the 2 days preceding an unannounced visit. This means that if there was a respiratory illness in the past week it would not have been reported. Moreover, these monthly unannounced visits were done to a different set of participants in each group!

Najnin 2019 (Continued)

Selective reporting (reporting bias)	High risk	Published protocol does not include respiratory illness as an outcome.
--------------------------------------	-----------	--

Morton 2004
Study characteristics

Methods	Cross-over study to evaluate the effectiveness of an alcohol gel as an adjunct to regular hand-washing for decreasing absenteeism amongst elementary children by reducing specific communicable diseases such cold, flu, and conjunctivitis. The study was conducted in an elementary school in New England, USA. In the cross-over design, classrooms in each grade level were randomised to begin as the experimental group (alcohol gel) or the control group (regular hand-washing). A study protocol for hand hygiene was introduced following the germ unit education. The hand-washing product was a soap-and-water alternative that is approximately 60% ethyl alcohol. In phase 1 (46 days) children in 9 classrooms were in the experimental group, and children in 8 classrooms were in the control group. After a 1-week washout period when no children had access to the alcohol gel, phase 2 (47 days) started, and the classroom that had participated before as experimental group passed into the control group and vice versa. Data were collected by the parents, who informed the secretary or the school nurse of the reasons for a child's absence, including symptoms of any illness. Respiratory illnesses were defined by symptoms of URTI.
Participants	253 children, 120 girls and 133 boys, from kindergarten to 3rd grade. Of the eligible 285 students, 32 children dropped out (10 due to skin irritation and 22 because of lack of parental consent). No denominator breakdown by arm is reported because the study used a cross-over design.
Interventions	Use of an alcohol gel as an adjunct to regular hand-washing and educational programme versus regular hand-washing and educational programme
Outcomes	Laboratory: no Effectiveness: days of absences from school for respiratory illness Safety: N/A
Notes	Risk of bias: high (no description of randomisation; partial reporting of outcomes, numerators and denominators) Note: the authors conclude that significantly fewer children became ill whilst using the alcohol gel as an adjunct to regular hand-washing than when using regular hand-washing only (decreased school absenteeism of 43% with the use of alcohol gel on top of hand-washing). The authors also described, as a limitation of the study, the fact that the school nurse served as the data collector, which could be perceived as bias in measurement of the outcome variable. Randomisation and allocation are not described; no cluster coefficients were reported; and attrition was not taken into consideration during the analysis. Unit of randomisation and analysis are different. No reporting by arm. No ORs, no CIs reported.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Unclear risk	Insufficient information
Blinding of participants and personnel (performance bias)	Unclear risk	"A cross-over design was used. In the crossover design, classrooms in each grade level were randomized to begin as the experimental group (regular hand washing)."

Morton 2004 (Continued)

All outcomes

Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"The school nurse served as the data collector for the duration of the study. This could be perceived as bias in the measurement of the outcome variable, absenteeism related to infectious illness."
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information
Selective reporting (reporting bias)	Unclear risk	Insufficient information

Najnin 2019

Study characteristics

Methods	Cluster-RCT, parallel assignment
Participants	<p>Residents of the high-risk, cholera-prone study areas. Low-income communities in Mirpur area of urban Dhaka defined by low per capita income, poor sanitation, unsafe water use, sharing of water source, and poor living conditions. 90 geographic clusters were included, with 30-metre buffer zones.</p> <p>A total of 7842 households, with 52,237 individuals analysed</p> <p>Vaccine-only area: data were analysed for 1965 households consisting of 13,148 individuals</p> <p>Vaccine-plus-behaviour-change area: data were analysed for 3886 households consisting of 25,566 individuals</p> <p>Control area: data were analysed for 1991 households consisting of 13,523 individuals</p> <p>Study criteria from published protocol:</p> <p>Inclusion criteria: apparently healthy residents of selected vaccination sites, aged 1 year and above, non-pregnant women, written informed consent</p> <p>Exclusion criteria: age less than 1 year and pregnant women</p>
Interventions	Hand-washing and water treatment promotion. See Table 1 for details.
Outcomes	<p>Laboratory: none used</p> <p>Effectiveness: prevalence of respiratory illness. People were classified as having respiratory illness if they reported having fever plus either cough or nasal congestion or fever plus breathing difficulty in the past 2 days of unannounced home visits: in each intervention group and amongst those who had soap/soapy water with water present in the hand-washing station (35% of all groups combined) versus those without this (regardless of the intervention group). Planned secondary outcome: prevalence of reported respiratory illness during 2-year intervention period</p> <p>Safety: no adverse effects planned or reported</p>
Notes	<p>The period study conducted: 2011 to 2013</p> <p>Funding: government and private Bill & Melinda Gates Foundation</p>

Risk of bias

Najnin 2019 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated randomisation sequence was used to allocate 90 geographical clusters to 1 of 3 groups. Before randomisation, clusters were stratified blocked into 2 categories according to the distance to the hospital. (parent article: Lancet. 2015 Oct 3;386(10001):1362-1371)
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	All trial participants and investigators were aware of group assignment. Several in and out migrations across all groups before, after, and during outcome monitoring, and large number of changes in intervention areas
Blinding of outcome assessment (detection bias) All outcomes	High risk	Several in and out migrations across all groups before, after, and during outcome monitoring, and large number of changes in intervention areas
Incomplete outcome data (attrition bias) All outcomes	High risk	High migration movement. This could have distorted the baseline characteristics even more. Very hard to assess because the numbers in the index paper are different from the parent paper (Qadri 2015). In addition to that, for each intervention, data were analysed for 15% to 30% of those allocated on start date. Each group started with approximately 80,000 people; the number analysed is much lower (237,216 people were in the study area on start date of outcome monitoring, the total number analysed across all groups was 52,237). No info about data on migrated individuals or on those who changed intervention areas was dealt with? Also data for prevalence of ARI adjusted for age and wealth were not shown. The outcome is addressed in the 2 days preceding an unannounced visit. This means that if there was a respiratory illness in the past week it would not have been reported. Moreover, these monthly unannounced visits were done to a different set of participants in each group!
Selective reporting (reporting bias)	High risk	Published protocol does not include respiratory illness as an outcome.

Nicholson 2014
Study characteristics

Methods	Cluster-RCT
Participants	70 low-income communities in Mumbai, India (35 communities per arm) were randomised to intervention arm (N = 1025) and control arm (N = 1026). Households located in low-income urban communities in west and south Mumbai, India. Each household contains 1 target child in the first year of a municipal school (typically aged 5 years).
Interventions	Combination of hand-washing promotion with provision of free soap aimed at 5-year-olds with provision of free soap. See Table 1 for details.
Outcomes	Laboratory: none reported Effectiveness: Primary outcomes: episodes of diarrhoea, ARIs, and school absences amongst target children, and episodes of diarrhoea and ARIs among their families

Nicholson 2014 (Continued)

Secondary outcomes: episodes of eye infections, vomiting, abscesses or boils, headaches, and earache

Operational definitions for all the illnesses were taken from *Black's Medical Dictionary* (MacPherson 1999). ARIs as "pneumonia, cough, fever, chest pain and shortness of breath, cold, inflammation of any or all of the airways, that is, nose, sinuses, throat, larynx, trachea and bronchi"

Safety: no safety measures planned or reported by the investigators

Notes

The period study conducted: 22 October 2007 to 2 August 2008

Funding: multinational corporate company (Unilever plc.)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Coin tossing used, which could have led to a large imbalance.
Allocation concealment (selection bias)	Low risk	"a coin toss was used to assign one community in each pair to intervention and one to control"
Blinding of participants and personnel (performance bias) All outcomes	High risk	Participants knew to which arm they had been recruited. Households were removed from the study if they provided no data for 5 consecutive weeks.
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data collectors were independent of the behaviour change intervention. Each was assigned exclusively to either households in the intervention group or to control households. However, communities, where very low literacy levels exist, were replaced after randomisation.
Incomplete outcome data (attrition bias) All outcomes	High risk	Data for non-completers were available and similar across groups. ITT and PP were performed. However, households were removed from the study if they provided no data for 5 consecutive weeks.
Selective reporting (reporting bias)	Unclear risk	No information to judge

Pandejpong 2012
Study characteristics

Methods	Cluster-RCT, single study centre
Participants	<p>Children (total number = 1437) were randomised to alcohol hand gel every 60 minutes (N = 452 children), every 120 minutes (N = 447 children), and once before lunch (N = 540 children).</p> <p>Inclusion criteria: all children in a large private school in suburban Bangkok, Thailand, all ages, both genders with parental consent to participate.</p> <p>Exclusion criteria: an allergy to alcohol hand gel</p>
Interventions	3 disinfection interventions: Alcohol hand gel applied every 60 minutes vs every 120 minutes vs once before lunch (3 groups). The current school standard for hand hygiene (q lunch group). See Table 1 for details.
Outcomes	Laboratory: none

Pandejpong 2012 (Continued)

Effectiveness:

Primary: rates of absenteeism from physician-confirmed ILI

Secondary: rate of absenteeism caused by total reported ILI (with and without a doctor's confirmation)

In case the child was sick but did not see a doctor, the parents were asked to report any of the following symptoms: runny nose or cough, fever or chills, sore throat, headache, diarrhoea, and presence of hand, foot, or mouth ulcers. If 2 or more of these symptoms were reported, then the child's illness was documented as an ILI.

Safety: investigators reported that no adverse reaction to the alcohol hand gel was reported in any participants

Notes

The period study conducted: December 2009 to February 2010

Funding: Royal College of Physicians of Thailand

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No information provided.
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Parents and teachers are aware of the assignment. Teachers were responsible for recording the absenteeism case record forms. Parents would report child sickness. No diagnostic tests, even in the case of physician-confirmed ILI
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome is physician-confirmed ILI.
Incomplete outcome data (attrition bias) All outcomes	Low risk	"No students were lost to follow-up or discontinued the intervention during the study period."
Selective reporting (reporting bias)	Low risk	All outcomes were reported.

Pandejpong 2012

Study characteristics

Methods	Cluster-RCT, single study centre
Participants	<p>Children (total number = 1437) were randomised to alcohol hand gel every 60 minutes (N = 452 children), every 120 minutes (N = 447 children), and once before lunch (N = 540 children).</p> <p>Inclusion criteria: all children in a large private school in suburban Bangkok, Thailand, all ages, both genders with parental consent to participate.</p> <p>Exclusion criteria: an allergy to alcohol hand gel</p>

Pandejpong 2012 (Continued)

Interventions	3 disinfection interventions: Alcohol hand gel applied every 60 minutes vs every 120 minutes vs once before lunch (3 groups). The current school standard for hand hygiene (q lunch group). See Table 1 for details.
Outcomes	<p>Laboratory: none</p> <p>Effectiveness:</p> <p>Primary: rates of absenteeism from physician-confirmed ILI</p> <p>Secondary: rate of absenteeism caused by total reported ILI (with and without a doctor's confirmation)</p> <p>In case the child was sick but did not see a doctor, the parents were asked to report any of the following symptoms: runny nose or cough, fever or chills, sore throat, headache, diarrhoea, and presence of hand, foot, or mouth ulcers. If 2 or more of these symptoms were reported, then the child's illness was documented as an ILI.</p> <p>Safety: investigators reported that no adverse reaction to the alcohol hand gel was reported in any participants</p>
Notes	<p>The period study conducted: December 2009 to February 2010</p> <p>Funding: Royal College of Physicians of Thailand</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No information provided.
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Parents and teachers are aware of the assignment. Teachers were responsible for recording the absenteeism case record forms. Parents would report child sickness. No diagnostic tests, even in the case of physician-confirmed ILI
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome is physician-confirmed ILI.
Incomplete outcome data (attrition bias) All outcomes	Low risk	"No students were lost to follow-up or discontinued the intervention during the study period."
Selective reporting (reporting bias)	Low risk	All outcomes were reported.

Priest 2014
Study characteristics

Methods	A cluster-RCT
---------	---------------

Priest 2014 (Continued)

Participants	<p>Study included children aged 5 to 11 years at 68 primary schools in New Zealand. Schools were randomised to hand sanitiser + education session arm (34 schools and 8859 children) and education session arm (34 schools and 7386 children).</p> <p>Inclusion criteria:</p> <p>School-level inclusion: at least 100 children of primary school age (school years 1 to 6; children will generally range in age from 5 years to 11 years) at November 2008. Schools that are not currently using hand sanitiser products or are willing to not use them for the period of the trial. Schools are within the City boundaries of Christchurch, Dunedin, or Invercargill in New Zealand. The principal of the school consents to the school being included in the trial. Not “special schools” (e.g. schools for children with deafness or disability) and either not currently using hand sanitiser products or willing to not use them for the period of the trial if they were randomised to the control group were eligible to participate in the trial.</p> <p>Student-level inclusion (follow-up children): children were eligible to participate in the follow-up group, for whom more detailed information on absences was collected, if they attended a school year 1 to 6 class in 1 of the included schools at the beginning of the second school term in 2009 (the end of April), and their caregivers completed the consent form indicating that they were willing to be telephoned following their child’s absences and that they were able to take part in telephone interviews in English</p> <p>Exclusion criteria:</p> <p>School-level exclusion: special needs schools</p> <p>Student-level exclusion (follow-up children): children of the principal investigators and study personnel of the trial. Or, children of families that the principal of the primary school directs us not to approach</p>
Interventions	<p>Hand sanitiser provision (in addition to hand hygiene education session also provided to control group) in schoolchildren. See Table 1 for details.</p>
Outcomes	<p>Laboratory: none</p> <p>Effectiveness:</p> <p>Primary outcome: the incidence rate of absence episodes from school (reported by the parents during telephone calls) due to any illness during the study period (winter term)</p> <p>Secondary outcomes: assessing whether hand sanitiser was effective in reducing the:</p> <ol style="list-style-type: none"> 1. incidence rate of respiratory illness absence episodes, 2. incidence rate of gastrointestinal illness absence episodes, 3. incidence rate of absence for any reason, 4. length of illness episode, 5. length of illness absence episode, and 6. incidence rate of subsequent illness amongst other children or adults in the household. <p>Definition of respiratory illness: at least 2 of the following caregiver-reported symptoms for 1 day, or 1 of the following symptoms for 2 days (but not fever alone): runny nose, stuffy or blocked nose or noisy breathing, cough, fever, sore throat, or sneezing</p> <p>Safety: examined whether the use of hand sanitiser was associated with an increased risk of any skin reactions during the intervention period. Skin reactions: dryness, redness, flakiness, itchiness, eczema, and any other skin reactions</p>
Notes	<p>The period study conducted: 27 April to 25 September 2009</p> <p>Funding: government (Health Research Council of New Zealand)</p>

Priest 2014 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stata/MP 10.1 for Windows was used to generate the random numbers"
Allocation concealment (selection bias)	Low risk	Done by trial statistician provided with school codes and district and randomised the schools to either "A" or "B"
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Outcome assessors were blinded to the group allocation until the analysis was completed.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors were blinded to the group allocation until the analysis was completed.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	The flow diagram gives a clear account on follow-up, with numbers of those lost to follow-up and those who discontinued the intervention along with the reasons for doing so. No child was excluded from the analysis. Only PP analysis was reported.
Selective reporting (reporting bias)	Low risk	All outcomes stated in the published protocol were reported in the study. The exception was "1 planned secondary outcome (that is irrelevant to our study) that was not collected and 2 collected secondary outcomes that were not planned in the original protocol".

Priest 2014
Study characteristics

Methods	A cluster-RCT
Participants	<p>Study included children aged 5 to 11 years at 68 primary schools in New Zealand. Schools were randomised to hand sanitiser + education session arm (34 schools and 8859 children) and education session arm (34 schools and 7386 children).</p> <p>Inclusion criteria:</p> <p>School-level inclusion: at least 100 children of primary school age (school years 1 to 6; children will generally range in age from 5 years to 11 years) at November 2008. Schools that are not currently using hand sanitiser products or are willing to not use them for the period of the trial. Schools are within the City boundaries of Christchurch, Dunedin, or Invercargill in New Zealand. The principal of the school consents to the school being included in the trial. Not "special schools" (e.g. schools for children with deafness or disability) and either not currently using hand sanitiser products or willing to not use them for the period of the trial if they were randomised to the control group were eligible to participate in the trial.</p> <p>Student-level inclusion (follow-up children): children were eligible to participate in the follow-up group, for whom more detailed information on absences was collected, if they attended a school year 1 to 6 class in 1 of the included schools at the beginning of the second school term in 2009 (the end of April), and their caregivers completed the consent form indicating that they were willing to be telephoned following their child's absences and that they were able to take part in telephone interviews in English</p>

Priest 2014 (Continued)

Exclusion criteria:

School-level exclusion: special needs schools

Student-level exclusion (follow-up children): children of the principal investigators and study personnel of the trial. Or, children of families that the principal of the primary school directs us not to approach

Interventions	Hand sanitiser provision (in addition to hand hygiene education session also provided to control group) in schoolchildren. See Table 1 for details.
Outcomes	<p>Laboratory: none</p> <p>Effectiveness:</p> <p>Primary outcome: the incidence rate of absence episodes from school (reported by the parents during telephone calls) due to any illness during the study period (winter term)</p> <p>Secondary outcomes: assessing whether hand sanitiser was effective in reducing the:</p> <ol style="list-style-type: none"> 1. incidence rate of respiratory illness absence episodes, 2. incidence rate of gastrointestinal illness absence episodes, 3. incidence rate of absence for any reason, 4. length of illness episode, 5. length of illness absence episode, and 6. incidence rate of subsequent illness amongst other children or adults in the household. <p>Definition of respiratory illness: at least 2 of the following caregiver-reported symptoms for 1 day, or 1 of the following symptoms for 2 days (but not fever alone): runny nose, stuffy or blocked nose or noisy breathing, cough, fever, sore throat, or sneezing</p> <p>Safety: examined whether the use of hand sanitiser was associated with an increased risk of any skin reactions during the intervention period. Skin reactions: dryness, redness, flakiness, itchiness, eczema, and any other skin reactions</p>
Notes	<p>The period study conducted: 27 April to 25 September 2009</p> <p>Funding: government (Health Research Council of New Zealand)</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stata/MP 10.1 for Windows was used to generate the random numbers"
Allocation concealment (selection bias)	Low risk	Done by trial statistician provided with school codes and district and randomised the schools to either "A" or "B"
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Outcome assessors were blinded to the group allocation until the analysis was completed.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome assessors were blinded to the group allocation until the analysis was completed.
Incomplete outcome data (attrition bias)	Unclear risk	The flow diagram gives a clear account on follow-up, with numbers of those lost to follow-up and those who discontinued the intervention along with the

Priest 2014 (Continued)

All outcomes		reasons for doing so. No child was excluded from the analysis. Only PP analysis was reported.
Selective reporting (reporting bias)	Low risk	All outcomes stated in the published protocol were reported in the study. The exception was "1 planned secondary outcome (that is irrelevant to our study) that was not collected and 2 collected secondary outcomes that were not planned in the original protocol".

Radonovich 2019
Study characteristics

Methods	Cluster-RCT, multicentre, pragmatic effectiveness trial
Participants	<p>Study included 280 clusters randomly assigned to N95 respirators (189 clusters and 1993 HCPs) and medical masks (191 clusters and 2058 HCPs).</p> <p>All participants in a cluster worked in the same outpatient clinic or outpatient setting. All participants were permitted to participate for 1 or more years and gave written consent for each year of participation.</p> <p>Inclusion criteria: healthcare workers in outpatient settings serving adult and paediatric patients with a high prevalence of acute respiratory illness. Participants were aged at least 18 years and employed at 1 of the 7 participating health systems, and self-identified as routinely positioned within 6 feet (1.83 m) of patients. Participants were full-time employees (defined as direct patient care for approximately ≥ 24 hours weekly) and worked primarily at the study site (defined as $\geq 75\%$ of working hours).</p> <p>Exclusion criteria: medical conditions precluding safe participation or anatomic features that could interfere with respirator fit, such as facial hair or third-trimester pregnancy. Participants self-identified race and sex using fixed categories; these variables were collected because facial anthropometrics related to race and sex may influence N95 respirator fit.</p>
Interventions	Fit-tested N95 respirators versus medical masks when near patients with respiratory illness. See Table 1 for details.
Outcomes	<p>Laboratory. Primary outcome: the incidence of laboratory-confirmed influenza, defined as:</p> <ol style="list-style-type: none"> 1. detection of influenza A or B virus by RT-PCR in an upper respiratory specimen collected within 7 days of symptom onset; 2. detection of influenza from a randomly obtained swab from an asymptomatic participant; and 3. influenza seroconversion (symptomatic or asymptomatic), defined as at least a 4-fold rise in haemagglutination inhibition antibody titres to influenza A or B virus between pre-season and post-season serological samples deemed not attributable to vaccination. <p>Effectiveness. Secondary outcomes: the incidence of 4 measures of viral respiratory illness or infection as follows:</p> <ol style="list-style-type: none"> 1. acute respiratory illness with or without laboratory confirmation; 2. laboratory-detected respiratory infection, defined as detection of a respiratory pathogen by PCR or serological evidence of infection with a respiratory pathogen during the study surveillance period(s), which was added to the protocol prior to data analysis; 3. laboratory-confirmed respiratory illness, identified as previously described (defined as self-reported acute respiratory illness plus the presence of at least PCR-confirmed viral pathogen in a specimen collected from the upper respiratory tract within 7 days of the reported symptoms and/or at least a 4-fold rise from pre-intervention to post-intervention serum antibody titres to influenza A or B virus; and 4. influenza-like illness, defined as temperature of at least 100 °F (37.8 °C) plus cough and/or a sore throat, with or without laboratory confirmation.

Radonovich 2019 (Continued)

Safety: no serious study-related adverse events were reported. 19 participants reported skin irritation or worsening acne during years 3 and 4 at 1 site in the N95 respirator group.

Notes

The study was conducted from September 2011 to May 2015, with final follow-up on 28 June 2016.

Funding: government

Compliance: adherence was reported on daily surveys 22,330 times in the N95 respirator group and 23,315 times in the medical mask group. "Always" was reported 14,566 (65.2%) times in the N95 respirator group and 15,186 (65.1%) times in the medical mask group; "sometimes" 5407 (24.2%) times in the N95 respirator group and 5853 (25.1%) times in the medical mask group; "never" 2272 (10.2%) times in the N95 respirator group and 2207 (9.5%) times in the medical mask group; and "did not recall" 85 (0.4%) times in the N95 respirator group and 69 (0.3%) times in the medical mask group. Participant-reported adherence could not be assessed in 784 participants (31.2%) in the N95 respirator group and 822 (30.8%) in the medical mask group ($P = 0.84$) because of lack of response to surveys or lack of adherence opportunities (i.e. participants did not encounter an individual with respiratory signs or symptoms). Analysed post hoc, participant adherence was reported as always or sometimes 89.4% of the time in the N95 respirator group and 90.2% of the time in the medical mask group.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated random sequences by an individual not involved in the study implementation and data analyses. Used stratified randomisation
Allocation concealment (selection bias)	Low risk	Used constrained randomisation
Blinding of participants and personnel (performance bias) All outcomes	Low risk	The participants cannot be blinded, but it seems that all the measures otherwise were the same with meticulous follow-up. Besides, the primary outcome was lab based (an objective outcome), which is unlikely to be affected by lack of blinding. Investigators were blinded to the randomisation until completion of the study and analysis.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Primary outcome is laboratory-confirmed diagnosis.
Incomplete outcome data (attrition bias) All outcomes	Low risk	"Missing outcomes were imputed using standard multiple imputation techniques, creating multiple imputed data sets with no missing values for each analysis"
Selective reporting (reporting bias)	Low risk	Reported study outcomes matched the published protocol. Every outcome was accounted for.

Radonovich 2019
Study characteristics

Methods	Cluster-RCT, multicentre, pragmatic effectiveness trial
Participants	Study included 280 clusters randomly assigned to N95 respirators (189 clusters and 1993 HCPs) and medical masks (191 clusters and 2058 HCPs).

Radonovich 2019 (Continued)

All participants in a cluster worked in the same outpatient clinic or outpatient setting. All participants were permitted to participate for 1 or more years and gave written consent for each year of participation.

Inclusion criteria: healthcare workers in outpatient settings serving adult and paediatric patients with a high prevalence of acute respiratory illness. Participants were aged at least 18 years and employed at 1 of the 7 participating health systems, and self-identified as routinely positioned within 6 feet (1.83 m) of patients. Participants were full-time employees (defined as direct patient care for approximately ≥ 24 hours weekly) and worked primarily at the study site (defined as $\geq 75\%$ of working hours).
Exclusion criteria: medical conditions precluding safe participation or anatomic features that could interfere with respirator fit, such as facial hair or third-trimester pregnancy. Participants self-identified race and sex using fixed categories; these variables were collected because facial anthropometrics related to race and sex may influence N95 respirator fit.

Interventions	Fit-tested N95 respirators versus medical masks when near patients with respiratory illness. See Table 1 for details.
Outcomes	<p>Laboratory. Primary outcome: the incidence of laboratory-confirmed influenza, defined as:</p> <ol style="list-style-type: none"> 1. detection of influenza A or B virus by RT-PCR in an upper respiratory specimen collected within 7 days of symptom onset; 2. detection of influenza from a randomly obtained swab from an asymptomatic participant; and 3. influenza seroconversion (symptomatic or asymptomatic), defined as at least a 4-fold rise in haemagglutination inhibition antibody titres to influenza A or B virus between pre-season and postseason serological samples deemed not attributable to vaccination. <p>Effectiveness. Secondary outcomes: the incidence of 4 measures of viral respiratory illness or infection as follows:</p> <ol style="list-style-type: none"> 1. acute respiratory illness with or without laboratory confirmation; 2. laboratory-detected respiratory infection, defined as detection of a respiratory pathogen by PCR or serological evidence of infection with a respiratory pathogen during the study surveillance period(s), which was added to the protocol prior to data analysis; 3. laboratory-confirmed respiratory illness, identified as previously described (defined as self-reported acute respiratory illness plus the presence of at least PCR-confirmed viral pathogen in a specimen collected from the upper respiratory tract within 7 days of the reported symptoms and/or at least a 4-fold rise from pre-intervention to postintervention serum antibody titres to influenza A or B virus; and 4. influenza-like illness, defined as temperature of at least 100 °F (37.8 °C) plus cough and/or a sore throat, with or without laboratory confirmation. <p>Safety: no serious study-related adverse events were reported. 19 participants reported skin irritation or worsening acne during years 3 and 4 at 1 site in the N95 respirator group.</p>
Notes	<p>The study was conducted from September 2011 to May 2015, with final follow-up on 28 June 2016.</p> <p>Funding: government</p> <p>Compliance: adherence was reported on daily surveys 22,330 times in the N95 respirator group and 23,315 times in the medical mask group. “Always” was reported 14,566 (65.2%) times in the N95 respirator group and 15,186 (65.1%) times in the medical mask group; “sometimes” 5407 (24.2%) times in the N95 respirator group and 5853 (25.1%) times in the medical mask group; “never” 2272 (10.2%) times in the N95 respirator group and 2207 (9.5%) times in the medical mask group; and “did not recall” 85 (0.4%) times in the N95 respirator group and 69 (0.3%) times in the medical mask group. Participant-reported adherence could not be assessed in 784 participants (31.2%) in the N95 respirator group and 822 (30.8%) in the medical mask group ($P = 0.84$) because of lack of response to surveys or lack of adherence opportunities (i.e. participants did not encounter an individual with respiratory signs or symptoms). Analysed post hoc, participant adherence was reported as always or sometimes 89.4% of the time in the N95 respirator group and 90.2% of the time in the medical mask group.</p>

Risk of bias

Radonovich 2019 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated random sequences by an individual not involved in the study implementation and data analyses. Used stratified randomisation
Allocation concealment (selection bias)	Low risk	Used constrained randomisation
Blinding of participants and personnel (performance bias) All outcomes	Low risk	The participants cannot be blinded, but it seems that all the measures otherwise were the same with meticulous follow-up. Besides, the primary outcome was lab based (an objective outcome), which is unlikely to be affected by lack of blinding. Investigators were blinded to the randomisation until completion of the study and analysis.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Primary outcome is laboratory-confirmed diagnosis.
Incomplete outcome data (attrition bias) All outcomes	Low risk	"Missing outcomes were imputed using standard multiple imputation techniques, creating multiple imputed data sets with no missing values for each analysis"
Selective reporting (reporting bias)	Low risk	Reported study outcomes matched the published protocol. Every outcome was accounted for.

Ram 2015
Study characteristics

Methods	RCT
Participants	<p>377 household compounds (index cases) completed the study. Control arm has 184 compounds with 1607 contacts, and intervention group has 193 compounds with 1814 contacts. Final analysis was performed on 193 index cases and 1661 contacts in the intervention group and 184 index cases and 1498 contacts in the control group.</p> <p>In 2009, index case-patients with symptom onset within 7 days preceding enrolment were eligible. Eligibility criteria changed in 2010 to include index case-patient with symptom onset within 48 hours preceding enrolment.</p> <p>Inclusion criteria:</p> <ol style="list-style-type: none"> 1. Individuals ≥ 5 years old: ILI, defined as history of fever and either cough or sore throat with fever onset within the previous 24 hours. 2. Individuals < 5 years old: any child with acute fever with onset within the previous 24 hours. 3. Return to home within 24 hours of presentation to Upazilla Health Complex, Jahurul Islam Medical College Hospital or the local pharmacies, i.e. the index case cannot be admitted for treatment. If admitted, the patient would not be eligible. 4. No fever in any bari resident during the 7 days preceding the patient's presentation to hospital (see definition below). 5. At least 2 individuals (in addition to the index case-patient) who intend to reside in the bari during the subsequent 20 days. 6. Residence within 30 minutes travel time (1-way) from the Upazilla Health Complex or Jahurul Islam Medical College Hospital or the local pharmacy. <p>Exclusion criteria: compounds were excluded if any compound member(s) was reported to have fever within 3 days before index case-patient enrolment. At another time point, compounds were excluded</p>

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Ram 2015 (Continued)

if any primary household member was reported to have fever (fever occurring within 48 hours prior to enrolment recorded).

Interventions	Promoting intensive hand-washing in households to prevent transmission of ILI. See Table 1 for details.
Outcomes	<p>Laboratory: PCR for influenza A and B, with further subtyping of influenza A isolates for all ILI amongst contacts</p> <p>Effectiveness: incidence of ILI. An age-based definition of ILI was used as follows.</p> <ol style="list-style-type: none"> 1. For individuals > 5 years old, ILI was defined as history of fever with cough or sore throat. 2. For children < 5 years old, ILI was defined as fever (the authors used this relatively liberal case definition in order to include influenza cases with atypical presentations in children). <p>Safety: no safety data planned or reported by investigators</p>
Notes	<p>Inclusion/exclusion criteria changed 3 times during the study conduct.</p> <p>The period study conducted: June 2009 to December 2010</p> <p>Funding: government</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Block randomisation, with a block size of 4, in order to promote random and even allocation of household compounds to the 2 treatment arms. The list of random assignments was generated by an investigator with no contact with the participants.
Allocation concealment (selection bias)	Low risk	Once baseline data collection was complete, the data collector notified the field research officer, who consulted the block randomisation list to make the assignment of the household compound to intervention or control.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Relied on symptom reporting from the head of family. Inclusion/exclusion criteria changed 3 times during the study conduct. Given the provision of a hand-washing station as part of the intervention, it was not possible to ensure blinding of participants, intervention staff, or data collectors.
Blinding of outcome assessment (detection bias) All outcomes	High risk	Relied on symptom reporting from the head of family. Inclusion/exclusion criteria changed 3 times during the conduct of the study. Given the provision of a hand-washing station as part of the intervention, it was not possible to ensure blinding of participants, intervention staff, or data collectors.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Flow chart followed all households and individuals from recruitment to analysis.
Selective reporting (reporting bias)	Low risk	The specified outcomes are clearly accounted for. Investigators report all outcomes for each modified enrolment.

Ram 2015
Study characteristics
Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Ram 2015 (Continued)

Methods	RCT
Participants	<p>377 household compounds (index cases) completed the study. Control arm has 184 compounds with 1607 contacts, and intervention group has 193 compounds with 1814 contacts. Final analysis was performed on 193 index cases and 1661 contacts in the intervention group and 184 index cases and 1498 contacts in the control group.</p> <p>In 2009, index case-patients with symptom onset within 7 days preceding enrolment were eligible. Eligibility criteria changed in 2010 to include index case-patient with symptom onset within 48 hours preceding enrolment.</p> <p>Inclusion criteria:</p> <ol style="list-style-type: none"> 1. Individuals ≥ 5 years old: ILI, defined as history of fever and either cough or sore throat with fever onset within the previous 24 hours. 2. Individuals < 5 years old: any child with acute fever with onset within the previous 24 hours. 3. Return to home within 24 hours of presentation to Upazilla Health Complex, Jahurul Islam Medical College Hospital or the local pharmacies, i.e. the index case cannot be admitted for treatment. If admitted, the patient would not be eligible. 4. No fever in any bari resident during the 7 days preceding the patient's presentation to hospital (see definition below). 5. At least 2 individuals (in addition to the index case-patient) who intend to reside in the bari during the subsequent 20 days. 6. Residence within 30 minutes travel time (1-way) from the Upazilla Health Complex or Jahurul Islam Medical College Hospital or the local pharmacy. <p>Exclusion criteria: compounds were excluded if any compound member(s) was reported to have fever within 3 days before index case-patient enrolment. At another time point, compounds were excluded if any primary household member was reported to have fever (fever occurring within 48 hours prior to enrolment recorded).</p>
Interventions	Promoting intensive hand-washing in households to prevent transmission of ILI. See Table 1 for details.
Outcomes	<p>Laboratory: PCR for influenza A and B, with further subtyping of influenza A isolates for all ILI amongst contacts</p> <p>Effectiveness: incidence of ILI. An age-based definition of ILI was used as follows.</p> <ol style="list-style-type: none"> 1. For individuals > 5 years old, ILI was defined as history of fever with cough or sore throat. 2. For children < 5 years old, ILI was defined as fever (the authors used this relatively liberal case definition in order to include influenza cases with atypical presentations in children). <p>Safety: no safety data planned or reported by investigators</p>
Notes	<p>Inclusion/exclusion criteria changed 3 times during the study conduct.</p> <p>The period study conducted: June 2009 to December 2010</p> <p>Funding: government</p>
Risk of bias	
Bias	Authors' judgement Support for judgement
Random sequence generation (selection bias)	<p>Low risk</p> <p>Block randomisation, with a block size of 4, in order to promote random and even allocation of household compounds to the 2 treatment arms. The list of random assignments was generated by an investigator with no contact with the participants.</p>

Ram 2015 (Continued)

Allocation concealment (selection bias)	Low risk	Once baseline data collection was complete, the data collector notified the field research officer, who consulted the block randomisation list to make the assignment of the household compound to intervention or control.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Relied on symptom reporting from the head of family. Inclusion/exclusion criteria changed 3 times during the study conduct. Given the provision of a hand-washing station as part of the intervention, it was not possible to ensure blinding of participants, intervention staff, or data collectors.
Blinding of outcome assessment (detection bias) All outcomes	High risk	Relied on symptom reporting from the head of family. Inclusion/exclusion criteria changed 3 times during the conduct of the study. Given the provision of a hand-washing station as part of the intervention, it was not possible to ensure blinding of participants, intervention staff, or data collectors.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Flow chart followed all households and individuals from recruitment to analysis.
Selective reporting (reporting bias)	Low risk	The specified outcomes are clearly accounted for. Investigators report all outcomes for each modified enrolment.

Roberts 2000
Study characteristics

Methods	Open cluster-RCT carried out between March and November 1996 (the Southern Hemisphere winter season) in 23 childcare centres caring for a minimum of 50 children 10 hours a day, 5 days a week in Australia. The study assessed the effects of an Australian national hand-washing programme compared to standard procedure. Randomisation was according to a random-number table, and cluster coefficients are reported.
Participants	Children (299 in the intervention arm and 259 in the control arm) aged 3 or younger attending the centres at least 3 days a week. Attrition was 51 children in the intervention arm and 72 children in the control arm due mainly to staff leaving the centres.
Interventions	Hand-washing programme with training for staff and children. It is unclear whether any extra hand-cleansing agents were used, as GloGerm (?) is mentioned when it was used in a preliminary study.
Outcomes	Laboratory: N/A Effectiveness: ARI (runny nose, cough, and blocked nose) Follow-up was via a parental phone interview every 2 weeks. Safety: N/A
Notes	Risk of bias: low (cluster coefficients and analysis by unit of randomisation) Note: the authors conclude that although there was no overall decrease in respiratory illness (RR 0.95, 95% CI 0.89 to 1.01), in children up to 24 months the decrease was statistically significant (RR 0.90, 95% CI 0.83 to 0.97). The authors speculated that this was because maximum benefits are likely from this age group due to their limited ability to wipe their nose and hands without a structured programme. Analyses by 3 compliance levels are also reported. A so-so reported and well-conducted trial

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Roberts 2000 (Continued)

Random sequence generation (selection bias)	Low risk	Randomisation was according to a random-number table.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	It was not possible to blind the intervention.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"The observer was not informed of the content of the training sessions or the intervention status of the centres."
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Recruitment rate 88% (23 of 26 CCCs); loss to follow-up not clear, as no denominator given
Selective reporting (reporting bias)	Low risk	Centres were comparable at baseline.

Sandora 2005
Study characteristics

Methods	Single-blind, cluster-RCT carried around the Boston area, USA, in the period of November 2002 to April 2003. The trial tested the effects of using a hand sanitiser and a programme of instruction on the transmissions of GI infections (data not extracted) and ARI in families. Units of randomisation were child-care centres and were carried out on enrolment by an investigator using random block size generated by computer. Assignment was single-blind (i.e. investigator blinded to the status of the centre). Cluster correlation was 0.01.
Participants	292 families with 1 or more children aged 6 months to 5 years who were in child care for 10 or more hours a week 155 children in 14 centres were allocated to the intervention arm and 137 children in 12 centres to the control arm. The mean age was 3 to 2.7 years. Attrition was respectively 15 (3 lost to follow-up and 12 who discontinued the intervention) and 19 (8 lost to follow-up and 11 who discontinued the intervention). ITT analysis was carried out.
Interventions	Alcohol-based hand sanitiser with biweekly hand hygiene educational materials over 5 months versus biweekly educational material on healthy diet
Outcomes	Effectiveness: ARI (2 of the following symptoms for 1 day or 1 of the following symptoms for 2 days: runny nose, cough, sneezing, stuffy or blocked nose, fever, sore throat). An illness episode had to be separated by 2 symptom-free days from a previous episode. A secondary illness was when it followed a similar illness in another family member by 2 to 7 days. Follow-up was by means of biweekly phone calls to caregivers. Safety: dry skin (71 reports), stinging (11 reports), bad smell (7 reports), dislike (2 reports), allergic reaction (2 reports), slippery feel (1 report), and irritation (20 reports)
Notes	Risk of bias: low Note: the authors conclude that although the rate of GI illnesses was significantly lower in the intervention group, the IRR was not significantly different for ARI (0.97, 95% CI 0.72 to 1.30). Compliance and droplet route spread may account for this apparent lack of effect. A well-reported trial

Sandora 2005 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Random assignments were generated by computer using a permuted-blocks design with random block sizes."
Allocation concealment (selection bias)	Low risk	"Assignments were concealed in opaque envelopes, and centers were assigned to control or intervention groups by a study investigator as they were enrolled."
Blinding of participants and personnel (performance bias) All outcomes	High risk	"Teachers in the intervention classrooms were responsible for encouraging the use of the disinfecting wipes and hand sanitizer according to the study protocol ... Given that no placebo was provided and sanitizer use was recorded, neither families nor data collectors could be blinded as to the group assignment of the family."
Blinding of outcome assessment (detection bias) All outcomes	High risk	"Given that no placebo was provided and sanitizer use was recorded, neither families nor data collectors could be blinded as to the group assignment of the family."
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition was 15 in intervention arm (3 lost to follow-up and 12 who discontinued the intervention) and 19 in the control arm (8 lost to follow-up and 11 who discontinued the intervention). ITT analysis was carried out.
Selective reporting (reporting bias)	Unclear risk	Well-reported

Sandora 2008
Study characteristics

Methods	Cluster-RCT carried out in a single elementary school system located in Avon, Ohio, USA to assess the effectiveness of a multifactorial infection-control intervention, including alcohol-based hand sanitiser and surface disinfection, in reducing absenteeism caused by gastrointestinal and respiratory illnesses amongst elementary school students. The study also aimed to describe the viral and bacterial contamination of common surfaces in the school classroom and to assess the impact of an environmental disinfectant on the presence of selected viruses and bacteria on these surfaces. Clustering was described as "teams of 3-4 classes depending on the class year".
Participants	<p>A total of 363 students in 15 different classrooms were eligible to participate and received letters about the study.</p> <p>A sample of 285 of these students provided written informed consent and were randomly assigned to the intervention group (146) or to the control group (139) and contributed to final analysis.</p> <p>No students were lost to follow-up or discontinued the intervention during the study period.</p> <p>Baseline demographic characteristics were similar in the intervention and control groups. Most families were white and non-Hispanic and in excellent or very good health at baseline.</p>
Interventions	Alcohol-based hand sanitiser to use at school and quaternary ammonium wipes to disinfect classroom surfaces daily for 8 weeks versus usual hand-washing and cleaning practices
Outcomes	<p>Laboratory:</p> <p>Serological evidence: no</p> <p>Swabs for bacteria and viruses from 3 types of classroom surfaces were taken.</p>

Sandora 2008 (Continued)

Effectiveness:

Respiratory illness defined as days absent as measured by a (blinded) school worker who routinely recorded reason for absenteeism either for gastrointestinal or respiratory causes.

Safety: N/A

Notes

The authors conclude that the multifaceted intervention that included alcohol-based hand sanitiser use and disinfection of common classroom surfaces reduced absenteeism from gastrointestinal illness amongst elementary school students. The intervention did not impact on absenteeism from respiratory illness. In addition, norovirus was detected less frequently on classroom surfaces in the group receiving the intervention. The study is of good quality with low risk of bias. The authors checked compliance by counting discarded wipes. Reasons given for the apparent lack of effect against ARIs but good effect on GI illness are that disinfecting the classroom surfaces (daily at lunchtime with alkali) was important, as were the alcohol wipes. The authors measured the norovirus concentration on surfaces and found this to be reduced. Other reasons may be that droplets are not affected by this method, or that contamination of hands by respiratory infections is likely to be continuous (in orofaecal transmission is mostly at the time of defecation).

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"The allocation sequence was generated by computer ..."
Allocation concealment (selection bias)	Unclear risk	"...and teams were assigned to study groups by a study investigator (Dr Shih)." Blinding of allocation cannot be guaranteed.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"All of the students absences were recorded in the usual fashion by the school employee who normally answers this dedicated telephone line. This employee was blinded to the group assignment of the child."
Incomplete outcome data (attrition bias) All outcomes	Low risk	No students were lost to follow-up or discontinued the intervention during the study period.
Selective reporting (reporting bias)	Unclear risk	Well-reported

Satomura 2005
Study characteristics
Methods

RCT. Randomisation was achieved by simple computer-generated random digit. Allocation was concealed using sealed, opaque envelopes. Not clear if there was a central randomisation centre. Post hoc exchange of envelopes was prevented by writing both the name of each participant and the number on the envelope he/she drew before breaking the seal. Participants were not blinded to the intervention; however, disease incidence was determined by 1 study physician who was not informed of the results of assignment. Analysis was done based on the intention-to-treat principle. The study targeted community healthcare all over Japan and was conducted between December 2002 and March 2003 for a follow-up period of 60 days.

Satomura 2005 (Continued)

Participants	<p>387 participants at 18 sites were recruited, 384 were included in the analysis: water gargling (N = 122), povidone-iodine gargling (N = 132), and control (N = 130).</p> <p>Follow-up was completed on 338 participants. Attrition was fully explained for URTI analysis; however, 2 participants were not accounted for in the ILI analysis. 46 participants did not complete the follow-up due to either discontinuation of diary use (n = 9) or contracting ILI (n = 37). Of the 37 participants with ILI, 11 were in the povidone-iodine group, 12 in the water group, and 14 in the control group. Analysis was performed on 35 participants (Kitamura 2007).</p>
Interventions	<p>Participants were randomised to 1 of the following: water gargling, n = 122 (20 mL of water for about 15 seconds 3 times consecutively, at least 3 times a day); povidone-iodine gargling, n = 133 (20 mL of 15 to 30 times diluted 7% povidone-iodine (as indicated by the manufacturer) in the same way as water gargling); and control, n = 132 (retain their previous gargling habits).</p> <p>All groups were asked to fill a daily gargling diary (standardised form to record: gargling habits, hand-washing, and influenza complaints).</p> <p>The frequency of gargling in the water group was higher (3.6); the frequency of hand-washing was similar amongst the 3 groups.</p> <p>URTI symptom was classified according to Jackson methods. Diary recording was continued throughout the follow-up period and for 1 week after the onset of URTI.</p> <p>ILI was reported separately.</p>
Outcomes	<p>Laboratory: none</p> <p>Effectiveness:</p> <p>Primary outcome: incidence of first URTI. Index cases were defined as all of the following conditions:</p> <ol style="list-style-type: none"> 1. both nasal and pharyngeal symptoms, 2. severity of at least 1 symptom increased by 2 grades or more, and 3. worsening of a symptom of 1 increment or more for > 3 days. <p>Secondary outcome: severity of URTI of the incident cases was assessed by grading each symptom during the initial 7 days after the onset of URTI in numeric scores: none = 0, mild = 1, moderate = 2, and severe = 3</p> <p>ILI was defined as both developing a fever of 38 °C or higher and worsening arthralgia in addition to some respiratory symptoms (Kitamura 2007).</p> <p>Safety: no harm was reported. However, 2 participants in the povidone-iodine group switched to water gargling (analysed in their assignment group).</p>
Notes	<p>The authors concluded that simple water gargling is effective in preventing URTIs amongst healthy people. However, no statistically significant difference was observed against ILIs.</p> <p>The study was well-conducted; blinding would have added to the validity of the results. In addition, the study was not powered enough to detect a statistically significant preventative effect against ILI.</p> <p>The study demonstrates that in addition to hand-washing, simple gargling even with water can reduce URTI, but not ILI. However, during periods of endemic influenza, multiple inexpensive and simple modalities (hand-washing, masks, gargling) can be utilised together to reduce infection and transmission.</p> <p>Overall, the reporting of the 2 combined studies together is highly confusing. In the first study (Satomura 2005), the main outcome is URTI defined as fever and arthralgia. The second study (which is a presentation of further data from the 2005 publication in the guise of a short report) introduces the outcome ILI with a definition similar to that of URTI in the first study but referring to the earlier outcome as common cold. Also of note is reporting of significance without confidence intervals. Overall, this potentially important study should be repeated with a larger denominator.</p> <p>Unclear risk of bias because of confused reporting and absence of double-blinding</p>
Risk of bias	
Bias	Authors' judgement Support for judgement

Satomura 2005 (Continued)

Random sequence generation (selection bias)	Low risk	"Group assignment was based on simple computer-generated random digits..."
Allocation concealment (selection bias)	Low risk	"By an individual drawing of sealed opaque envelopes, subjects were randomly assigned to the following three groups" "allocation was completely concealed from study administrators"
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not blinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"To prevent post hoc exchange of the envelopes, local administrators wrote down both the name of each subject and the number on the envelope he/she drew before breaking the seal."
Incomplete outcome data (attrition bias) All outcomes	Low risk	338 of 385 randomised followed up; reasons reported.
Selective reporting (reporting bias)	Unclear risk	Confusing reporting

Savolainen-Kopra 2012
Study characteristics

Methods	Open cluster-RCT, 3-arm intervention trial
Participants	<p>A total of 21 clusters (683 individuals) were randomised to implement hand hygiene with soap and water (257 individuals), alcohol-based hand rub (202 individuals), or control (224 individuals).</p> <p>The study was conducted in distinct office work units in 6 corporations in the Helsinki Region that together employed some 10,000 staff. All employees (age ≥ 18 years, both genders) were contacted by email survey.</p> <p>Inclusion criteria: "Volunteers working in defined units"</p> <p>Exclusion criteria: "Persons with open wounds or chronic eczema in hands"</p> <p>The designated 21 study clusters were identified as operationally distinct working units, each containing at least 50 people.</p>
Interventions	Hand hygiene with soap and water and standardised instructions on how to limit the transmission of infections. Usual hand hygiene (control). See Table 1 for details.
Outcomes	<p>Laboratory:</p> <p>"Between November 2008 and May 2010, the seven occupational health clinics serving the six participating corporations were advised to collect, using standard techniques, two to three respiratory samples per week from typical RTI patients and also faecal samples from a few representative patients with gastrointestinal symptoms when a GIT outbreak was suspected. The samples could originate from the study participants and also from work units not included in the study. In the laboratory, viral nucleic acids were extracted with well-characterized commercial kits and tested by validated real-time PCR methods to detect influenza A and B viruses, respiratory syncytial virus, parainfluenza virus types 1, 2, and 3, adenoviruses, human rhinoviruses and human enteroviruses from respiratory specimens, and norovirus from faecal specimens (detailed descriptions of the test procedures are available from the authors)."</p>

Savolainen-Kopra 2012 (Continued)

Effectiveness:

Predefined primary endpoints:

1. Number of reported infection episodes in a cluster per total reported weeks.
2. Number of reported sick leave episodes in a cluster per total reported weeks.

Secondary endpoints and outcome measures:

1. Number of days with reported symptoms of RTI and/or GTI in a cluster within a time frame of 100 reporting weeks.
2. Number of days-off due to own RTI or GTI in a cluster within a time frame of 100 reporting weeks.

Safety: reported 0 adverse events

Notes

The period study conducted: January 2009 to May 2010

Funding: government

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Low risk	"clusters were matched and randomized prior to onset of the interventions"
Blinding of participants and personnel (performance bias) All outcomes	High risk	The interventions were not blinded to any party involved (i.e. the study group, participants, or the occupational health services). Subjective reporting of disease episodes
Blinding of outcome assessment (detection bias) All outcomes	High risk	Subjective reporting of disease episodes
Incomplete outcome data (attrition bias) All outcomes	High risk	24% loss to follow-up. However, new recruiting in most clusters; the total number of reporting participants at the end of the trial was 91.7% compared to that at the beginning. Attrition was reported, and 76% of volunteers who started reporting continued to do so until the end of the study. Because of new recruiting in most clusters, the total number of reporting participants at the end of the trial was 626, or 91.7%, compared to that at the beginning. This means that 15.7% of the participants were replaced during the study!!! Raw data on the effects of the interventions on the occurrence of respiratory infections and vomiting/diarrhoea diseases were not reported. Zero adverse effects were reported.
Selective reporting (reporting bias)	Low risk	All planned outcomes were reported.

Roberts 2000
Study characteristics

Methods

Open cluster-RCT carried out between March and November 1996 (the Southern Hemisphere winter season) in 23 childcare centres caring for a minimum of 50 children 10 hours a day, 5 days a week in

Roberts 2000 (Continued)

Australia. The study assessed the effects of an Australian national hand-washing programme compared to standard procedure. Randomisation was according to a random-number table, and cluster coefficients are reported.

Participants	Children (299 in the intervention arm and 259 in the control arm) aged 3 or younger attending the centres at least 3 days a week. Attrition was 51 children in the intervention arm and 72 children in the control arm due mainly to staff leaving the centres.
Interventions	Hand-washing programme with training for staff and children. It is unclear whether any extra hand-cleansing agents were used, as GloGerm (?) is mentioned when it was used in a preliminary study.
Outcomes	Laboratory: N/A Effectiveness: ARI (runny nose, cough, and blocked nose) Follow-up was via a parental phone interview every 2 weeks. Safety: N/A
Notes	Risk of bias: low (cluster coefficients and analysis by unit of randomisation) Note: the authors conclude that although there was no overall decrease in respiratory illness (RR 0.95, 95% CI 0.89 to 1.01), in children up to 24 months the decrease was statistically significant (RR 0.90, 95% CI 0.83 to 0.97). The authors speculated that this was because maximum benefits are likely from this age group due to their limited ability to wipe their nose and hands without a structured programme. Analyses by 3 compliance levels are also reported. A so-so reported and well-conducted trial

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation was according to a random-number table.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	High risk	It was not possible to blind the intervention.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"The observer was not informed of the content of the training sessions or the intervention status of the centres."
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Recruitment rate 88% (23 of 26 CCCs); loss to follow-up not clear, as no denominator given
Selective reporting (reporting bias)	Low risk	Centres were comparable at baseline.

Sandora 2005
Study characteristics

Methods	Single-blind, cluster-RCT carried around the Boston area, USA, in the period of November 2002 to April 2003. The trial tested the effects of using a hand sanitiser and a programme of instruction on the transmissions of GI infections (data not extracted) and ARIs in families. Units of randomisation were child-care centres and were carried out on enrolment by an investigator using random block size generated
---------	---

Sandora 2005 (Continued)

by computer. Assignment was single-blind (i.e. investigator blinded to the status of the centre). Cluster correlation was 0.01.

Participants	<p>292 families with 1 or more children aged 6 months to 5 years who were in child care for 10 or more hours a week</p> <p>155 children in 14 centres were allocated to the intervention arm and 137 children in 12 centres to the control arm. The mean age was 3 to 2.7 years. Attrition was respectively 15 (3 lost to follow-up and 12 who discontinued the intervention) and 19 (8 lost to follow-up and 11 who discontinued the intervention). ITT analysis was carried out.</p>
Interventions	Alcohol-based hand sanitiser with biweekly hand hygiene educational materials over 5 months versus biweekly educational material on healthy diet
Outcomes	<p>Effectiveness: ARI (2 of the following symptoms for 1 day or 1 of the following symptoms for 2 days: runny nose, cough, sneezing, stuffy or blocked nose, fever, sore throat). An illness episode had to be separated by 2 symptom-free days from a previous episode. A secondary illness was when it followed a similar illness in another family member by 2 to 7 days.</p> <p>Follow-up was by means of biweekly phone calls to caregivers.</p> <p>Safety: dry skin (71 reports), stinging (11 reports), bad smell (7 reports), dislike (2 reports), allergic reaction (2 reports), slippery feel (1 report), and irritation (20 reports)</p>
Notes	<p>Risk of bias: low</p> <p>Note: the authors conclude that although the rate of GI illnesses was significantly lower in the intervention group, the IRR was not significantly different for ARIs (0.97, 95% CI 0.72 to 1.30). Compliance and droplet route spread may account for this apparent lack of effect. A well-reported trial</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Random assignments were generated by computer using a permuted-blocks design with random block sizes."
Allocation concealment (selection bias)	Low risk	"Assignments were concealed in opaque envelopes, and centers were assigned to control or intervention groups by a study investigator as they were enrolled."
Blinding of participants and personnel (performance bias) All outcomes	High risk	"Teachers in the intervention classrooms were responsible for encouraging the use of the disinfecting wipes and hand sanitizer according to the study protocol ... Given that no placebo was provided and sanitizer use was recorded, neither families nor data collectors could be blinded as to the group assignment of the family."
Blinding of outcome assessment (detection bias) All outcomes	High risk	"Given that no placebo was provided and sanitizer use was recorded, neither families nor data collectors could be blinded as to the group assignment of the family."
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition was 15 in intervention arm (3 lost to follow-up and 12 who discontinued the intervention) and 19 in the control arm (8 lost to follow-up and 11 who discontinued the intervention). ITT analysis was carried out.
Selective reporting (reporting bias)	Unclear risk	Well-reported

Sandora 2008

Study characteristics

Methods	Cluster-RCT carried out in a single elementary school system located in Avon, Ohio, USA to assess the effectiveness of a multifactorial infection-control intervention, including alcohol-based hand sanitiser and surface disinfection, in reducing absenteeism caused by gastrointestinal and respiratory illnesses amongst elementary school students. The study also aimed to describe the viral and bacterial contamination of common surfaces in the school classroom and to assess the impact of an environmental disinfectant on the presence of selected viruses and bacteria on these surfaces. Clustering was described as "teams of 3-4 classes depending on the class year".
Participants	<p>A total of 363 students in 15 different classrooms were eligible to participate and received letters about the study.</p> <p>A sample of 285 of these students provided written informed consent and were randomly assigned to the intervention group (146) or to the control group (139) and contributed to final analysis.</p> <p>No students were lost to follow-up or discontinued the intervention during the study period.</p> <p>Baseline demographic characteristics were similar in the intervention and control groups. Most families were white and non-Hispanic and in excellent or very good health at baseline.</p>
Interventions	Alcohol-based hand sanitiser to use at school and quaternary ammonium wipes to disinfect classroom surfaces daily for 8 weeks versus usual hand-washing and cleaning practices
Outcomes	<p>Laboratory: Serological evidence: no Swabs for bacteria and viruses from 3 types of classroom surfaces were taken.</p> <p>Effectiveness: Respiratory illness defined as days absent as measured by a (blinded) school worker who routinely recorded reason for absenteeism either for gastrointestinal or respiratory causes.</p> <p>Safety: N/A</p>
Notes	The authors conclude that the multifaceted intervention that included alcohol-based hand sanitiser use and disinfection of common classroom surfaces reduced absenteeism from gastrointestinal illness amongst elementary school students. The intervention did not impact on absenteeism from respiratory illness. In addition, norovirus was detected less frequently on classroom surfaces in the group receiving the intervention. The study is of good quality with low risk of bias. The authors checked compliance by counting discarded wipes. Reasons given for the apparent lack of effect against ARIs but good effect on GI illness are that disinfecting the classroom surfaces (daily at lunchtime with alkali) was important, as were the alcohol wipes. The authors measured the norovirus concentration on surfaces and found this to be reduced. Other reasons may be that droplets are not affected by this method, or that contamination of hands by respiratory infections is likely to be continuous (in orofaecal transmission is mostly at the time of defecation).

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"The allocation sequence was generated by computer ..."
Allocation concealment (selection bias)	Unclear risk	"...and teams were assigned to study groups by a study investigator (Dr Shih)." Blinding of allocation cannot be guaranteed.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not possible

Sandora 2008 (Continued)

Blinding of outcome assessment (detection bias) All outcomes	Low risk	"All of the students absences were recorded in the usual fashion by the school employee who normally answers this dedicated telephone line. This employee was blinded to the group assignment of the child."
Incomplete outcome data (attrition bias) All outcomes	Low risk	No students were lost to follow-up or discontinued the intervention during the study period.
Selective reporting (reporting bias)	Unclear risk	Well-reported

Satomura 2005
Study characteristics

Methods	<p>RCT. Randomisation was achieved by simple computer-generated random digit. Allocation was concealed using sealed, opaque envelopes. Not clear if there was a central randomisation centre. Post hoc exchange of envelopes was prevented by writing both the name of each participant and the number on the envelope he/she drew before breaking the seal. Participants were not blinded to the intervention; however, disease incidence was determined by 1 study physician who was not informed of the results of assignment. Analysis was done based on the intention-to-treat principle. The study targeted community healthcare all over Japan and was conducted between December 2002 and March 2003 for a follow-up period of 60 days.</p>
Participants	<p>387 participants at 18 sites were recruited, 384 were included in the analysis: water gargling (N = 122), povidone-iodine gargling (N = 132), and control (N = 130).</p> <p>Follow-up was completed on 338 participants. Attrition was fully explained for URTI analysis; however, 2 participants were not accounted for in the ILI analysis. 46 participants did not complete the follow-up due to either discontinuation of diary use (n = 9) or contracting ILI (n = 37). Of the 37 participants with ILI, 11 were in the povidone-iodine group, 12 in the water group, and 14 in the control group. Analysis was performed on 35 participants (Kitamura 2007).</p>
Interventions	<p>Participants were randomised to 1 of the following: water gargling, n = 122 (20 mL of water for about 15 seconds 3 times consecutively, at least 3 times a day); povidone-iodine gargling, n = 133 (20 mL of 15 to 30 times diluted 7% povidone-iodine (as indicated by the manufacturer) in the same way as water gargling); and control, n = 132 (retain their previous gargling habits).</p> <p>All groups were asked to fill a daily gargling diary (standardised form to record: gargling habits, hand-washing, and influenza complaints).</p> <p>The frequency of gargling in the water group was higher (3.6); the frequency of hand-washing was similar amongst the 3 groups.</p> <p>URTIs symptom was classified according to Jackson methods. Diary recording was continued throughout the follow-up period and for 1 week after the onset of URTI.</p> <p>ILI was reported separately.</p>
Outcomes	<p>Laboratory: none</p> <p>Effectiveness:</p> <p>Primary outcome: incidence of first URTI. Index cases were defined as all of the following conditions:</p> <ol style="list-style-type: none"> 1. both nasal and pharyngeal symptoms, 2. severity of at least 1 symptom increased by 2 grades or more, and 3. worsening of a symptom of 1 increment or more for > 3 days. <p>Secondary outcome: severity of URTI of the incident cases was assessed by grading each symptom during the initial 7 days after the onset of URTI in numeric scores: none = 0, mild = 1, moderate = 2, and severe = 3</p>

Satomura 2005 (Continued)

ILI was defined as both developing a fever of 38 °C or higher and worsening arthralgia in addition to some respiratory symptoms (Kitamura 2007).

Safety: no harm was reported. However, 2 participants in the povidone-iodine group switched to water gargling (analysed in their assignment group).

Notes	<p>The authors concluded that simple water gargling is effective in preventing URTIs amongst healthy people. However, no statistically significant difference was observed against ILIs.</p> <p>The study was well-conducted; blinding would have added to the validity of the results. In addition, the study was not powered enough to detect a statistically significant preventative effect against ILI.</p> <p>The study demonstrates that in addition to hand-washing, simple gargling even with water can reduce URTI, but not ILI. However, during periods of endemic influenza, multiple inexpensive and simple modalities (hand-washing, masks, gargling) can be utilised together to reduce infection and transmission.</p> <p>Overall, the reporting of the 2 combined studies together is highly confusing. In the first study (Satomura 2005), the main outcome is URTI defined as fever and arthralgia. The second study (which is a presentation of further data from the 2005 publication in the guise of a short report) introduces the outcome ILI with a definition similar to that of URTI in the first study but referring to the earlier outcome as common cold. Also of note is reporting of significance without confidence intervals. Overall, this potentially important study should be repeated with a larger denominator.</p> <p>Unclear risk of bias because of confused reporting and absence of double-blinding</p>
-------	--

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Group assignment was based on simple computer-generated random digits..."
Allocation concealment (selection bias)	Low risk	"By an individual drawing of sealed opaque envelopes, subjects were randomly assigned to the following three groups" "allocation was completely concealed from study administrators"
Blinding of participants and personnel (performance bias) All outcomes	High risk	Not blinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"To prevent post hoc exchange of the envelopes, local administrators wrote down both the name of each subject and the number on the envelope he/she drew before breaking the seal."
Incomplete outcome data (attrition bias) All outcomes	Low risk	338 of 385 randomised followed up; reasons reported.
Selective reporting (reporting bias)	Unclear risk	Confusing reporting

Savolainen-Kopra 2012
Study characteristics

Methods	Open cluster-RCT, 3-arm intervention trial
Participants	A total of 21 clusters (683 individuals) were randomised to implement hand hygiene with soap and water (257 individuals), alcohol-based hand rub (202 individuals), or control (224 individuals).

Savolainen-Kopra 2012 (Continued)

The study was conducted in distinct office work units in 6 corporations in the Helsinki Region that together employed some 10,000 staff. All employees (age \geq 18 years, both genders) were contacted by email survey.

Inclusion criteria: "Volunteers working in defined units"

Exclusion criteria: "Persons with open wounds or chronic eczema in hands"

The designated 21 study clusters were identified as operationally distinct working units, each containing at least 50 people.

Interventions	Hand hygiene with soap and water and standardised instructions on how to limit the transmission of infections. Usual hand hygiene (control). See Table 1 for details.
Outcomes	<p>Laboratory:</p> <p>"Between November 2008 and May 2010, the seven occupational health clinics serving the six participating corporations were advised to collect, using standard techniques, two to three respiratory samples per week from typical RTI patients and also faecal samples from a few representative patients with gastrointestinal symptoms when a GIT outbreak was suspected. The samples could originate from the study participants and also from work units not included in the study. In the laboratory, viral nucleic acids were extracted with well-characterized commercial kits and tested by validated real-time PCR methods to detect influenza A and B viruses, respiratory syncytial virus, parainfluenza virus types 1, 2, and 3, adenoviruses, human rhinoviruses and human enteroviruses from respiratory specimens, and norovirus from faecal specimens (detailed descriptions of the test procedures are available from the authors)."</p> <p>Effectiveness:</p> <p>Predefined primary endpoints:</p> <ol style="list-style-type: none"> 1. Number of reported infection episodes in a cluster per total reported weeks. 2. Number of reported sick leave episodes in a cluster per total reported weeks. <p>Secondary endpoints and outcome measures:</p> <ol style="list-style-type: none"> 1. Number of days with reported symptoms of RTI and/or GTI in a cluster within a time frame of 100 reporting weeks. 2. Number of days-off due to own RTI or GTI in a cluster within a time frame of 100 reporting weeks. <p>Safety: reported 0 adverse events</p>
Notes	<p>The period study conducted: January 2009 to May 2010</p> <p>Funding: government</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information
Allocation concealment (selection bias)	Low risk	"clusters were matched and randomized prior to onset of the interventions"
Blinding of participants and personnel (performance bias) All outcomes	High risk	The interventions were not blinded to any party involved (i.e. the study group, participants, or the occupational health services). Subjective reporting of disease episodes
Blinding of outcome assessment (detection bias)	High risk	Subjective reporting of disease episodes

Savolainen-Kopra 2012 (Continued)

All outcomes

Incomplete outcome data (attrition bias) All outcomes	High risk	24% loss to follow-up. However, new recruiting in most clusters; the total number of reporting participants at the end of the trial was 91.7% compared to that at the beginning. Attrition was reported, and 76% of volunteers who started reporting continued to do so until the end of the study. Because of new recruiting in most clusters, the total number of reporting participants at the end of the trial was 626, or 91.7%, compared to that at the beginning. This means that 15.7% of the participants were replaced during the study!!! Raw data on the effects of the interventions on the occurrence of respiratory infections and vomiting/diarrhoea diseases were not reported. Zero adverse effects were reported.
Selective reporting (reporting bias)	Low risk	All planned outcomes were reported.

Simmerman 2011
Study characteristics

Methods	Randomised controlled study
Participants	<p>Study recruited 348 households and 885 members and randomised them as follows:</p> <ol style="list-style-type: none"> Control (index household = 119, with 302 family members) Hand-washing (index household = 119, with 292 family members) Hand-washing and face mask (index household = 110, with 291 family members) <p>The household members of children (index cases) presenting with ILI at the outpatient department of the Queen Sirikit National Institute of Child Health (QSNICH) in Bangkok, the largest public paediatric hospital in Thailand</p> <p>Inclusion criteria:</p> <p>For index cases: children aged 1 month through 15 years, residents of the Bangkok metropolitan area, and had an onset of illness < 48 hours before respiratory specimens tested positive for influenza by an RIDT that was later confirmed by qualitative real-time RT-PCR (rRT-PCR)</p> <p>Eligible index cases' households must have had at least 2 other members aged ≥ 1 month who planned to sleep inside the house for a period of at least 21 days from the time of enrolment.</p> <p>Exclusion criteria:</p> <p>For index cases: children at high risk for severe influenza complications (e.g. chronic lung disease, renal disease, and long-term aspirin therapy) and those treated with influenza antiviral medications</p> <p>Excluded households: those with any member reporting an ILI that preceded the index case by 7 days or less and households where any member had received influenza vaccination during the preceding 12 months</p>
Interventions	Hand-washing, or hand-washing plus paper surgical face mask, or control. See Table 1 for details.
Outcomes	<p>Laboratory:</p> <p>To identify index cases:</p> <p>QuickVue Influenza A+B rapid diagnostic kit (Quidel Co., San Diego, CA, USA), followed by rRT-PCR for influenza viral RNA</p> <p>Index cases and contacts tested with nasal swab and throat swab both processed for rRT-PCR.</p>

Simmerman 2011 (Continued)

2 blood samples for antibody seroconversion collected on Days 1 and 21 (seroconversion defined as a fourfold rise in HI titre between paired sera for any of the antigens assayed).

Effectiveness:

Laboratory-confirmed secondary influenza virus infections amongst household members described as the secondary attack rate (SAR). A secondary influenza virus infection was defined as a positive rRT-PCR result on Days 3 or 7 or a fourfold rise in influenza HI antibody titres with the virus type and subtype matching the index case.

SAR for ILI defined by the WHO as fever plus cough or sore throat, based on self-reported symptoms.

Safety: no safety measures planned or reported by the investigators

Adherence: participants in the control arm reported an average of 3.9 hand-washing episodes/day (on Day 7), whilst participants in the hand-washing arm reported an average of 4.7 hand-washing episodes/day (95% CI 4.3 to 5.0; $P = 0.002$ compared to controls), and participants in the hand-washing plus face mask arm reported 4.9 episodes/day (95% CI 4.5 to 5.3; $P < 0.001$ compared to controls). In the intervention arms, parents had the highest reported daily hand-washing frequency (5.7, 95% CI 5.3 to 6.0) followed by others (4.8, 95% CI 4.3 to 5.3), siblings (4.3, 95% CI 3.7 to 4.8), and the index cases (4.1, 95% CI 3.8 to 4.4). There was no difference in the average amount of soap used in a week in the hand-washing arm (54 mL per person) and the hand-washing plus face mask arm (58.1 mL per person) ($P = 0.15$). 289 participants in the hand-washing plus face mask arm used an average of 12 masks per person per week (median 11, IQR 7 to 16) and reported wearing a face mask a mean of 211 minutes/day (IQR 17 to 317 minutes/day). Parents wore their masks for a median of 153 (IQR 40 to 411) minutes per day, far more than other relations (median 59; IQR 9 to 266), the index patients themselves (median 35; IQR 4 to 197), or their siblings (median 17; IQR 6 to 107). The study authors note that differences in average usage may be an attenuated measure of appropriate use in relation to the actual unmeasured exposure risk such as proximity to the index case.

Notes

The period study conducted: April 2008 and August 2009

Funding: government

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomization was achieved using a block randomization method using a list of blocks each with 12 household IDs, four of which were assigned to each of the three study arms."
Allocation concealment (selection bias)	Unclear risk	"A study coordinator assigned each household to one study arm after consent was obtained"
Blinding of participants and personnel (performance bias) All outcomes	Low risk	Recruiting clinicians were blinded to the allocation of the specific intervention. The participants were not blinded, but it is unlikely that the outcome would have been affected by lack of blinding.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	The primary outcome is a laboratory-confirmed influenza.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Household flow chart provided with reasons for exclusions, all numbers provided. Analysis was done by ITT and PP.
Selective reporting (reporting bias)	Low risk	All outcomes are accounted for in the ITT analysis of the results.

Stebbins 2011
Study characteristics

Methods	Cluster-RCT, open-label
Participants	<p>Study included 3360 students from 10 Pittsburgh elementary schools. Intervention arm (5 schools, 1695 people) and control arm (5 schools, 1665 people)</p> <p>No inclusion or exclusion criteria were provided.</p>
Interventions	Training in hand and respiratory (cough) hygiene. Hand sanitiser was provided and encouraged to be used regularly. See Table 1 for details.
Outcomes	<p>Laboratory:</p> <p>Primary outcome: laboratory-confirmed influenza (RT-PCR) amongst children presenting with ILIs leading to their absence from school</p> <p>2 nasal swabs were obtained using test manufacturer-approved sterile Dacron swabs. 1 swab was employed for influenza testing using the QuickVue Influenza A+B test (Quidel Corp, San Diego, CA).</p> <p>The second nasal swab was delivered on cold pack to the University of Pittsburgh Medical Center Clinical Virology Laboratory, Pittsburgh, PA for RT-PCR testing (performed within 48 hours). The RT-PCR used viral nucleic acid extract (EasyMag; bioMerieux, Durham, NC)</p> <p>and primer/probe sequences for influenza A, influenza B, and influenza A H1 and H3 subtypes (CDC, Atlanta GA).</p> <p>Effectiveness:</p> <p>Secondary outcome: absence episodes and cumulative days of absence due to ILI, any illness, and all causes</p> <p>Safety: none mentioned</p>
Notes	<p>The period study conducted: 1 November 2007 through 24 April 2008</p> <p>Funding: unclear</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"constrained randomization algorithm"
Allocation concealment (selection bias)	Low risk	"Random allocation of schools to two arms was created by Dr. Cummings and concealed until intervention assignment". "At the beginning of the school year parents and guardians were given the opportunity to decline participation"
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	In 76% and 78% of illness in intervention and control group were laboratory confirmed. ILI is objectively defined.

Stebbins 2011 (Continued)

Incomplete outcome data (attrition bias) All outcomes	High risk	Only episodes of identified causes were analysed. Causes of absence episodes in 66% of the study participants were not identified (2092 in the intervention group and 2232 in the control group). The parents could be contacted in only 34% cases of absence. About half of them had an illness, and in one-third of these cases the illness met the criteria of ILI (361 cases (33%)). Of these, 279 (77%) were tested for influenza.
Selective reporting (reporting bias)	Unclear risk	Insufficient information to judge

Stebbins 2011
Study characteristics

Methods	Cluster-RCT, open-label
Participants	Study included 3360 students from 10 Pittsburgh elementary schools. Intervention arm (5 schools, 1695 people) and control arm (5 schools, 1665 people) No inclusion or exclusion criteria were provided.
Interventions	Training in hand and respiratory (cough) hygiene. Hand sanitiser was provided and encouraged to be used regularly. See Table 1 for details.
Outcomes	Laboratory: Primary outcome: laboratory-confirmed influenza (RT-PCR) amongst children presenting with ILIs leading to their absence from school 2 nasal swabs were obtained using test manufacturer-approved sterile Dacron swabs. 1 swab was employed for influenza testing using the QuickVue Influenza A+B test (Quidel Corp, San Diego, CA). The second nasal swab was delivered on cold pack to the University of Pittsburgh Medical Center Clinical Virology Laboratory, Pittsburgh, PA for RT-PCR testing (performed within 48 hours). The RT-PCR used viral nucleic acid extract (EasyMag; bioMerieux, Durham, NC) and primer/probe sequences for influenza A, influenza B, and influenza A H1 and H3 subtypes (CDC, Atlanta GA). Effectiveness: Secondary outcome: absence episodes and cumulative days of absence due to ILI, any illness, and all causes Safety: none mentioned
Notes	The period study conducted: 1 November 2007 through 24 April 2008 Funding: unclear

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"constrained randomization algorithm"

Stebbins 2011 (Continued)

Allocation concealment (selection bias)	Low risk	"Random allocation of schools to two arms was created by Dr. Cummings and concealed until intervention assignment". "At the beginning of the school year parents and guardians were given the opportunity to decline participation"
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded
Blinding of outcome assessment (detection bias) All outcomes	Low risk	In 76% and 78% of illness in intervention and control group were laboratory confirmed. ILI is objectively defined.
Incomplete outcome data (attrition bias) All outcomes	High risk	Only episodes of identified causes were analysed. Causes of absence episodes in 66% of the study participants were not identified (2092 in the intervention group and 2232 in the control group). The parents could be contacted in only 34% cases of absence. About half of them had an illness, and in one-third of these cases the illness met the criteria of ILI (361 cases (33%)). Of these, 279 (77%) were tested for influenza.
Selective reporting (reporting bias)	Unclear risk	Insufficient information to judge

Suess 2012
Study characteristics

Methods	Cluster-RCT, open-label, parallel design
Participants	<p>Study sample included 84 households randomised as follows:</p> <ol style="list-style-type: none"> 1. 30 control (index cases = 30, household contact = 82) 2. 26 mask group (index cases = 26, household contact = 69) 3. 28 mask and hand hygiene group (index cases = 28, household contact = 67) <p>Inclusion criteria: patients presenting to general practitioners or family physicians at the study sites within 2 days of symptom onset; had a positive rapid antigen test for influenza (later to be confirmed by quantitative RT-PCR (qRT-PCR)); and was at least 2 years old. Index cases also had to be the only household member suffering from respiratory disease within 14 days prior to symptom onset. Exclusion criteria were pregnancy, severely reduced health status, and HIV infection. 1-person households were also not eligible or inclusion.</p>
Interventions	Quote: "facemask and practising intensified hand hygiene (MH group), wearing facemask only (M group) and none of the 2 (control group)". See Table 1 for details.
Outcomes	<p>Primary outcomes: SAR of laboratory-confirmed (qRT-PCR) influenza infection amongst household members (secondary infection cases) presenting with ILI within the observation period (8 days from the date of onset). ILI was defined as fever (> 38.0 °C) + cough or sore throat. Nasal wash specimens (or if these were not possible, nasal swabs) from all participating household members</p> <p>Effectiveness:</p> <p>Secondary outcomes: laboratory-confirmed influenza infection in a household contact (secondary infection cases). The study authors defined a symptomatic secondary influenza virus infection as a laboratory-confirmed influenza infection in a household member who developed fever (> 38.0 °C), cough, or sore throat during the observation period. They termed all other secondary cases as subclinical. A</p>

Suess 2012 (Continued)

secondary outcome measure was the occurrence of ILI as defined by WHO as fever plus cough or sore throat.

Safety: study reported that the majority of participants (107/172, 62%) did not report any problems with mask-wearing. This proportion was significantly higher in the group of adults (71/100, 71%) compared to the group of children (36/72, 50%) ($P = 0.005$). The main problem reported by participants (adults as well as children) was "heat/humidity" (18/34, 53% of children; 10/29, 35% of adults) ($P = 0.1$), followed by "pain" and "shortness of breath" when wearing a face mask.

Notes

Period study conducted: November 2009 to April 2011

Funding: governmental

Adherence: in general, daily adherence was good, reaching a plateau of over 50% in nearly all groups (M and MH groups; 2009/10 and 2010/11) from the third day on (by then the intervention had been implemented in all households). A gradual decline towards lower adherence began around the sixth day of the index patient's illness.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"prepared lists of random numbers with Microsoft Excel 2003 (Microsoft™ Cooperation, Seattle, USA) which were divided between the three intervention groups. Each participating physician received a list of random numbers with the interventions represented in a 1:1:1 ratio"
Allocation concealment (selection bias)	Low risk	"the participating physician received a list of random numbers with the interventions represented in a 1:1:1 ratio. Eligible index patients were randomly assigned a number, which was then communicated to the study center. The resulting intervention was only communicated to the households with the physicians. Intervention material was given to the study sites in closed boxes marked only with the randomisation number. Recruiting physicians were not aware of the allocation of the numbers to the interventions and the boxes for the three intervention arms looked identical. After randomisation, participants were given their box by the physician's assistants"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Outcomes are very objective and therefore unlikely to be influenced by lack of blinding. In addition, "physicians (as well as laboratory personnel) blinded from the randomisation results".
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"physicians (as well as laboratory personnel) blinded from the randomisation results". Outcomes are very objective and therefore unlikely to be influenced by lack of blinding.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up. Daily follow-up home visits over the short period of data collection (8 days)
Selective reporting (reporting bias)	Low risk	The follow-up period is very short (8 days) with very good coverage, and the criteria for defining the outcome are highly objective. All planned outcomes were reported.

Suess 2012
Study characteristics
Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Suess 2012 (Continued)

Methods	Cluster-RCT, open-label, parallel design
Participants	<p>Study sample included 84 households randomised as follows:</p> <ol style="list-style-type: none"> 1. 30 control (index cases = 30, household contact = 82) 2. 26 mask group (index cases = 26, household contact = 69) 3. 28 mask and hand hygiene group (index cases = 28, household contact = 67) <p>Inclusion criteria: patients presenting to general practitioners or family physicians at the study sites within 2 days of symptom onset; had a positive rapid antigen test for influenza (later to be confirmed by quantitative RT-PCR (qRT-PCR)); and was at least 2 years old. Index cases also had to be the only household member suffering from respiratory disease within 14 days prior to symptom onset. Exclusion criteria were pregnancy, severely reduced health status, and HIV infection. 1-person households were also not eligible or inclusion.</p>
Interventions	Quote: "facemask and practising intensified hand hygiene (MH group), wearing facemask only (M group) and none of the 2 (control group)". See Table 1 for details.
Outcomes	<p>Primary outcomes: SAR of laboratory-confirmed (qRT-PCR) influenza infection amongst household members (secondary infection cases) presenting with ILI within the observation period (8 days from the date of onset). ILI was defined as fever (> 38.0 °C) + cough or sore throat. Nasal wash specimens (or if these were not possible, nasal swabs) from all participating household members</p> <p>Effectiveness:</p> <p>Secondary outcomes: laboratory-confirmed influenza infection in a household contact (secondary infection cases). The study authors defined a symptomatic secondary influenza virus infection as a laboratory-confirmed influenza infection in a household member who developed fever (> 38.0 °C), cough, or sore throat during the observation period. They termed all other secondary cases as subclinical. A secondary outcome measure was the occurrence of ILI as defined by WHO as fever plus cough or sore throat.</p> <p>Safety: study reported that the majority of participants (107/172, 62%) did not report any problems with mask-wearing. This proportion was significantly higher in the group of adults (71/100, 71%) compared to the group of children (36/72, 50%) (P = 0.005). The main problem reported by participants (adults as well as children) was "heat/humidity" (18/34, 53% of children; 10/29, 35% of adults) (P = 0.1), followed by "pain" and "shortness of breath" when wearing a face mask.</p>
Notes	<p>Period study conducted: November 2009 to April 2011</p> <p>Funding: governmental</p> <p>Adherence: in general, daily adherence was good, reaching a plateau of over 50% in nearly all groups (M and MH groups; 2009/10 and 2010/11) from the third day on (by then the intervention had been implemented in all households). A gradual decline towards lower adherence began around the sixth day of the index patient's illness.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"prepared lists of random numbers with Microsoft Excel 2003 (Microsoft™ Co-operation, Seattle, USA) which were divided between the three intervention groups. Each participating physician received a list of random numbers with the interventions represented in a 1:1:1 ratio"
Allocation concealment (selection bias)	Low risk	"the participating physician received a list of random numbers with the interventions represented in a 1:1:1 ratio. Eligible index patients were randomly assigned a number, which was then communicated to the study center. The resulting intervention was only communicated to the households with the

Suess 2012 (Continued)

physicians. Intervention material was given to the study sites in closed boxes marked only with the randomisation number. Recruiting physicians were not aware of the allocation of the numbers to the interventions and the boxes for the three intervention arms looked identical. After randomisation, participants were given their box by the physician's assistants"

Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Outcomes are very objective and therefore unlikely to be influenced by lack of blinding. In addition, "physicians (as well as laboratory personnel) blinded from the randomisation results".
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"physicians (as well as laboratory personnel) blinded from the randomisation results". Outcomes are very objective and therefore unlikely to be influenced by lack of blinding.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up. Daily follow-up home visits over the short period of data collection (8 days)
Selective reporting (reporting bias)	Low risk	The follow-up period is very short (8 days) with very good coverage, and the criteria for defining the outcome are highly objective. All planned outcomes were reported.

Talaat 2011
Study characteristics

Methods	Cluster-RCT
Participants	Children (N = 44,451) in the first 3 primary grades from 60 governmental elementary schools in Cairo, Egypt were included and randomised to 30 schools in the intervention arm (N = 20,882 students) and 30 control schools (N = 23,569 students). No exclusion criteria provided.
Interventions	Students were required to wash their hands at least twice during the school days for about 45 seconds, followed by proper rinsing and drying on a clean towel. Campaign material was developed, and posters were placed near sinks in the classroom and playground to encourage hand-washing with soap and water upon arriving at school, before and after meals, using the bathroom, and after coughing and sneezing. See Table 1 for details.
Outcomes	Laboratory: point-of-care influenza A and B viruses using QuickVue (QuickVue; Quidel Corp., San Diego, CA, USA). School nurses collected nasal swabs from children who visited the school clinic with ILI, and only for students who had prior written approval of a parent. Effectiveness: rates of absenteeism caused by ILI and laboratory-confirmed influenza. ILI defined as fever > 38 °C and either cough or sore throat. Safety: none planned or reported by the investigators
Notes	The period study conducted: 16 February to 12 May 2008 Funding: unclear

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Talaat 2011 (Continued)

Random sequence generation (selection bias)	Low risk	"computer-generated random number table"
Allocation concealment (selection bias)	Unclear risk	No information given.
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	The participants and study personnel were not blinded, although lack of blinding is unlikely to have influenced the outcome. Laboratory-confirmed influenza was only conducted only for students who had prior written approval of a parent.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"Differential interest of study teams may have contributed to the low rate of testing in students who were absent because of ILI in the control schools compared to the intervention schools (12% vs 22%)"
Incomplete outcome data (attrition bias) All outcomes	High risk	No flow chart of clusters flow during the study period. No information on withdrawal. Differential interest of study teams may have contributed to the low rate of testing in students who were absent because of ILI in the control schools compared to the intervention schools (12% vs 22%) incomplete or loss of data. The total number ILI episodes could be an underestimate, as there is no proactive method to look for symptoms of ILI amongst the students; it depends on the student being absent or in class with symptoms that are picked up by the teachers at school.
Selective reporting (reporting bias)	Unclear risk	Insufficient information to judge

Talaat 2011
Study characteristics

Methods	Cluster-RCT
Participants	Children (N = 44,451) in the first 3 primary grades from 60 governmental elementary schools in Cairo, Egypt were included and randomised to 30 schools in the intervention arm (N = 20,882 students) and 30 control schools (N = 23,569 students). No exclusion criteria provided.
Interventions	Students were required to wash their hands at least twice during the school days for about 45 seconds, followed by proper rinsing and drying on a clean towel. Campaign material was developed, and posters were placed near sinks in the classroom and playground to encourage hand-washing with soap and water upon arriving at school, before and after meals, using the bathroom, and after coughing and sneezing. See Table 1 for details.
Outcomes	Laboratory: point-of-care influenza A and B viruses using QuickVue (QuickVue; Quidel Corp., San Diego, CA, USA). School nurses collected nasal swabs from children who visited the school clinic with ILI, and only for students who had prior written approval of a parent. Effectiveness: rates of absenteeism caused by ILI and laboratory-confirmed influenza. ILI defined as fever > 38 °C and either cough or sore throat. Safety: none planned or reported by the investigators
Notes	The period study conducted: 16 February to 12 May 2008 Funding: unclear

Talaat 2011 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"computer-generated random number table"
Allocation concealment (selection bias)	Unclear risk	No information given.
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	The participants and study personnel were not blinded, although lack of blinding is unlikely to have influenced the outcome. Laboratory-confirmed influenza was only conducted only for students who had prior written approval of a parent.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"Differential interest of study teams may have contributed to the low rate of testing in students who were absent because of ILI in the control schools compared to the intervention schools (12% vs 22%)"
Incomplete outcome data (attrition bias) All outcomes	High risk	No flow chart of clusters flow during the study period. No information on withdrawal. Differential interest of study teams may have contributed to the low rate of testing in students who were absent because of ILI in the control schools compared to the intervention schools (12% vs 22%) incomplete or loss of data. The total number ILI episodes could be an underestimate, as there is no proactive method to look for symptoms of ILI amongst the students; it depends on the student being absent or in class with symptoms that are picked up by the teachers at school.
Selective reporting (reporting bias)	Unclear risk	Insufficient information to judge

Temime 2018
Study characteristics

Methods	2-arm cluster-RCT
Participants	All residents and staff of 27 privately held chains of nursing homes owned by Korian. 26 nursing homes (13 per arm), with an average of 80 residents per nursing home, were included in the study.
Interventions	"The intervention was based on a bundle of HH-related measures aimed at NH staff, residents, visitors, and outside care providers. These measures included facilitated access to handrub solution using pocket-sized containers and new dispensers, a campaign to promote HH with posters and event organization, the formation of local work groups in each NH to work on HH guidelines, and staff education using e-learning on infection control and HH training performed by the same nurse for all NHs." See Table 1 for details.
Outcomes	Laboratory: none used Effectiveness: Primary outcomes: incidence rate of ARIs and AGE reported in the context of episodes of clustered cases, defined as at least 5 cases within 4 days amongst nursing home residents or staff. ARIs were defined as the combination of at least 1 respiratory symptom with 1 symptom of systemic infection. AGE was defined as the sudden onset of diarrhoea or vomiting in the absence of a non-infectious aetiology.

Temime 2018 (Continued)

Secondary endpoints were mortality rate, hospitalisation rate, and antibiotic prescription rate (measured in defined daily doses (DDDs) per 100 resident days).

Safety: no adverse event surveillance planned or reported by the investigators

Notes

The period study conducted: 1 April 2014 to 1 April 2015

Funding: private (Institute of Ageing Well Korian (Institut du bien vieillir Korian), which runs the nursing homes included in the study)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	"simple" randomisation is used
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	"we suspected that underreporting occurred. The data were verified qualitatively after the end of the intervention through individual phone interviews with each participating NH. Based on these interviews, ARI clustered cases episodes had actually occurred in 12 out of 13 control NHs; however, only 1 had been notified to health authorities. No unreported clustered cases episodes were identified in the intervention NHs"
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data were collected at NH level and reported to centralised by the NH group headquarters in Paris through computerised databases. There was underreporting of ARI and AGE in the control groups. The trial authors suspected that underreporting occurred. Primary outcome: high risk. Secondary outcomes: low risk
Incomplete outcome data (attrition bias) All outcomes	High risk	For the primary outcome, there was underreporting of ARI and AGE in the control groups; no study flow chart was provided; and no reporting on any exclusions. Surveillance is based on voluntary and standardised notifications to health authorities of any AGE or ARI clustered case episode.
Selective reporting (reporting bias)	Low risk	Reported outcomes match planned outcomes published in the protocol.

Temime 2018
Study characteristics

Methods	2-arm cluster-RCT
Participants	All residents and staff of 27 privately held chains of nursing homes owned by Korian. 26 nursing homes (13 per arm), with an average of 80 residents per nursing home, were included in the study.
Interventions	"The intervention was based on a bundle of HH-related measures aimed at NH staff, residents, visitors, and outside care providers. These measures included facilitated access to handrub solution using pocket-sized containers and new dispensers, a campaign to promote HH with posters and event organization, the formation of local work groups in each NH to work on HH guidelines, and staff education using e-learning on infection control and HH training performed by the same nurse for all NHs."

Temime 2018 (Continued)

See [Table 1](#) for details.

Outcomes	<p>Laboratory: none used</p> <p>Effectiveness:</p> <p>Primary outcomes: incidence rate of ARIs and AGE reported in the context of episodes of clustered cases, defined as at least 5 cases within 4 days amongst nursing home residents or staff. ARIs were defined as the combination of at least 1 respiratory symptom with 1 symptom of systemic infection. AGE was defined as the sudden onset of diarrhoea or vomiting in the absence of a non-infectious aetiology.</p> <p>Secondary endpoints were mortality rate, hospitalisation rate, and antibiotic prescription rate (measured in defined daily doses (DDDs) per 100 resident days).</p> <p>Safety: no adverse event surveillance planned or reported by the investigators</p>
Notes	<p>The period study conducted: 1 April 2014 to 1 April 2015</p> <p>Funding: private (Institute of Ageing Well Korian (Institut du bien vieillir Korian), which runs the nursing homes included in the study)</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	"simple" randomisation is used
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	"we suspected that underreporting occurred. The data were verified qualitatively after the end of the intervention through individual phone interviews with each participating NH. Based on these interviews, ARI clustered cases episodes had actually occurred in 12 out of 13 control NHs; however, only 1 had been notified to health authorities. No unreported clustered cases episodes were identified in the intervention NHs"
Blinding of outcome assessment (detection bias) All outcomes	High risk	Data were collected at NH level and reported to centralised by the NH group headquarters in Paris through computerised databases. There was underreporting of ARI and AGE in the control groups. The trial authors suspected that underreporting occurred. Primary outcome: high risk. Secondary outcomes: low risk
Incomplete outcome data (attrition bias) All outcomes	High risk	For the primary outcome, there was underreporting of ARI and AGE in the control groups; no study flow chart was provided; and no reporting on any exclusions. Surveillance is based on voluntary and standardised notifications to health authorities of any AGE or ARI clustered case episode.
Selective reporting (reporting bias)	Low risk	Reported outcomes match planned outcomes published in the protocol.

Turner 2004a

Study characteristics

Turner 2004a (Continued)

Methods	Double-blind RCT conducted by Hill Top Research, Inc., Winnipeg, Canada, to assess the efficacy of acids with virucidal activity for the inactivation of virus and prevention of experimental rhinovirus colds. Participants in good health, aged 18 to 60, were recruited from Winnipeg and surrounding communities for participation. Qualified participants were randomised to treatment with vehicle (62% ethanol, 1% ammonium lauryl sulphate, and 1% Klucel), vehicle containing 3.5% salicylic acid, or vehicle containing 1% salicylic acid and 3.5% pyroglutamic acid. The volunteers' hands were disinfected, and then test product was applied to both hands of participant. 15 minutes after application, the fingerprints of each hand were contaminated with rhinovirus type 39. The volunteers touched conjunctiva and the nasal mucosa only with the right hand. Viral contamination of the fingers was assessed in the left hands of the volunteers, and viral infection was assessed by culture of nasal lavage specimens and blood samples.
Participants	85 volunteers; 31 control group, 27 used vehicle with 3.5% salicylic acid, 27 used vehicle with 1% salicylic acid and 3.5% pyroglutamic acid
Interventions	Use of salicylic acid versus salicylic acid and pyroglutamic acid versus "placebo" substance
Outcomes	Laboratory: yes Effectiveness: rhinovirus type 39 infection Safety: N/A
Notes	Risk of bias: unclear (no description of randomisation process, concealment or allocation) Note: the authors concluded that organic acids commonly used in over-the-counter skin care and cosmetic products have substantial virucidal activity against rhinovirus. These preparations provided effective residual antiviral activity on the hands. The virucidal effect of these hand treatments resulted in a reduction in the incidence of rhinovirus infection in the treated volunteers ($P = 0.025$). The utility of this observation in the natural setting remains to be determined. The volunteers were not allowed to use their hands in the interval between the hand treatment and the virus challenge, so the effect of normal use of the hands on the virucidal activity of these organic acids is not known. Similarly, the virus challenge method used in these experiments may not simulate the natural setting in all aspects. The effect of nasal secretions that would be transferred with the virus in the natural setting on the activity of the acids or on the transmission of virus was not tested in the model. We are unsure as to the practical significance of this study and the generalisability of its results to the real world. Poorly reported study

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"randomised" Sequence generation not described.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	"double blind", but no description
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"double blind", but no description
Incomplete outcome data (attrition bias) All outcomes	Low risk	All accounted for (short study).

Turner 2004a (Continued)

Selective reporting (reporting bias)	High risk	Poorly reported
--------------------------------------	-----------	-----------------

Turner 2004b
Study characteristics

Methods	<p>Double-blind RCT conducted by Hill Top Research, Inc., Winnipeg, Canada, to assess the residual virucidal activity of a skin cleanser wipe and its effectiveness in preventing experimental rhinovirus colds. Participants in good health, aged 18 to 60 years, were recruited from Winnipeg and surrounding communities for participation.</p> <p>The residual activity of a skin cleanser wipe containing 4% pyroglutamic acid formulated with 0.1% benzalkonium chloride was tested. The negative control treatment was 62% ethanol. Benzalkonium chloride had been previously tested and was found to have no virucidal activity. Volunteers were randomly assigned to use the control preparation or the active preparation. The study material was applied to hands with a towelette. 15 minutes later, when the fingers were completely dry, the fingertips of each hand of the control participants and the volunteers in the active treatment group were contaminated with rhinovirus type 39. An additional volunteer in the active group was challenged with virus 1 hour after application, and the final group of volunteers was challenged 3 hours after application. Viral infection was assessed by culture of nasal lavage specimens and blood samples.</p>
Participants	122 volunteers; 30 in control group, 92 in active group (30 tested after 15 minutes, 30 after 1 hour, 32 after 2 hours)
Interventions	Use of a skin cleanser wipe containing 4% pyroglutamic acid formulated with 0.1% benzalkonium chloride versus skin cleanser wipe containing ethanol
Outcomes	Laboratory: yes Effectiveness: rhinovirus type 39 infection Safety: N/A
Notes	Risk of bias: unclear (no description of randomisation process, concealment or allocation)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"randomised" Sequence generation not described.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	"double blind", but no description given
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"double blind", but no description given
Incomplete outcome data (attrition bias) All outcomes	Low risk	All accounted for (short study).

Turner 2004b (Continued)

Selective reporting (reporting bias)	High risk	Poorly reported
--------------------------------------	-----------	-----------------

Turner 2012
Study characteristics

Methods	Randomised controlled clinical trial
Participants	<p>A total of 212 participants were enrolled (116 in the treatment group, 96 in the control group).</p> <p>Healthy adult volunteers aged > 18 years from the University of Virginia community. Written informed consent was obtained, and volunteers were compensated for participation.</p> <p>Exclusion: individuals with skin conditions that would interfere with safety evaluations or medical conditions that could impact the person's well-being or affect study results, and those whose occupations required frequent hand-washing.</p>
Interventions	Antiviral hand treatment containing 2% citric acid, 2% malic acid, and 62% ethanol (n = 116) or to a no-treatment control group (n = 96). The hand treatment was applied every 3 hours and after hand-washing whilst the participants were awake. See Table 1 for details.
Outcomes	<p>Laboratory: PCR using AmpliTaq Gold DNA Polymerase from Applied Biosystems</p> <p>Effectiveness: reduction of rhinovirus-induced common colds; comparison of the number of RV-associated illnesses per 100 participants in the control group with that in the treatment group over 9 weeks. Definitions: a common cold illness was defined as the presence of any of the symptoms of nasal obstruction, rhinorrhoea, sore throat, or cough on at least 3 consecutive days. Illnesses separated by at least 3 symptom-free days were considered to be separate illnesses. Rhinovirus infection was defined as the detection of RV in nasal lavage. All volunteers were seen weekly for nasal lavage, and specimens were assayed by PCR for the presence of RV. PCR-positive specimens separated by at least 8 days and at least 1 negative PCR specimen were considered to be separate infections. RV-associated illnesses were based on detection of RV either at the time of the illness or at the first weekly visit after the illness.</p> <p>Safety: hand irritation occurred in 11 of the 116 volunteers (9%) in the treatment group, which met protocol criteria for removal from the study. An additional 8 participants who did not meet these protocol criteria voluntarily withdrew due to hand irritation. There was no hand irritation in the control group. No other adverse effects of the study treatment were noted.</p>
Notes	<p>The period study conducted: August 2009 to November 2009</p> <p>Funding: The Dial Corporation - a Henkel Company, Scottsdale, Arizona, USA</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"A randomization code generated using commercially available software was provided by the sponsor"
Allocation concealment (selection bias)	Low risk	"staff at the study site assigned sequential subject numbers as they enrolled volunteers into the study, and treatment assignment was determined by the subject number."
Blinding of participants and personnel (performance bias)	Low risk	The outcomes are unlikely to be influenced by lack of blinding.

Turner 2012 (Continued)

All outcomes

Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Personnel who conducted the laboratory assays were blinded to study groups and to whether the specimen was from a routine or illness related visit"
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition (and reasons for it) was reported. Study outcomes reported as ITT and PP.
Selective reporting (reporting bias)	Low risk	All planned outcomes in study protocol were reported on.

Turner 2004a
Study characteristics

Methods	Double-blind RCT conducted by Hill Top Research, Inc., Winnipeg, Canada, to assess the efficacy of acids with virucidal activity for the inactivation of virus and prevention of experimental rhinovirus colds. Participants in good health, aged 18 to 60, were recruited from Winnipeg and surrounding communities for participation. Qualified participants were randomised to treatment with vehicle (62% ethanol, 1% ammonium lauryl sulphate, and 1% Klucel), vehicle containing 3.5% salicylic acid, or vehicle containing 1% salicylic acid and 3.5% pyroglutamic acid. The volunteers' hands were disinfected, and then test product was applied to both hands of participant. 15 minutes after application, the fingerprints of each hand were contaminated with rhinovirus type 39. The volunteers touched conjunctiva and the nasal mucosa only with the right hand. Viral contamination of the fingers was assessed in the left hands of the volunteers, and viral infection was assessed by culture of nasal lavage specimens and blood samples.	
Participants	85 volunteers; 31 control group, 27 used vehicle with 3.5% salicylic acid, 27 used vehicle with 1% salicylic acid and 3.5% pyroglutamic acid	
Interventions	Use of salicylic acid versus salicylic acid and pyroglutamic acid versus "placebo" substance	
Outcomes	Laboratory: yes Effectiveness: rhinovirus type 39 infection Safety: N/A	
Notes	Risk of bias: unclear (no description of randomisation process, concealment or allocation) Note: the authors concluded that organic acids commonly used in over-the-counter skin care and cosmetic products have substantial virucidal activity against rhinovirus. These preparations provided effective residual antiviral activity on the hands. The virucidal effect of these hand treatments resulted in a reduction in the incidence of rhinovirus infection in the treated volunteers (P = 0.025). The utility of this observation in the natural setting remains to be determined. The volunteers were not allowed to use their hands in the interval between the hand treatment and the virus challenge, so the effect of normal use of the hands on the virucidal activity of these organic acids is not known. Similarly, the virus challenge method used in these experiments may not simulate the natural setting in all aspects. The effect of nasal secretions that would be transferred with the virus in the natural setting on the activity of the acids or on the transmission of virus was not tested in the model. We are unsure as to the practical significance of this study and the generalisability of its results to the real world. Poorly reported study	

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Turner 2004a (Continued)

Random sequence generation (selection bias)	Unclear risk	"randomised" Sequence generation not described.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	"double blind", but no description
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"double blind", but no description
Incomplete outcome data (attrition bias) All outcomes	Low risk	All accounted for (short study).
Selective reporting (reporting bias)	High risk	Poorly reported

Turner 2004b
Study characteristics

Methods	<p>Double-blind RCT conducted by Hill Top Research, Inc., Winnipeg, Canada, to assess the residual virucidal activity of a skin cleanser wipe and its effectiveness in preventing experimental rhinovirus colds. Participants in good health, aged 18 to 60 years, were recruited from Winnipeg and surrounding communities for participation.</p> <p>The residual activity of a skin cleanser wipe containing 4% pyroglutamic acid formulated with 0.1% benzalkonium chloride was tested. The negative control treatment was 62% ethanol. Benzalkonium chloride had been previously tested and was found to have no virucidal activity. Volunteers were randomly assigned to use the control preparation or the active preparation. The study material was applied to hands with a towelette. 15 minutes later, when the fingers were completely dry, the fingertips of each hand of the control participants and the volunteers in the active treatment group were contaminated with rhinovirus type 39. An additional volunteer in the active group was challenged with virus 1 hour after application, and the final group of volunteers was challenged 3 hours after application. Viral infection was assessed by culture of nasal lavage specimens and blood samples.</p>
Participants	122 volunteers; 30 in control group, 92 in active group (30 tested after 15 minutes, 30 after 1 hour, 32 after 2 hours)
Interventions	Use of a skin cleanser wipe containing 4% pyroglutamic acid formulated with 0.1% benzalkonium chloride versus skin cleanser wipe containing ethanol
Outcomes	Laboratory: yes Effectiveness: rhinovirus type 39 infection Safety: N/A
Notes	Risk of bias: unclear (no description of randomisation process, concealment or allocation)

Risk of bias

Bias	Authors' judgement	Support for judgement
------	--------------------	-----------------------

Turner 2004b (Continued)

Random sequence generation (selection bias)	Unclear risk	"randomised" Sequence generation not described.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	"double blind", but no description given
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"double blind", but no description given
Incomplete outcome data (attrition bias) All outcomes	Low risk	All accounted for (short study).
Selective reporting (reporting bias)	High risk	Poorly reported

Turner 2012
Study characteristics

Methods	Randomised controlled clinical trial
Participants	<p>A total of 212 participants were enrolled (116 in the treatment group, 96 in the control group).</p> <p>Healthy adult volunteers aged > 18 years from the University of Virginia community. Written informed consent was obtained, and volunteers were compensated for participation.</p> <p>Exclusion: individuals with skin conditions that would interfere with safety evaluations or medical conditions that could impact the person's well-being or affect study results, and those whose occupations required frequent hand-washing.</p>
Interventions	<p>Antiviral hand treatment containing 2% citric acid, 2% malic acid, and 62% ethanol (n = 116) or to a no-treatment control group (n = 96). The hand treatment was applied every 3 hours and after hand-washing whilst the participants were awake. See Table 1 for details.</p>
Outcomes	<p>Laboratory: PCR using AmpliTaq Gold DNA Polymerase from Applied Biosystems</p> <p>Effectiveness: reduction of rhinovirus-induced common colds; comparison of the number of RV-associated illnesses per 100 participants in the control group with that in the treatment group over 9 weeks. Definitions: a common cold illness was defined as the presence of any of the symptoms of nasal obstruction, rhinorrhoea, sore throat, or cough on at least 3 consecutive days. Illnesses separated by at least 3 symptom-free days were considered to be separate illnesses. Rhinovirus infection was defined as the detection of RV in nasal lavage. All volunteers were seen weekly for nasal lavage, and specimens were assayed by PCR for the presence of RV. PCR-positive specimens separated by at least 8 days and at least 1 negative PCR specimen were considered to be separate infections. RV-associated illnesses were based on detection of RV either at the time of the illness or at the first weekly visit after the illness.</p> <p>Safety: hand irritation occurred in 11 of the 116 volunteers (9%) in the treatment group, which met protocol criteria for removal from the study. An additional 8 participants who did not meet these protocol</p>

Turner 2012 (Continued)

criteria voluntarily withdrew due to hand irritation. There was no hand irritation in the control group. No other adverse effects of the study treatment were noted.

Notes

The period study conducted: August 2009 to November 2009

Funding: The Dial Corporation - a Henkel Company, Scottsdale, Arizona, USA

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"A randomization code generated using commercially available software was provided by the sponsor"
Allocation concealment (selection bias)	Low risk	"staff at the study site assigned sequential subject numbers as they enrolled volunteers into the study, and treatment assignment was determined by the subject number."
Blinding of participants and personnel (performance bias) All outcomes	Low risk	The outcomes are unlikely to be influenced by lack of blinding.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Personnel who conducted the laboratory assays were blinded to study groups and to whether the specimen was from a routine or illness related visit"
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition (and reasons for it) was reported. Study outcomes reported as ITT and PP.
Selective reporting (reporting bias)	Low risk	All planned outcomes in study protocol were reported on.

White 2001
Study characteristics

Methods	Double-blind, placebo-controlled, cluster-RCT that took place in 3 schools in California during March to April 1999. The study assessed the incremental value of using an alcohol hand rub together with water-and-soap hand-washing. Both arms were administered an educational programme beginning 2 weeks prior to start of the trial. Randomisation was by classroom, and the placebo hand rub was indistinguishable from the active ingredient. Details of randomisation are not given.
Participants	Of the 72 classes originally recruited, lack of compliance (use of supplementary product at least 3 times a day) reduced the classes to 32 (16 in both arms) and a total of 769 participants aged 5 to 12 (381 students who received the sanitiser, and 388 who received the placebo).
Interventions	Pump-activated antiseptic hand rub with benzalkonium chloride (SAB) (Woodward Laboratories) or inert placebo that "virtually" looked the same in batches of 4 colour-coded bottles. School staff, parents, and participants were blinded.
Outcomes	Laboratory: testing of virucidal and bactericidal activity of the active compound Effectiveness: ARI (cough, sneezing, sinus trouble, bronchitis, fever, red eye, headache, mononucleosis, acute exacerbations of asthma) Gastrointestinal and other illnesses (data not extracted) Follow-up and observation was carried out by classroom staff, and illnesses were described by parents.

White 2001 (Continued)

Safety: 7 students dropped out because of mild sensitivity to the rub

Notes

Risk of bias: high (no description of randomisation; partial reporting of outcomes, numerators and denominators)

Note: the authors conclude that addition of the rub led to a 30% to 38% decrease of illness and absenteeism (RR for illness absence incidence 0.69, RR for absence duration 0.71). Very high attrition, unclear randomisation procedure, educational programme and use of placebo hand rub make generalisability of the results debatable. No confidence intervals reported.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"randomised trial", but sequence generation not described
Allocation concealment (selection bias)	Unclear risk	Not described
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"To distinguish content, both the active and placebo formulations were distributed in four color-coded groups of 1oz spritz bottles. The content were and distribution patters were only know to the researchers and were indecipherable by the school staff or students."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"Teachers were responsible for recording attendance for each day during the study"
Incomplete outcome data (attrition bias) All outcomes	High risk	Partial reporting of outcomes, numerators and denominators
Selective reporting (reporting bias)	High risk	Poor reporting

Yeung 2011
Study characteristics

Methods

Clustered-RCT of a hand hygiene intervention involving pocket-sized containers of alcohol-based hand rub for the control of infections in long-term care facilities. Staff hand hygiene adherence was directly observed, and residents' infections necessitating hospitalisation were recorded. After a 3-month pre-intervention period, long-term care facilities (LTCFs) were randomised to receive pocket-sized containers of alcohol-based gel, reminder materials, and education for all HCWs (treatment group) or to receive basic life support education and workshops for all HCWs (control group). A 2-week intervention period (1 to 15 April 2007) was followed by 7 months of postintervention observations.

Participants

6 out of 7 community-based, private or semiprivate, residential LTCFs in Hong Kong agreed to participate and were randomised to:

1. hand hygiene group (3 LTCFs, 73 nursing staff and 244 residents analysed); or
2. control group (3 LTCFs, 115 nursing staff and 379 residents analysed).

All were nursing homes serving an elderly population. All LTCFs were situated in different regions of Hong Kong, including urban and rural areas. The targets of the intervention were all full- and part-time HCWs at these LTCFs.

Yeung 2011 (Continued)

The LTCFs employed 3 types of HCWs: nurses, nursing assistants, and physiotherapists.

Interventions	Pocket-sized containers of alcohol-based gel, reminder materials, and education (intervention group) or basic life-support education and workshop (control group). See Table 1 for details.
Outcomes	<p>Rates of infection (requiring hospitalisation)</p> <p>Outbreaks</p> <p>Death due to infection</p> <p>Diagnoses of infection coded into 6 categories, all of which were common endemic infections in LTCFs:</p> <ol style="list-style-type: none"> 1. pneumonia, 2. urinary tract infection, 3. septicaemia, 4. skin or soft-tissue infection (including cellulitis or pressure sores), 5. gastroenteritis, and 6. fever. <p>Infections recorded in death certificates were also included, regardless of whether the resident had been hospitalised. The causes of death were categorised as due to infection, not due to infection, or unknown. If the primary or the secondary diagnosis on the death certificate belonged to 1 of the 6 endemic infection categories, the death was coded as due to infection.</p> <p>No safety outcomes reported.</p>
Notes	University and industry funded

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	No details provided.
Allocation concealment (selection bias)	Unclear risk	No details provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Unblinded study
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded study
Incomplete outcome data (attrition bias) All outcomes	Low risk	No loss to follow-up
Selective reporting (reporting bias)	Unclear risk	No protocol available

Zomer 2015
Study characteristics

Methods	Cluster-RCT
Participants	<p>71 daycare centres (36 intervention DCCs, and 35 control) in Rotterdam-Rijnmond, Gouda and Leiden in the Netherlands</p> <p>Study enrolled 545 children (intervention = 278, control = 267).</p> <p>Inclusion/exclusion criteria: children who attended the DCC at least 2 days a week; were aged between 6 months and 3.5 years at start of the trial; intended to attend the DCC throughout the study period; and if their parents consented, were Dutch-speaking, and had access to email or regular post. Children were excluded if they had a chronic illness or medication that predisposed them to infection, a sibling taking part in the trial (i.e. 1 child per family could be included), or if they started attending CCC after the beginning of the trial).</p>
Interventions	<p>4 components:</p> <ol style="list-style-type: none"> 1. HH products, paper towel dispensers, soap, alcohol-based hand sanitiser, and hand cream were provided for 6 months. 2. Training and a booklet outlining the training. 3. 2 team training sessions aimed at specific HH improvement activities. 4. Posters and stickers for caregivers and children as reminders. <p>See Table 1 for details.</p>
Outcomes	<p>Laboratory: none</p> <p>Effectiveness: incidence of respiratory infections in children monitored by parents. The common cold was defined as a blocked or runny nose with at least 1 of the following symptoms: coughing, sneezing, fever, sore throat, or earache.</p> <p>Safety: none planned or reported by the investigators</p>
Notes	<p>The period study conducted: September 2011 to April 2012</p> <p>Funding: mixed. The Netherlands Organisation for Health Research and Development (ZonMw). Dispensers and refills were sponsored by SCA Hygiene Products, Sweden.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stratified randomization is performed by assigning each DCC to one of six strata based on size (i.e. small < 46 children per day versus large ≥ 46 children per day) and geographic location (i.e. highly urban versus urban versus slightly/non-urban). DCCs are assigned to either intervention or control group by means of computer generation with a 1:1 ratio in each of the strata"
Allocation concealment (selection bias)	Unclear risk	No information provided.
Blinding of participants and personnel (performance bias) All outcomes	High risk	Outcome is subjective.
Blinding of outcome assessment (detection bias)	High risk	Symptoms were reported by parents, no validation.

Zomer 2015 (Continued)

All outcomes

Incomplete outcome data (attrition bias) All outcomes	Low risk	Very few children were excluded or lost to follow-up (reasons for exclusions provided).
Selective reporting (reporting bias)	Low risk	All planned outcomes are reported. However, between published protocol and the paper, secondary outcomes became the primary outcome in the published paper!

AEs: adverse events
 AFH: Armed Forces Hospital
 AGE: acute gastroenteritis
 ALRI: acute lower respiratory infection
 ARI: acute respiratory infection
 ASR: adverse skin reactions
 A&E: accident and emergency
 BIPAP: bilevel positive airway pressure
 CCC: childcare centre
 CDC: Centers for Disease Control and Prevention
 CG: control group
 CHG: chlorhexidine gluconate
 CI: confidence interval
 CMF: citric acid: malic acid: sodium lauryl sulphate (a virucidal mixture added to tissue paper)
 CoV: coronavirus
 cluster-RCT: cluster-randomised controlled trial
 CRI: clinical respiratory illness
 CXR: chest X-ray
 DCC: daycare centre
 EG: experimental group
 FRI: febrile respiratory illness
 GI: gastrointestinal
 GTI: gastrointestinal infection
 GP: general practitioner
 HCW: healthcare worker
 HFH: Hanoi French Hospital
 HH: hand hygiene
 HR: high risk
 HSG: hand sanitiser group
 ICD-9: International Classification of Disease, 9th Revision, Clinical Modification
 ICU: intensive care unit
 ILI: influenza-like illness
 IQR: interquartile range
 IRR: incident rate ratio
 ITT: intention-to-treat
 LRTI: lower respiratory tract infection
 LTCF: long-term care facility
 MCU: medical convalescent unit
 MDCK: Madin Darby canine kidney cell line
 M group: face mask group
 MH group: face mask and hand hygiene group
 MS: monkey-derived cell line
 N/A: not applicable
 NAT: nucleic acid testing
 NH: nursing home
 NICU: neonatal intensive care unit
 NOS: Newcastle-Ottawa Scales
 NTS: nasal and throat swab
 OR: odds ratio
 PCR: polymerase chain reaction

PCU: physical conditioning unit
 POCT: point-of-care testing
 PP: per protocol
 PPE: personal protective equipment
 QNAF: Qatar National Research Fund
 RCT: randomised controlled trial
 RDS: respiratory distress syndrome
 RI: respiratory infection
 RIDT: rapid influenza diagnostic test
 RNA: ribonucleic acid
 RR: risk ratio
 rRT-PCR: real-time reverse transcription-polymerase chain reaction
 RTI: respiratory tract infection
 RT-PCR: reverse-transcriptase polymerase chain reaction
 RSV: respiratory syncytial virus
 RV: rhinovirus
 SAB: surfactant, allantoin, and benzalkonium chloride
 SAR: secondary attack rate
 SARS: severe acute respiratory syndrome
 SCBU: special care baby unit
 SD: standard deviation
 SHEWA-B: Sanitation, Hygiene Education and Water Supply in Bangladesh
 SOB: shortness of breath
 SOPs: standard operating procedures
 S/S: signs/symptoms
 SSTI: skin and soft-tissue infection
 STH: soil-transmitted helminth
 SWG: soap and water group
 TIDieR: Template for Intervention Description and Replication
 UHR-I: ultra high-risk infection
 UHR-S: ultra high-risk SARS
 URI: upper respiratory infection
 URTI: upper respiratory tract infection
 WBC: white blood cell
 WHO: World Health Organization

Characteristics of excluded studies *[ordered by study ID]*

Study	Reason for exclusion
Abou El Hassan 2004	Topic completely extraneous
Amirav 2005	Randomised controlled trial of aerosol treatment
Anderson 2004	Mathematical model with interesting discussion of interaction between public health measures
Anonymous 2002	News item
Anonymous 2004	News item
Anonymous 2005a	News item
Anonymous 2005b	News item
Anonymous 2005c	News item
Apisarnthanarak 2009	Intervention bundle not broken down.
Apisarnthanarak 2010	Participants took antivirals.

Study	Reason for exclusion
Aragon 2005	Descriptive paper (non-comparative). Has no viral outcomes
Azor-Martinez 2014	Results reported as respiratory and gastrointestinal infections. No extractable respiratory data
Barros 1999	Correlational study between incidence of URTI and factors such as overcrowding
Bauer 2009	Historical comparison with RSV gammaglobulin amongst interventions
Bell 2004	Has unpublished entry exit screening data and extensive references but no comparative data
Bellissimo-Rodrigues 2009	Intervention is chlorhexidine.
Ben-Abraham 2002	Exclude - bacterial illness only
Black 1981	Diarrhoea only outcome
Borkow 2010	No human beings involved.
Bouadma 2010	Hospital-based ventilator routine
Bowen 2007	Outcomes of composite infections. Respiratory infections are not reported separately.
Breugelmans 2004	Description of risk factors in aircraft
Cai 2009	Compliance study
Cantagalli 2010	Outcome outside inclusion criteria
Carbonell-Estrany 2008	Immunoglobulin intervention and descriptive review
Carter 2002	News item
Castillo-Chavez 2003	Editorial
Cava 2005a	Survey of quarantinees' views
Cava 2005b	Personal experiences of quarantine
CDC 2003a	Case reports
CDC 2003b	No data presented.
Chai 2005	Letter - about MRSA
Chami 2012	Outcomes of composite infections. Respiratory infections are not reported separately.
Chaovavanich 2004	Case report
Chau 2003	No original retrievable data. Mathematical model fitting expected to observed cases with quarantine in the SARS of Hong Kong
Chau 2008	Audit of infection control procedures and compliance with guidelines
Chen 2007	An assessment of the impact of different hand-washing teaching methods. No clinical outcomes

Study	Reason for exclusion
Cheng 2010	Confounded by antiviral use for postexposure prophylaxis
Chia 2005	Knowledge survey
Clynes 2010	Letters
Cowling 2007	Epidemiology, non-comparative, non-interventions study
Daniels 2010	Commentary
Daugherty 2008	No free data presented.
Davies 1994	Antibody titres as outcomes with so many biases that interpretation of study is problematic
Day 1993	No acute respiratory infection outcome data
Day 2006	Mathematical model; no new data
Dell'Omodarme 2005	Probabilistic and Bayesian mathematical model of screening at entry
Denbak 2018	Outcomes of composite infections. Respiratory infections are not reported separately.
Desenclos 2004	Description of transmission
DiGiovanni 2004	Qualitative study of compliance factors in quarantine
Doebbeling 1992	RCT respiratory data not present. Only 3 viruses isolated in total with no viral typing available.
Dwosh 2003	Case series
Edmonds 2010	Lab study
Fendler 2002	Cohort study badly biased with differential health profiles and healthcare workers dependency in intervention and control semi-cohorts. No attempt to adjust for confounders was made. No denominators available.
Flint 2003	Description of spread in aircraft and non-comparative data
Fung 2004	Non-comparative
Garcia 2010	Commentary
Gaydos 2001	Editorial linked to Ryan 2001. (Ryan 2001 was an included trial in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Gensini 2004	Interesting historical review
Girou 2002	Non-clinical outcomes
Glass 2006	Mathematical model - no original data presented
Goel 2007	Non-comparative study
Gomersall 2006	Non-comparative study

Study	Reason for exclusion
Gore 2001	Summary of Dyer 2000. (Dyer 2000 was a prospective, cluster open-label cross-over cohort study included in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Gostin 2003	Not an analytical study
Gralton 2010	Review
Guinan 2002	It would appear that 9 classes took part and "acted as their own controls", but it is not clear if there was cross-over of classes or not. In addition, the outcome is combined gastrointestinal/respiratory. The clue lies in the presence of a nested economic analysis which shows considerable savings in time for staff and pupils if the soap is used: in other words this is a (covert) publicity study.
Gupta 2005	Economic model - no new data
Gwaltney 1982	No breakdown of cases given by arm.
Han 2003	Non-comparative
Hayden 1985	This is an RCT with laboratory-induced colds, small numbers, and uncertain numerators, but almost certainly because of the unique laboratory conditions (placebo tissues not being a placebo at all) of impossible generalisation. It was a pilot to the far bigger trial by Farr 1988a ; Farr 1988b .
Hendley 1988	Inappropriate intervention
Hens 2009	Model
Heymann 2009	Already included in review as Heymann 2004. (Heymann 2004 was a controlled before and after study included in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Hilburn 2003	No ARI/viral outcomes (e.g. URTIs)
Hilmarsson 2007	Animal study
Hirsch 2006	Study tested pharmacological interventions.
Ho 2003	Descriptive review
Hsieh 2007	Mathematical model
Hugonnet 2007	Letter without any data
Jiang 2003	2 papers that are probably different versions of the same paper: Jiang SP, Huang LW, Wang JF, Wu W, Yin SM, Chen WX, et al. A study of the architectural factors and the infection rates of healthcare workers in isolation units for severe acute respiratory syndrome. <i>Chung-Hua Chieh Ho Ho Hu Hsi Tsa Chih [Chinese Journal of Tuberculosis & Respiratory Diseases]</i> . 26(10):594-7, 2003 Oct
Johnson 2009	Outcomes are non-clinical.
Jones 2005	Historical account
Kaydos-Daniels 2004	Not an analytical study
Kelso 2009	Model

Study	Reason for exclusion
Khaw 2008	Assessing the efficacy of O ₂ delivery
Kilabuko 2007	Aetiological study
Kosugi 2004	Non-comparative study
Lam 2004	Outcomes were generic (infection rates). No laboratory data available for viral diagnosis.
Lange 2004	No data presented.
Larson 2004a	Inappropriate outcomes
Larson 2004b	Inappropriate outcomes
Larson 2005	Cluster-RCT comparing the effects of 2 hand hygiene regimens on infection rates and skin condition and microbial counts of nurses' hands in neonatal intensive care units. Outcomes were generic (e.g. pneumonia and microbial counts of participants' skin). No laboratory data available for viral diagnosis.
Lau 2004	Attitude survey
Lau 2005	Herbal remedy effectiveness assessment
Lee 2005	Descriptive study of risk and protective factors of transmission in households. No assignment took place.
Lee 2010	Cohort study; unclear numbers were vaccinated against influenza
Lennell 2008	Measured absenteeism due to non-specific infection
Lipsitch 2003	Mathematical model fit to evidence
Luckingham 1984	Historical report on Tucson experience during Spanish flu pandemic
Ma 2004	Case-control study of risk factors for SARS
MacIntyre 2010	Commentary on Cowling 2009
Malone 2009	Model
Marin 1991	Viral resistance study
McSweeney 2007	Historical description
Mielke 2009	Review
Mikolajczyk 2008	No intervention
Monsma 1992	Non-comparative study
Nandrup-Bus 2009	The trial had only 2 clusters.
Nishiura 2009	Model

Study	Reason for exclusion
O'Callaghan 1993	Letter linked to Isaacs 1991. (Isaacs 1991 was a retrospective and prospective cohort study included in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Olsen 2003	Description of transmission
Ooi 2005	Descriptive study, but with interesting organisational chart
Orellano 2010	Confounded by antiviral use
Panchabhai 2009	Pharma intervention
Pang 2004	Descriptive study of Beijing outbreak. Some duplicate data in common with Pang 2003. (Pang 2003 was an ecological study included in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Patel 2012	Although within each district the participating schools and households were randomly selected, the allocation of districts to the intervention and comparison arms was not randomly assigned.
Pittet 2000	Analysis of relationship between hand-washing compliance campaign and nosocomial bacterial infections (e.g. MRSA)
Prasad 2004	Letter about retrospective cohort - behavioural
Rabenau 2005	In vitro test of several disinfectants
Reynolds 2008	Describes the psychological effects of quarantine
Richardson 2010	Non-clinical study
Riley 2003	Mathematical model fit to evidence
Rodriguez 2009	A "reasonable attempt at minimizing bias" (see inclusion criteria) does not include absenteeism
Rosen 2006	Non-specific outcome. Measured absenteeism
Rosenthal 2005	Outcomes were generic (e.g. pneumonia, URIs). No laboratory data available for viral diagnosis.
Safiulin 1972	Non-comparative set of studies with no clinical outcomes
Sandrock 2008	Review
Sattar 2000	Experiment assessing virucidal activity of fingertip surface - no clinical outcome data
Schull 2007	Describes the impact of SARS in a Toronto study
Seal 2010	Lab study
Seale 2009	Study looking at whether using respirators in A&E department is feasible
Sizun 1996	This is a review; no original data presented.
Slayton 2016	Compares hand-washing plus (antibacterial) towel versus hand-washing without towel
Stebbins 2009	Attitude survey

Study	Reason for exclusion
Stedman-Smith 2015	Composite outcome. No data on separate respiratory illnesses reported.
Stoner 2007	No study data available.
Stukel 2008	Impact of the SARS disruption on care/mortality for other pathologies (e.g. acute myocardial infarction). There are no interventions, and outcomes are unrelated to acute respiratory infections.
Svoboda 2004	Descriptive study with before-and-after data but shifting denominators
Tracht 2010	Model
Ueno 1990	Experimental study. No clinical intervention
Uhari 1999	No respiratory illness data to be extracted
van der Sande 2008	Laboratory study without any clinical outcomes
Vessey 2007	Composite outcome. No data on separate respiratory illnesses reported.
Viscusi 2009a	Lab study
Viscusi 2009b	Lab study
Wang 2003	Descriptive study
Wang 2005	Case-control study of susceptibility factors
Weber 2004	Editorial linked to Larson 2004a
Wen 2010	Lab study
White 2005	Redundant publication of White 2003. (White 2003 was a prospective, open, cohort study included in the previous version of this review (2011). Non-RCTs were removed in this 2020 update).
Wilczynski 1997	Clinical trial of the effects of breastfeeding
Wilder-Smith 2003	Description of risk factors in aircraft
Wilder-Smith 2005	Descriptive review
Wong 2005	Attitude survey
Yen 2010	Model
Yu 2004	Description of transmission
Zamora 2006	Head-to-head comparison of 2 sets of PPEs with no controls and no clinical outcomes
Zhai 2007	Non-comparative study
Zhao 2003	CCT of SARS treatment

A&E: accident and emergency
 ARI: acute respiratory infection
 CCT: controlled clinical trial

MRSA: methicillin-resistant *Staphylococcus aureus*

RCT: randomised controlled trial

RSV: respiratory syncytial virus

PPE: personal protective equipment

SARS: severe acute respiratory syndrome

URTI: upper respiratory tract infection

Characteristics of ongoing studies [ordered by study ID]

NCT03454009

Study name	Appropriate time-interval application of alcohol hand gel on reducing influenza-like illness among preschool children: a randomized, controlled trial
Methods	<p>This is a comprehensive randomised cluster hand-hygiene improvement intervention to reduce self-reported ARI/ILI and GI illness, absenteeism, presenteeism and related behavioural and attitudinal change over a 90-day trial. The intervention group will receive hand hygiene supplies and a variety of educational materials, including environmental posters in common areas. The control group will perform their usual hygiene activities and will not receive an intervention.</p> <p>Identical weekly surveys will be administered to the intervention and control groups to measure self-reported illness, absenteeism, presenteeism, along with behaviour and attitudes measured at specified intervals during the study. The intervention and control groups were randomised by work floors before the onset of the enrolment period. It is hypothesised that employees in the intervention group will experience reduced self-reported illness, absenteeism, and presenteeism along with improved protective hygiene behaviours and related attitudes, relative to those in the control group over the 90-day trial.</p>
Participants	<p>Inclusion criteria:</p> <ol style="list-style-type: none"> 1. At least 18 years of age or older 2. No known allergies to alcohol or surface disinfecting wipes 3. Works at least 30% of office hours at the study host site 4. Consent to receiving emails from Kent State University <p>Exclusion criteria:</p> <ol style="list-style-type: none"> 1. Under 18 years of age 2. Known allergies to alcohol or surface disinfecting wipes 3. Works less than 30% of office hours at the study host site 4. Does not consent to receiving emails from Kent State University
Interventions	The intervention group will receive hand hygiene supplies and a variety of educational materials, including environmental posters in common areas. The control group will perform their usual hygiene activities and will not receive an intervention.
Outcomes	Self-reported ARI/ILI and GI illness, absenteeism, presenteeism and related behavioural and attitudinal change over a 90-day trial
Starting date	5 February 2018
Contact information	Maggie Stedman-Smith, PhD, Kent State University College of Public Health
Notes	Recruitment completed. Last update in ClinicalTrials.gov was 1 May 2019. NCT03454009

NCT04267952

Study name	Hand hygiene intervention program on primary school students' health outcomes and absenteeism in school
Methods	<p>Study Type: interventional (clinical trial)</p> <p>Estimated enrolment: 200 participants</p> <p>Allocation: randomised</p> <p>Intervention model: parallel assignment</p> <p>Masking: single (participant)</p> <p>Masking description: participation will not know whether they are in the experimental or control group</p>
Participants	<p>Inclusion criteria: primary school student (especially third- and fourth-class student)</p> <p>Exclusion criteria: people with chronic disease</p>
Interventions	<p>Experimental: first group</p> <p>Hand hygiene intervention programme prepared by using planned behaviour theory will be applied to the students in this group.</p> <p>Active comparator: second group</p> <p>Students in this group will be given classic hand hygiene training.</p>
Outcomes	<p>Primary outcome measure: children with symptoms of infection will be referred to the family physician to have a rapid antigen test and to report the result to the researcher.</p> <p>10 identified upper respiratory tract symptoms (fever, sore throat, runny nose, etc.) will be recorded weekly by family of children. The researcher will receive symptom information from the family via weekly SMS.</p> <p>The number of days the child does not attend school due to illness and the percentage of absenteeism</p> <ol style="list-style-type: none"> 1. Group A streptococcal infections in rapid antigen test (time frame: total 20 weeks) 2. Incidence of symptoms of acute upper respiratory tract infection (time frame: total 20 weeks) 3. School absenteeism (time frame: total 20 weeks) <p>Secondary outcome measures: Glo Germ gel applied hands will shine areas containing micro-organisms. Contamination rate will be calculated by taking a photo of the hands and performing brightness analysis in Adobe Photoshop program.</p> <ol style="list-style-type: none"> 1. Pollution rate of hands (time frame: from date of randomisation until the date of first documented progression assessed up to 7 months)
Starting date	9 September 2019
Contact information	Contact: Uyanik +905068949969; gulcinyelten@hotmail.com
Notes	Recruitment is ongoing. Last update in ClinicalTrials.gov was 13 February 2020. NCT04267952

NCT04296643

Study name	Medical masks vs N95 respirators for COVID-19
------------	---

NCT04296643 (Continued)

Methods	A RCT in which nurses will be randomised to either medical masks or N95 respirators when providing medical care to patients with COVID-19. This Canadian multicentre RCT will assess whether medical masks are non-inferior to N95 respirators when nurses provide care involving non-aerosol generating procedures. Nurses will be randomised to either use of a medical mask or to a fit-tested N95 respirator when providing care for patients with febrile respiratory illness. The primary outcome is laboratory confirmed COVID-19 amongst nurse participants.
Participants	<p>Inclusion criteria:</p> <ol style="list-style-type: none"> 1. Nurses who work > 37 hours per week in medical, emergency, paediatric units <p>Exclusion criteria:</p> <ol style="list-style-type: none"> 1. Nurses with 1 or more comorbidities 2. Nurses who cannot pass an N95 respirator fit-test
Interventions	<p>Experimental: medical mask</p> <p>Medical mask worn when providing care to patient with febrile respiratory illness</p> <p>Active comparator: N95 respirator</p> <p>N95 respirator worn when providing care to patient with febrile respiratory illness</p>
Outcomes	<p>Primary outcome measures:</p> <ol style="list-style-type: none"> 1. Number of participants with RT-PCR confirmed COVID-19 infection 2. RT-PCR confirmed COVID-19 infection (time frame: 6 months) <p>Secondary outcome measures:</p> <ol style="list-style-type: none"> 1. Number of participants with acute respiratory illness 2. Number of participants with absenteeism 3. Number of participants with lower respiratory infection 4. Number of participants with pneumonia 5. Number of participants with ICU admission 6. Number of participants needing mechanical ventilation 7. Number of participants that died
Starting date	Started 2 April 2020
Contact information	Contact: Mark Loeb, MD, 9053340010; loebm@mcmaster.ca
Notes	NCT04296643

NCT04337541

Study name	Reduction in COVID-19 infection using surgical facial masks outside the healthcare system
Methods	Parallel RCT
Participants	<p>18 years of age and older. The participants recruited are people working outside of their home, who have not previously been infected with COVID-19 and who do not wear facial masks (e.g. healthcare personnel) when working. They will be randomised for:</p> <ol style="list-style-type: none"> 1. normal behaviour according to the authority's recommendations; or

NCT04337541 (Continued)

2. normal behaviour according to the authority's recommendations and use of facial masks.

Interventions	All participants will follow authority recommendations and be randomised to either wear facial masks or not. They will perform swab-test if they experience symptoms during the study as well as at the end of study. Participants will be instructed in using the facial mask consistently when outside their home (and at home when receiving visits from others). The instruction is given in writing and via an instruction video. The participants will be contacted once weekly to optimise compliance. It will be registered if the participants are diagnosed with COVID-19. Participants will perform antibody screening at study start and end. Participants who do not test positive for COVID-19 during the study period will perform a swab self-test if experiencing symptoms or when the study ends (instruction video).
Outcomes	Primary outcome: reduction in COVID-19 infection frequency Secondary outcome: number of participants testing positive in antibody screening at study start and study end, respectively
Starting date	2 April 2020
Contact information	Prof Henning Bundgaard, DMSc +4526112290; henning.bundgaard@regionh.dk
Notes	NCT04337541. Published <i>Annals of Internal Medicine</i> , https://www.acpjournals.org/doi/10.7326/M20-6817 , 18 Nov 2020).

NCT04471766

Study name	Evaluation of locally produced cloth face mask on COVID-19 and respiratory illnesses prevention at the community level - a cluster-RCT
Methods	Study type: interventional (clinical trial) Estimated enrolment: 66,000 participants Allocation: randomised Intervention model: parallel assignment Masking: single (outcomes assessor) Primary purpose: prevention
Participants	Ages eligible for study: 10 years and older (child, adult, older adult) Sexes eligible for study: all Accepts healthy volunteers: no Criteria Inclusion criteria: 1. Household resident 2. Age 10 years and older Exclusion criteria: 1. Refusal to participate
Interventions	Experimental: certified cloth face mask plus preventive information

NCT04471766 (Continued)

	Active comparator: information on COVID-19 prevention
Outcomes	<p>Self-reported main symptoms of COVID-19 (3 or more of fever, cough, fatigue, shortness of breath, loss of smell/taste)</p> <p>Consultation for COVID-19 like illness or reported positive test, or both</p> <p>Self reported COVID-19 like illness plus hospitalisation or death</p> <p>Any death during the follow-up period:</p> <ol style="list-style-type: none"> 1. Reported COVID-19 like illness (time frame: 4 months' follow-up) 2. Consultation (time frame: 4 months' follow-up) 3. Severe illness (time frame: 4 months' follow-up) 4. Mortality (time frame: 4 months' follow-up)
Starting date	Estimated study start date: July 2020
Contact information	Amabelia Rodrigues, PhD, 00245966078659; a.rodrigues@bandim.org
Notes	<p>The number of cases of COVID-19 is still increasing, and transmission of SARS-CoV-2 seems to occur mainly through person-to-person transmission through respiratory droplets, indirect contact with infected people and surfaces. The use of face masks is recommended as a public health measure, but in many settings only domestic cloth made masks are available to the majority of the people. However, masks can be of different quality, and very little is known about the utility of cloth face masks at the community level.</p> <p>In Bandim Health Project's Health and Demographic Surveillance System we will evaluate the effect of providing locally produced cloth face masks on the severity of COVID-19 like illness and mortality in an urban population. The locally produced cloth mask is made according to a laboratory-certified model and will be provided to the intervention group alongside information of how the risk of transmission can be reduced. The control group will receive information alone.</p> <p>Follow-up will be implemented through telephone calls and postepidemic home visits.</p>

Wang 2015

Study name	A cluster-RCT to test the efficacy of face masks in preventing respiratory viral infection among Hajj pilgrims
Methods	Cluster-randomised trial, randomising worshippers' accommodation tents during several seasons of the Hajj pilgrimage in Saudi Arabia. In the intervention tents, free face masks will be distributed to be worn for 7 days.
Participants	Pilgrims to the Hajj
Interventions	Standard surgical masks distributed free of charge to pilgrims in the intervention accommodation tents. Control group receives no intervention.
Outcomes	Flu-like illness, recorded in diaries, with laboratory confirmation where possible
Starting date	
Contact information	
Notes	Australian New Zealand Clinical Trials Registry (ANZCTR), ACTRN12613001018707

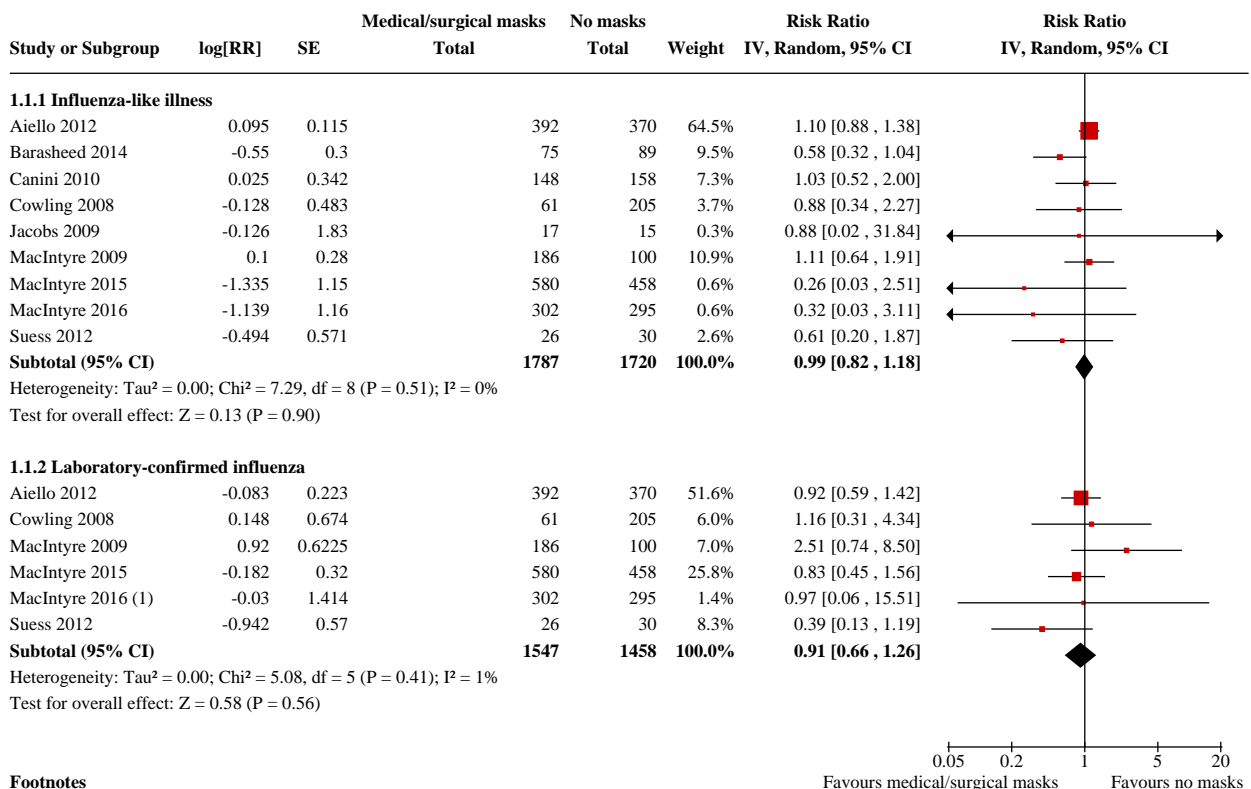
ARI: acute respiratory tract infections
ICU: intensive care unit
ILI: influenza-like illness
GI: gastrointestinal
RCT: randomised controlled trial
RT-PCR: reverse-transcriptase polymerase chain reaction

DATA AND ANALYSES

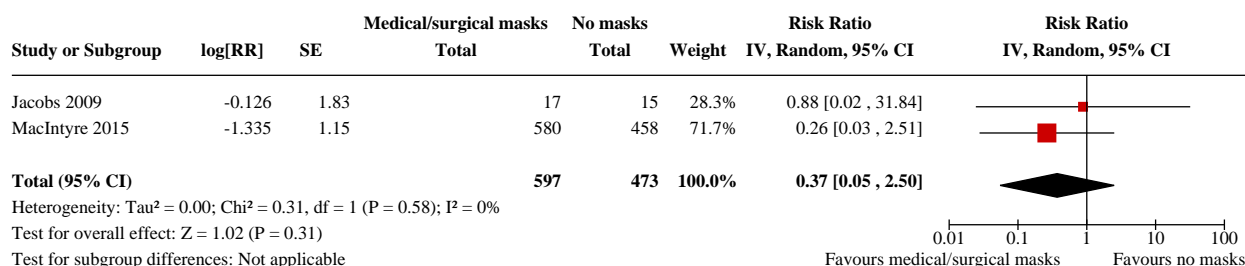
Comparison 1. Randomised trials: medical/surgical masks versus no masks

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1.1 Viral illness	9		Risk Ratio (IV, Random, 95% CI)	Subtotals only
1.1.1 Influenza-like illness	9	3507	Risk Ratio (IV, Random, 95% CI)	0.99 [0.82, 1.18]
1.1.2 Laboratory-confirmed influenza	6	3005	Risk Ratio (IV, Random, 95% CI)	0.91 [0.66, 1.26]
1.2 Influenza-like illness in healthcare workers	2	1070	Risk Ratio (IV, Random, 95% CI)	0.37 [0.05, 2.50]

Analysis 1.1. Comparison 1: Randomised trials: medical/surgical masks versus no masks, Outcome 1: Viral illness



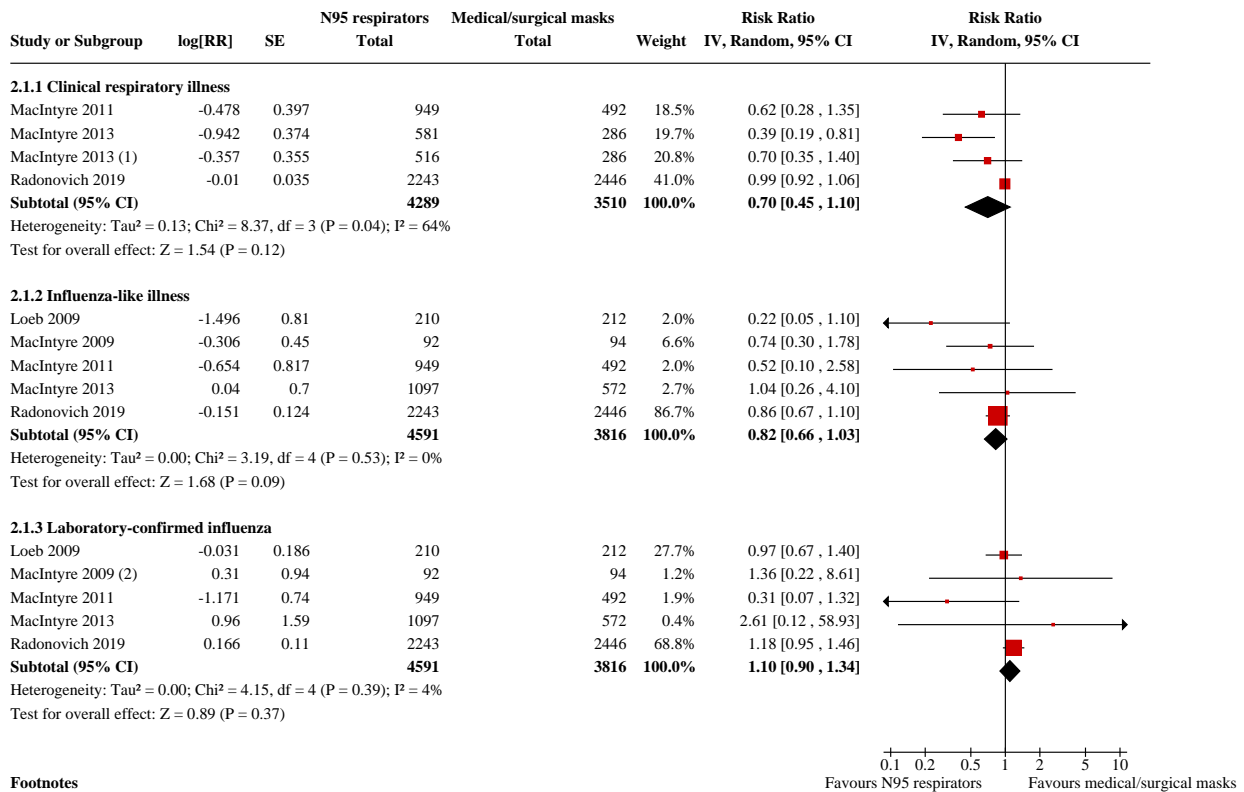
Analysis 1.2. Comparison 1: Randomised trials: medical/surgical masks versus no masks, Outcome 2: Influenza-like illness in healthcare workers



Comparison 2. Randomised trials: N95 respirators compared to medical/surgical masks

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
2.1 Viral illness	5		Risk Ratio (IV, Random, 95% CI)	Subtotals only
2.1.1 Clinical respiratory illness	3	7799	Risk Ratio (IV, Random, 95% CI)	0.70 [0.45, 1.10]
2.1.2 Influenza-like illness	5	8407	Risk Ratio (IV, Random, 95% CI)	0.82 [0.66, 1.03]
2.1.3 Laboratory-confirmed influenza	5	8407	Risk Ratio (IV, Random, 95% CI)	1.10 [0.90, 1.34]
2.2 Viral illness in healthcare workers	4		Risk Ratio (IV, Random, 95% CI)	Subtotals only
2.2.1 Clinical respiratory illness	3	7799	Risk Ratio (IV, Random, 95% CI)	0.70 [0.45, 1.10]
2.2.2 Influenza-like illness	4	8221	Risk Ratio (IV, Random, 95% CI)	0.81 [0.59, 1.11]
2.2.3 Laboratory-confirmed influenza	4	8221	Risk Ratio (IV, Random, 95% CI)	1.05 [0.79, 1.40]

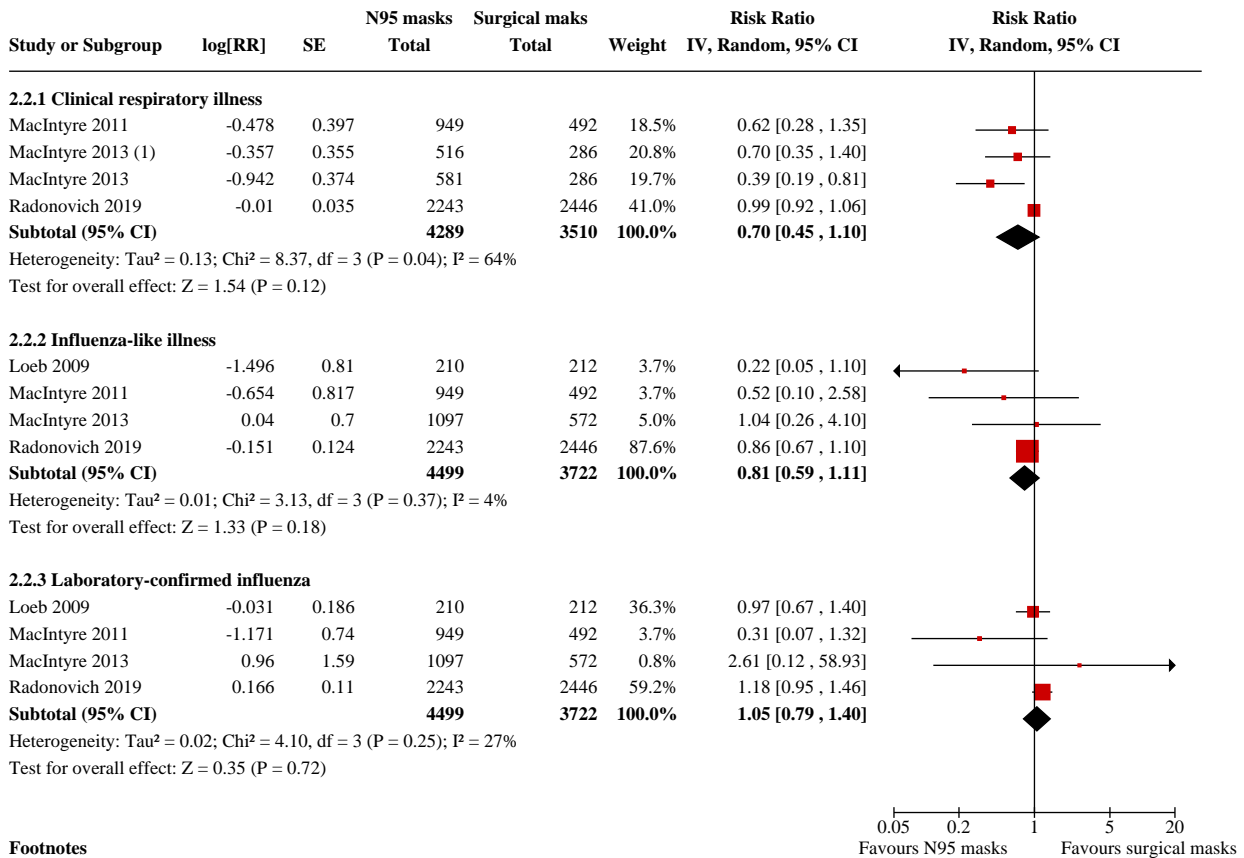
Analysis 2.1. Comparison 2: Randomised trials: N95 respirators compared to medical/surgical masks, Outcome 1: Viral illness



Footnotes

- (1) MacIntyre 2013 includes 2 comparisons: N95 vs surgical masks and targeted N95 vs surgical masks
- (2) MacIntyre 2009 reported on outcome laboratory confirmed infections

Analysis 2.2. Comparison 2: Randomised trials: N95 respirators compared to medical/surgical masks, Outcome 2: Viral illness in healthcare workers



Footnotes

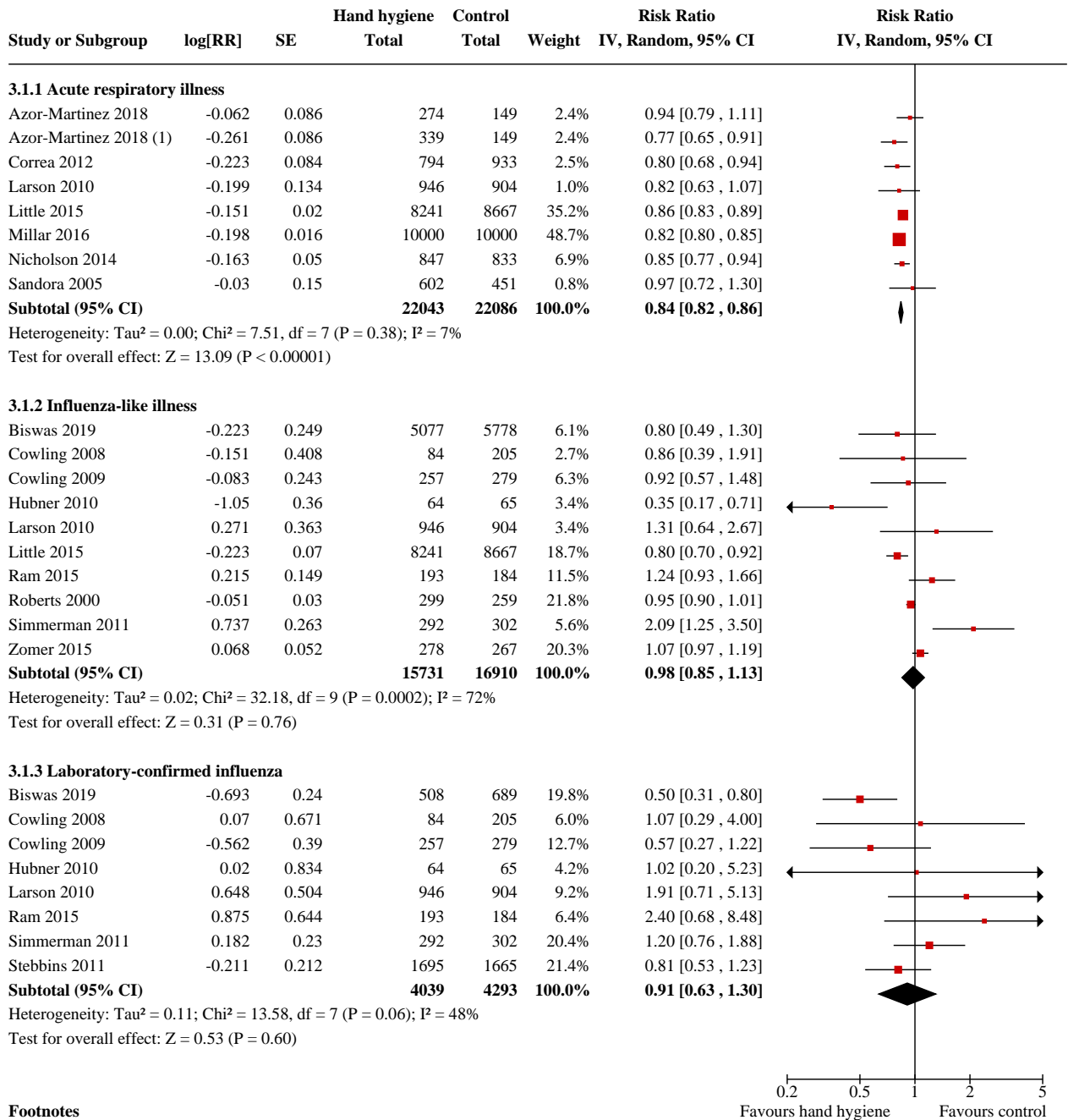
(1) MacIntyre 2013 includes 2 comparisons: N95 vs surgical masks and targeted N95 vs surgical masks

Comparison 3. Randomised trials: hand hygiene compared to control

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
3.1 Viral illness	16		Risk Ratio (IV, Random, 95% CI)	Subtotals only
3.1.1 Acute respiratory illness	7	44129	Risk Ratio (IV, Random, 95% CI)	0.84 [0.82, 0.86]
3.1.2 Influenza-like illness	10	32641	Risk Ratio (IV, Random, 95% CI)	0.98 [0.85, 1.13]
3.1.3 Laboratory-confirmed influenza	8	8332	Risk Ratio (IV, Random, 95% CI)	0.91 [0.63, 1.30]
3.2 ARI or ILI or influenza (including outcome with most events from each study)	16	61372	Risk Ratio (IV, Random, 95% CI)	0.89 [0.84, 0.95]
3.3 Influenza or ILI: sensitivity analysis including outcomes with the most precise and unequivocal definitions	11	26343	Risk Ratio (IV, Random, 95% CI)	0.92 [0.80, 1.05]
3.4 ARI or ILI or influenza: subgroup analysis	16	61372	Risk Ratio (IV, Random, 95% CI)	0.89 [0.84, 0.95]

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
3.4.1 Children	9	21283	Risk Ratio (IV, Random, 95% CI)	0.92 [0.84, 1.01]
3.4.2 Adults	7	40089	Risk Ratio (IV, Random, 95% CI)	0.85 [0.79, 0.92]
3.5 Absenteeism	3	3150	Risk Ratio (IV, Random, 95% CI)	0.64 [0.58, 0.71]

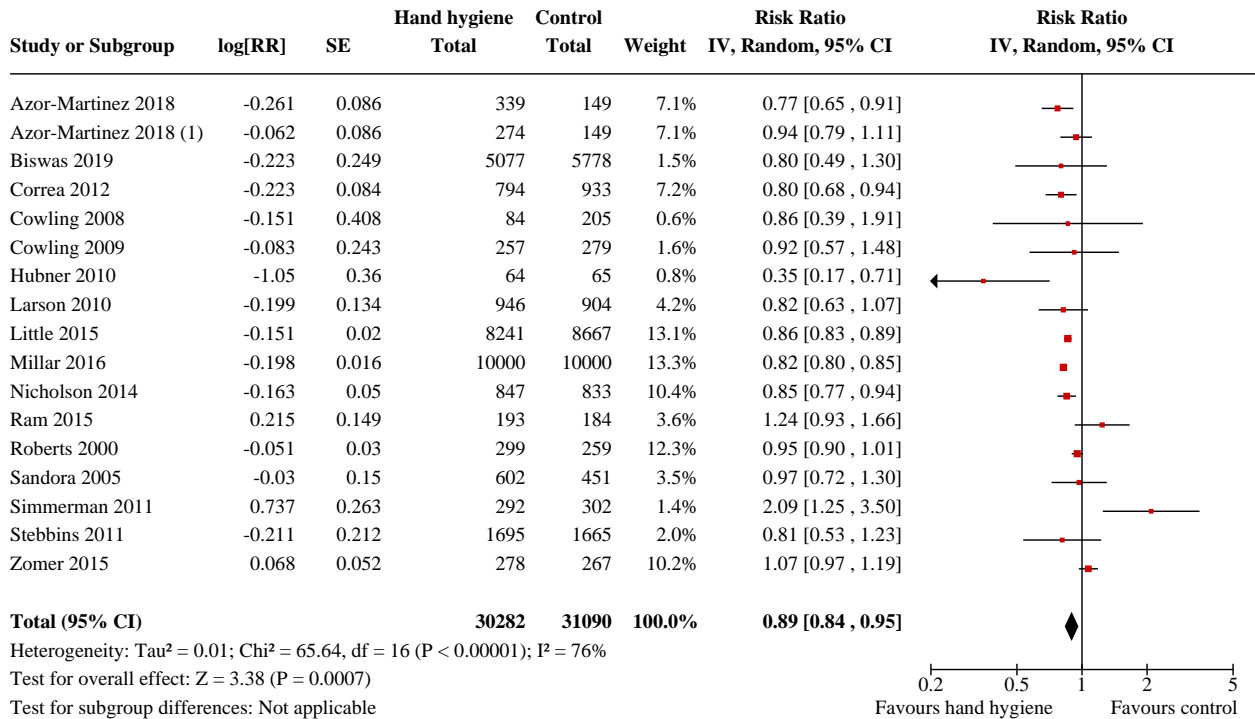
Analysis 3.1. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 1: Viral illness



Footnotes

(1) Azor 2018 included 2 hand-washing groups: one using soap and water (RR 0.94) and the other using hand sanitizer (RR 0.77)

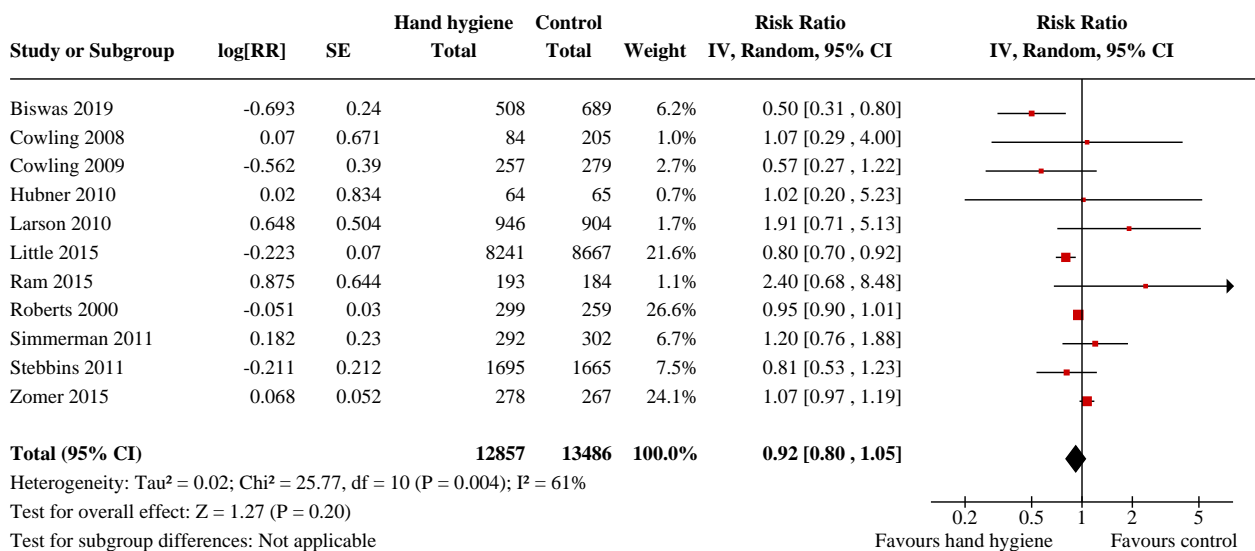
Analysis 3.2. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 2: ARI or ILI or influenza (including outcome with most events from each study)



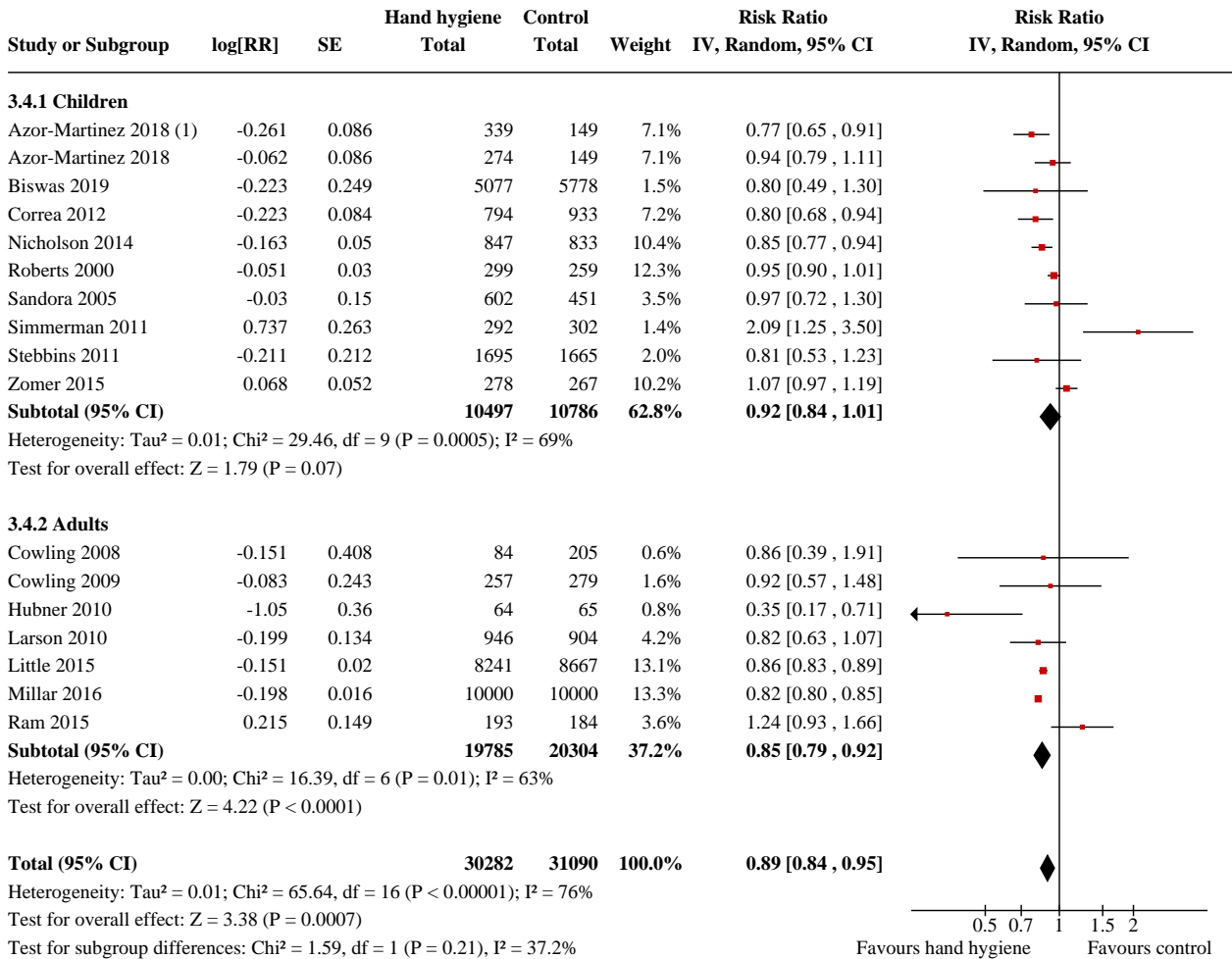
Footnotes

(1) Azor 2018 included 2 treatment groups: soap and water (RR 0.94); and hand sanitizer (RR 0.77)

Analysis 3.3. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 3: Influenza or ILI: sensitivity analysis including outcomes with the most precise and unequivocal definitions



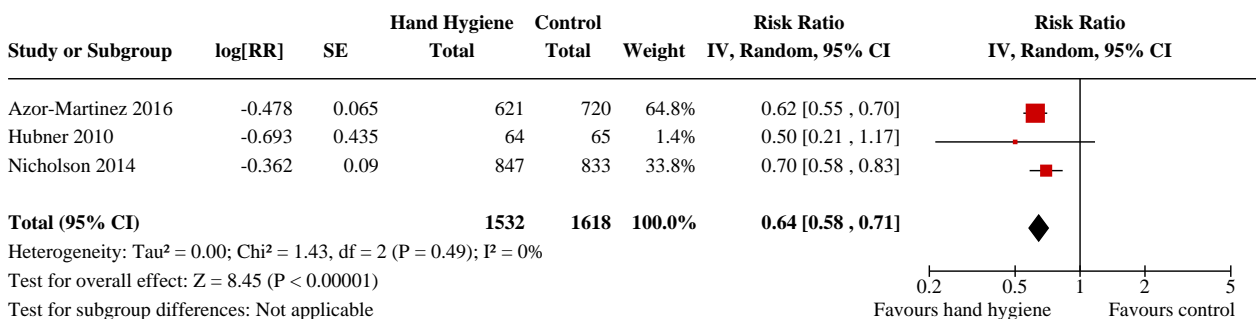
Analysis 3.4. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 4: ARI or ILI or influenza: subgroup analysis



Footnotes

(1) Azor 2018 includes 2 intervention groups: soap and water (RR 0.94) and hand sanitizer (RR 0.77)

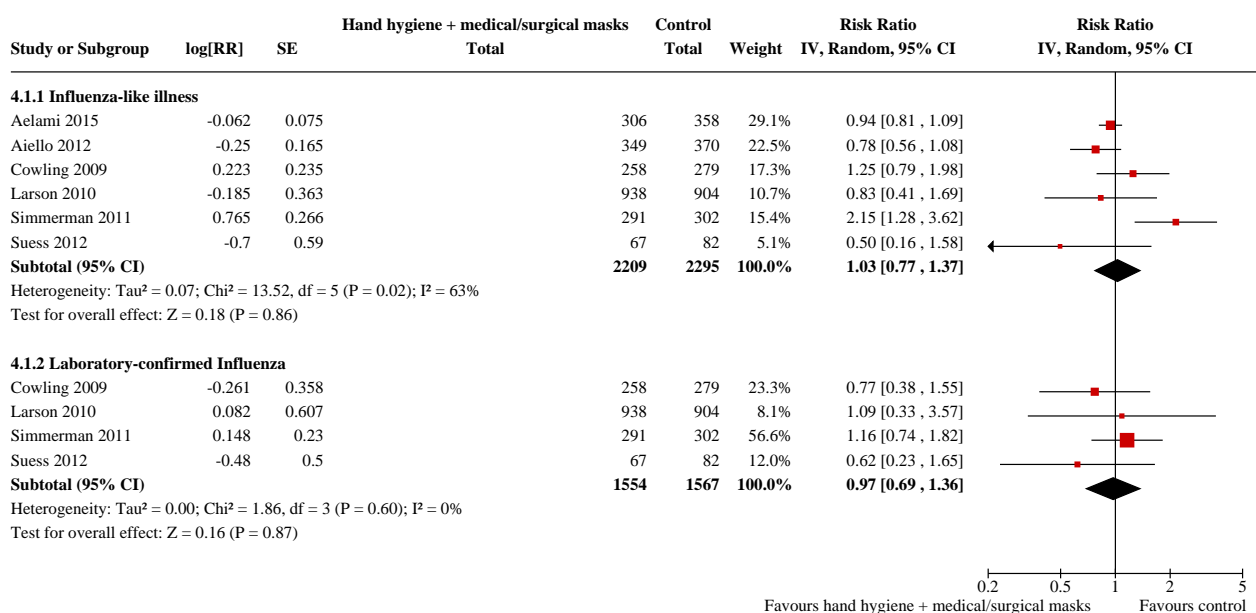
Analysis 3.5. Comparison 3: Randomised trials: hand hygiene compared to control, Outcome 5: Absenteeism



Comparison 4. Randomised trials: hand hygiene + medical/surgical masks compared to control

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
4.1 Viral illness	6		Risk Ratio (IV, Random, 95% CI)	Subtotals only
4.1.1 Influenza-like illness	6	4504	Risk Ratio (IV, Random, 95% CI)	1.03 [0.77, 1.37]
4.1.2 Laboratory-confirmed Influenza	4	3121	Risk Ratio (IV, Random, 95% CI)	0.97 [0.69, 1.36]

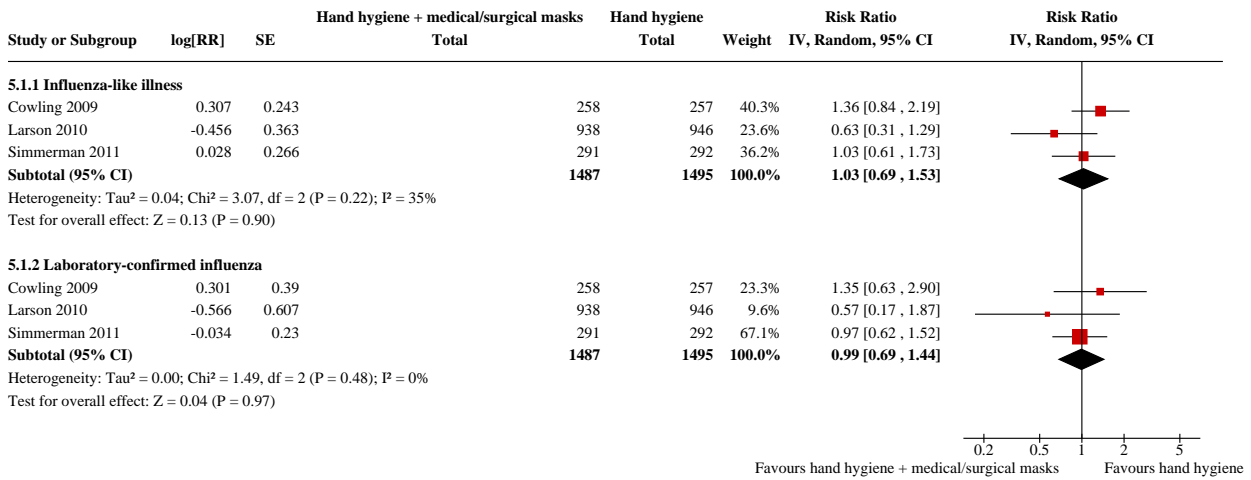
Analysis 4.1. Comparison 4: Randomised trials: hand hygiene + medical/surgical masks compared to control, Outcome 1: Viral illness



Comparison 5. Randomised trials: hand hygiene + medical/surgical masks compared to hand hygiene

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
5.1 Viral illness	3		Risk Ratio (IV, Random, 95% CI)	Subtotals only
5.1.1 Influenza-like illness	3	2982	Risk Ratio (IV, Random, 95% CI)	1.03 [0.69, 1.53]
5.1.2 Laboratory-confirmed influenza	3	2982	Risk Ratio (IV, Random, 95% CI)	0.99 [0.69, 1.44]

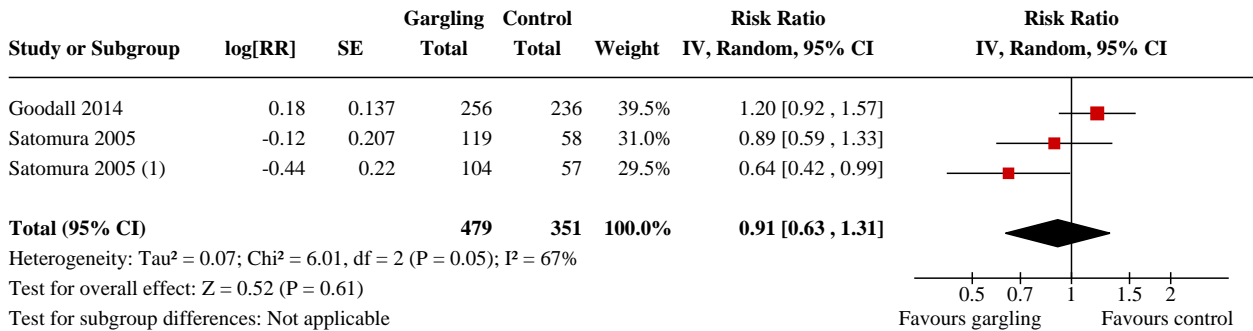
Analysis 5.1. Comparison 5: Randomised trials: hand hygiene + medical/ surgical masks compared to hand hygiene, Outcome 1: Viral illness



Comparison 6. Randomised trials: gargling compared to control

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
6.1 Viral illness	2	830	Risk Ratio (IV, Random, 95% CI)	0.91 [0.63, 1.31]

Analysis 6.1. Comparison 6: Randomised trials: gargling compared to control, Outcome 1: Viral illness



Footnotes

(1) Satomura 2005 included 2 intervention groups

ADDITIONAL TABLES

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist

Author, year	Brief name	Recipient	Why	What (materials)	What (procedures)	Who provided	How	Where	When and how much	Tailoring	Modification of intervention through-out trial	Strategies to improve or maintain intervention fidelity	Extent of intervention fidelity
Masks compared to either no masks or different mask types													
Barashe 2014	Su-pervised mask use	Religious pilgrims ≥ 15 years	Prevent respiratory virus infections at mass gatherings through mask use	Plain surgical face masks (3M Standard Tie-On Surgical Mask, Cat No: 1816) manufactured by 3M company, USA; 5 masks per day Written instructions on face mask use Special polythene bags for disposal	Masks provided to index case and their contacts with advice on mask use (before prayers, in seminars, and after meals). Written instructions provided on face mask use, need to change them, and disposal.	Not described, presumably the medical researchers	Face-to-face provision of masks, Valleys, Saudi Arabia and reminders	Tents of pilgrimage site (Mina, Jeddah, Saudi Arabia)	Advice on mask use given throughout pilgrimage stay (5 days)	None reported.	None reported.	The medical researchers followed pilgrims each day to remind participants about recording their mask usage in health diary.	Face mask use: mask group: 56/75 (76%), control group: 11/89 (12%) (P < 0.001) 76% of intervention tents wore masks. 10 of 75 (13%) pilgrims in 'mask' tents wore face masks during sleep.
Canini 2010	Surgical face masks	Householders (over 5 years)	Limit transmission of influenza by large	Initial supply of 30 masks: for adults and children > 10: surgery masks with earloops, 3 plys, anti fog (AEROKYN, LCH medical products, Paris, France)	Masks given immediately on home visit by attending general practitioner with demonstration of proper use	General practitioners	Face-to-face individually	Households in France	One-off provision of masks worn for 5 days	None described.	None described.	Not described, but reported mask usage was measured	34/51 (66%) wore masks > 80% of the duration.



Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

			droplets produced during coughing in households	Children 5 to 10: face mask KC47127, (Kimberly-Clark, Dallas, TX, USA) Closed plastic bags for disposal	and instruction to be worn for 5 days in presence of another household member or in confined space (e.g. car) and to change every 3 hours or if damaged.								Reported mask-wearing: 11 ± 7.2 masks during 4.0 ± 1.6 days with an average use of 2.5 ± 1.3 masks per day and duration of use of 3.7 ± 2.7 hours/day
Ja-cobs 2009	Face masks	Hospital health-care providers (nurses, doctors, and medical personnel)	Decrease risk of infection through limiting droplet spread through masks	Hospital-standard disposable surgical Mask MA-3 (Ozu Sangyo, Tokyo, Japan); quantity not specified	Provision of masks and instructions for use	Not described, presumably research team	Face-to-face	Tertiary care hospital in Tokyo, Japan Face masks worn whilst on hospital property.	77 days	None described.	None described.	Self-reported compliance	Self-reported compliance for both groups reported as good, with full compliance by 84.3% and remainder complying 79.2% to 98.7%.
Loeb 2009	2 active interventions A. surveillance	Health-care workers (nurses)	Reduce transmission of influenza in healthcare settings through coughing	A. Surgical masks B. N95 respirators	Provision of masks or N95 respirators Instruction in use and proper placement of devices	Provided by research team (not further	In-person face-to-face	Tertiary hospitals in Ontario,	1 influenza season (12 weeks) Use of mask as required[2] when pro-	Fit-testing of nurses not already	Ceased before season	Compliance audits during peak of season by trained auditor who stood	18 episodes: N95: 6/7 participants (85.7%) wearing assigned

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	gical masks B. N95 respirators	or sneezing with protective masks	Fit-testing and demonstration of positioning of N95 using standard protocol and procedure (details provided)	Qualitative fit-testing using saccharin or Bitrex protocol ^[1]	de-scribed) Fit-testing by technician for N95	Canada	viding care to or within 1 m of patient with febrile respiratory illness, $\geq 38^\circ\text{C}$, and new or worsening cough or shortness of breath Nurses to wear N95 when caring for patients with “febrile respiratory illness”	fit-tested	short distance from patient isolation room	device versus 100% for masks			
MacIntyre 2009	2 active interventions in addition to infection control guidelines A. Surgical masks (SM)	Householders with a child with fever and respiratory symptoms	Prevent or reduce respiratory virus transmission in the community through non-pharmaceutical interventions	A. 3M surgical mask, catalogue no. 1820; St Paul, MN, USA for adults B. P2 masks (3M flat-fold P2 mask, catalogue no. 9320; Bracknell, Berkshire, UK) A and B: health guidelines and pamphlets about infection control	Provision of masks and pamphlets and education about infection prevention and mask use Telephone calls and exit interviews to record adherence to mask use All groups: health guidelines, pamphlets about infection control were provided	Not described, presumably research team	Face-to-face and telephone	Households in Sydney, Australia	2 winter seasons (3 months and 6 months) 2 weeks of follow-up Masks to be worn at all times when in same room as index child, regardless of distance from child	None described.	None described.	Daily telephone calls to record mask use throughout day Exit interviews about adherence	Reported mask use: Day 1 SM: 36/94 (38%) P2: 42/92 (46%) stated wearing “most or all” of the time. Other participants were wearing face masks rarely or never. Day 5:

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	B. P2 masks (P2)												SM: 29/94 (31%) P2: 23/92 (25%)
MacIntyre 2011	3 active interventions A. Medical masks B. N95 respirators fit-tested C. N95 respirators non-fit-tested	Health-care workers	Protect HCWs by preventing transmission of influenza and other respiratory viruses from patients through mask wearing	Daily supply of A. 3 medical masks (3M catalogue number 1820, St Paul, MN, USA) 2 respirators: B. N95 fit-tested mask (3M flat-fold N95 respirator, catalogue number 9132) fit-tested with 3M FT-30 Bitrex Fit Test kit according to manufacturer's instructions (3M, St Paul, MN, USA) C. N95 non-fit-tested mask (3M flat-fold N95 respirator, catalogue number 9132) Diary cards for usage recording	Supply of masks or respirators. Instruction in when to wear it, correct fitting, and storage (in paper bag in personal locker) Instruction in importance of hand hygiene before and after removal For fit-tested group: fit-testing procedure	Masks provided to hospitals. Training of staff provided by 1 member of research team.	Masks and training provided to face, not described if training was individually or in groups.	Emergency departments and respiratory wards in Beijing, China	Entire work shift for 4 weeks	Taken off for toilet and meal breaks and at end of shift	None described.	Mask/respirator use monitored by: (i) observed compliance by head ward nurse recorded daily; (ii) self-report diary cards carried during day recording; (i) no. hours; (ii) usage. Exit interviews	Adherence for usage was high for all and not significantly different amongst arms. Medical mask: 76%, 5 hours N95 fit-tested: 74%, 5.2 hours N95 non-fit-tested: 68%, 4.9 hours
MacIntyre 2013	3 active interventions A. N95 respirators at all times	Health-care workers (nurses and doctors)	Protect HCWs from respiratory infections through mask use	Daily supply of: A. and B. 2 respirators (3M Health Care N95 Particulate Respirator; catalogue number 1860) 3M FT-30 Bitrex Fit Test Kit C. 3 masks 3 masks	Supply of respirators Instructions in use including times and fit Fit-testing procedure according to the manufacturer's instructions (3M) For targeted N95:	3M supplied respirators and masks. Provider instructions	Masks and training provided to face, not described	Emergency departments and respiratory wards of tertiary	For 4 weeks, A and B worn at all times on shift; B. targeted (intermittent) use of N95 respirators only whilst	None described.	None described.	Self-reported daily record of number of hours worked, mask or respirator use, number of high-	Compliance highest for targeted N95 (82%; 422/516) versus N95 (57%; 333/581) versus medical mask

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	B. N95 respirators targeted use C. Medical masks			(3M Standard Tie-On Surgical Mask catalogue number mask 1817; 3M, St Paul, MN, USA) Pocket-sized diary card with tick boxes for mask use	checklist of defined high-risk procedures, including common aerosol-generating procedures	not specified.	if training was individually or in groups.	tiary hospitals in Beijing, China	performing high-risk procedures or barrier.			risk procedures undertaken collected by study staff.	(66%; 380/572).
MacIntyre 2015	2 active interventions A. Cloth masks B. Medical masks	Hospital health-care workers	Prevent respiratory infections in HCWs from patients through mask-wearing	A. 5 cloth masks for study duration (2-layer, cotton) B. 2 medical masks daily for each 8-hour shift for study duration (3 layers, non-woven material) All masks locally manufactured. Written instructions on cleaning cloth masks	Cloth or medical masks to be worn at all times on shift. Cloth masks to be washed with soap and water daily after shifts, and the process of cleaning to be documented. Provision of written instructions for cloth mask cleaning	Researcher arranged supply of masks and instructions and any training of staff assisting the delivery.	Masks and written instructions provided to face.	Hospital wards in Vietnam	4 weeks (25 days) of face mask use	Masks not worn while toilet or during tea or lunch breaks.	None described.	Monitored compliance with mask use by self-report diary card and exit survey and interviews with a sub-sample (AC-TRN1261000023/25)	Mask-wearing compliance: cloth mask: 56.8%; medical mask: 56.6%; Reported cloth mask washing: 92%
MacIntyre 2016	Medical mask use	Sick householders with ILI (index of res-	Protect well people in the community from transmission of res-	21 medical masks (3M 1817 surgical mask) Diary cards for mask use	Supply of masks Instructions for mask wearing and hand-washing protocol Provision of diary cards	Study staff member provided masks and	Masks and instructions provided face-	Fever clinics of major hospitals in	3 masks/day for 21 days Mask wearing: whenever in the same	Allowed to remove their masks during	None reported.	Self-reported daily record of mask use using diary card	Mask use: mask group: 4.4 hours; control group: 1.4 hours

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	cas- es) and their well con- tacts of the same house- hold	piratory pathogens by con- tacts with ILI through mask use				in- struc- tions in use.	to- face and indi- vidually.	Bei- jing, Chi- na	room as a household member or a visi- tor to the household Hand- washing: before putting on and after taking off	meal- times and whilst asleep and to cease wear- ing once symp- toms re- solved		
Radonovitch 2019	Health- care person- nel of out- patient sites with in medical centres (MM)	Prevent HCP from acquiring workplace viral respi- ratory in- fections and trans- mitting them to others by effective respira- tory pro- tection by N95 respira- tors which reduce aerosol exposure and in- halation of small airborne particles, meet fil- tration	A. N95 respirators: 3M Corporation 1860, 1860S, and 1870 (St Paul, MN, USA) or Kim- berly Clark Technol Flu- idshield PFR95-270, PFR95-274 (Dallas, TX, USA) B. Medical mask Precept 15320 (Arden, NC, USA) or Kimberly Clark Technol Fluidshield 47107 (Dal- las, TX, USA). Reminder signs posted at each site A portable comput- er equipped with da- ta recording software (HandyAudit; Toronto, Canada) to document	Participants in- structed to wear assigned pro- tective devices whenever they were positioned within 6 feet (1.83 m) of patients with sus- pected or con- firmed respiratory ill- ness and to don a new N95/MM with each patient in- teraction. Hand hygiene recommended to all participants in accordance with Centers for Disease Control	Centres pro- vided de- vice sup- plied by study to HCP. Study per- son- nel post- ed re- va- minder signs and emails and con- duct- ed ad-	Face- to- face indi- vid- ual pro- vi- sion of de- vices and ad- her- ence ob- ser- va- tions On- site post- ing of signs Oth- er re- mind- ers	Out- pa- tient sites with- in med- ical cen- tres in USA	As in- structed, for each new pa- tient in- teraction during 12- week peri- od of peak viral respi- ratory ill- ness each year for 4 years (to- tal of 48 weeks)	Fit- ting of N95 masks	None de- scribed.	Reminder signage posted at study sites, and emails sent by study per- sonnel. Self-re- ported daily de- vice wear- ing of "al- ways", "some- times", "never", or "did not recall" Observa- tion of de- vice wear- ing behav- iours as parti- cipants en- tered and	Device wearing: N95: 89.4% re- ported "al- ways" or "some- times" versus MM: 90.2% "Never" N95: 10.2% MM: 9.5%

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

requirements, and fit tightly	adherence (Radonovich 2016)	and Prevention guidelines.	her-ence ob-ser-vations.	by email		exited care rooms conducted during unannounced, inconspicuous visits to randomly selected sites documented on portable computer
		Infection prevention policies				
		were followed at each study site.				
		Reminder signs posted at sites and emails sent.				
		Annual fit-testing conducted for all participants.				
		Filtration testing performed on the device models in the study. Further details in protocol (Radonovich 2016).				

Hand hygiene

Alz-her 2018	Hand hygiene workshop	Pri-ary school girls	Target-ed school children to improve hand hygiene to reduce school absences due to up-per res-pirato-ry infec-tion and spread of infection in schools	6-minute video-clip of 2 siblings that attended school-based health education about hand hygiene Short interactive lecture about: common infections in schools, methods of transmis-sion, hand-washing pro-cedure using soap and water including when to wash hands	Delivery of work-shop and distri-bution of sup-ported materi-als (games and posters) to school and students	Study in-vesti-gator deliv-ered work-shop.	De-liv-ered face-to-face in group for the work-shop	2 pri-ary girls' schools in Sau-di Ara-bia	1-hour once-off workshop; posters and games provided to school	Not de-scribed	Not de-scribed	Posters in restrooms as re-minders of hand-washing hygiene during 5-week fol-low-up pe-riod after workshop	Not re-ported
--------------	-----------------------	----------------------	---	--	---	--	---	--	--	----------------	----------------	--	---------------

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

			and to families	Puzzle games related to hand hygiene									
				Posters with cartoon princesses' picture promoting hand-washing									
Arbo-gast 2016	Multimodal hand hygiene intervention programme in addition to control of brief video	Office buildings and employees of insurance company	Reduce hand-to-mouth germ transmission from shared workspaces and workplace facilities and thereby healthcare claims and absenteeism through improved workplace hand hygiene	Alcohol-based hand sanitiser (PURELL Advanced, GOJO Industries Inc, Akron, OH, USA) installed as wall-mounted dispensers, stands, or free-standing bottles One 8-ounce bottle of hand sanitiser (PURELL Advanced) per cubicle One 100-count canister of hand wipes (PURELL Wipes) per cubicle Replenishment products stored in supply room (in addition to existing foam hand wash (GOJO Green Certified Foam Handwash) and an alcohol-based hand sanitiser foam wall-mounted dispenser (PURELL, GOJO Industries) already provided near the restroom exits prior to intervention) Identical soap in all restrooms Intervention and control group: brief (< 1-minute educational video) about	Hand hygiene supplies installed in offices. Replenishment product was made easily available to individual employees upon request via a simple process. Monitoring of product shipments into sites Physical collection and full replacement of soap, sanitiser, and wipes Intervention and control group: educational video embedded at end of baseline online knowledge survey	Not described, presumably investigation and installations	Hand hygiene supplies provided in office environments and individually at staff cubicles/offices. Video provided individually via email.	High-traffic common areas of 2 US health insurance companies (e.g. near elevators, at entrances) and appropriate public spaces (e.g. coffee area, break rooms,	13.5 months overall One-off email video 11 days before study hand hygiene supplies installed. 13 months of provision of supplies 2 times evening collection and full replacement of products	Sanitizer installed in high-use areas of offices.	Not described	Employee survey at 4 months included questions about hand hygiene practice compliance. Monitoring of product shipments into the sites and physical collection of the soap, sanitiser, and wipes 2 times in the study; collected samples were measured and usage rates were estimated	Intervention group employees: reported 40% more cleaning of work area regularly; significantly more likely to keep the hand sanitiser with them and use it throughout the day; significant increases in hand sanitiser use for at-risk activities ^[3] Estimated use by average employee from sample collection:

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

parents and teachers	and to families through non-pharmaceutical intervention of hand-washing programme in schools	Stories, songs, and classroom posters about hand hygiene and infection transmission Hand sanitiser (ALCO ALOE GEL hand sanitiser by Americo Góvantes Burguete, S.L. Madrid, Spain containing 0.2% chlorhexidine digluconate, 1% phenoxylethanol, 0.1% benzalkonium chloride, 5% denat ethyl alcohol, excipients quantity sufficient for 100 mL alcohol 70%, pH 7.0 to 7.5) Informational poster about when and how to wash hands Written and verbal guidance to teachers, parents, and students on properties, possible side effects, and precautionary measures for gel use and storage	Workshop provided for pupils and teachers: frequent infections in schools, transmission and prevention, instructions on correct hand-washing (water and soap, soaping > 20 s, drying hands), use of hand sanitisers and possible side effects Classroom activities linked to hand hygiene and infection transmission Reinforcement of hand hygiene by teachers Hand sanitiser dispensers fixed to walls with an informational poster about hand-washing Supervision of younger children when using hand sanitiser and administration of sanitiser if needed Instruction of children in hand-washing proce-	struction. Workshop and verbal and written information presumably provided by the study research assistant. Classroom activities provided by research assistant and teachers. Supervision	ual parents. Workshops and classroom activities delivered in groups face-to-face. Teacher reinforcement of hand hygiene provided to class face-to-face. Hand sanitiser use supervised was	tails not provided) 2-hour workshop held 1 month before study commencement Fortnightly classroom activities As required, teacher supervision and administration of hand sanitiser Daily reinforcement of hand hygiene by teachers	hand sanitiser dispensers 2-hour workshop held 1 month before study commencement Fortnightly classroom activities As required, teacher supervision and administration of hand sanitiser Daily reinforcement of hand hygiene by teachers	struction of hand sanitiser as needed by teachers, especially for younger children	Fortnightly support by research assistant promoting hand-washing Self-reported correct hand-washing procedure (water and soap, soaping > than 20 s, drying hands)	sis but not separately reported.
----------------------	--	---	---	---	---	--	---	--	--	----------------------------------

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

															dures after toilet and when dirty and correct hand sanitiser use ^[4]	and admin-istratio-n of hand san-iti-ser for younger chil-dren by teach-ers	pro-vided indi-vidu-ally and face-to-face.
Azor-Martinez 2018	Edu-cation-al and hand hy-giene pro-gramme	Day care cen-tres and their at-tend-ing chil-dren, their par-ents, and DCC staff	Prevent trans-mission of respira-tory infec-tions by im-proved hand hy-giene of children, par-ents, and staff through hand-washing prac-tices and use of hand sani-tiser due to its bac-tericide and viru-cide prop-erties	A. Liquid soap (no spe-cific antibac-terial com-pon-ents (pH = 5.5)) OR B. Hand sani-tiser (70% ethyl alcohol (pH = 7.0 to 7.5)) for home use and in dis-pensers for school class-room	Installation of liq-uid soap or hand saniti-ser dis-pensers in class-rooms Supervision and admin-istration of hand sani-tiser if re-quired	Work-shop deliv-ered by re-searchers	Work-shops deliv-ered face-to-face in groups to par-ents and staff.	Class-room of DCCs (in Spain) for child inter-ven-tions	8 months overall Initial 1-hour work-shop 1 month be-fore study com-mence-ment	Ad-min-istra-tion of hand saniti-ser in the case of young chil-dren	Not de-scribed	Not de-scribed	Not de-scribed	Reported that no moni-toring of compli-ance through con-tinu-ous obser-vation of hand hy-giene	behav-iours was done, but amount of hand sani-tiser was measured	Families or DCC staff, or both, used 1660 L of hand sani-tiser, esti-mated use by each child of dose 6 to 8 times/day.	
	2 ac-tive inter-ven-tions: A. soap and wa-ter B. hand sani-tiser			Workshop content hand-out Stories, songs, and posters about hand hy-giene and infec-tion trans-mission	3 hand hy-giene work-shops for par-ents and DCC staff: 1. Hand-washing prac-tices, hand saniti-ser use, pos-sible side ef-fects and pre-caution-ary mea-sures (HSG only) 2. RIs and their treat-ments	Re-search as-sis-tant pro-vided hand hy-giene ma-ter-ials to DCCs and par-ents. Par-ents and staff	Work-shop con-tent em-ailed to at-tendees indi-vid-u-ally.	Work-shops pro-vid-ed at DCCs.	3 fur-ther iden-ti-cal ses-sions/DCC provided again 1 month apart	DCC staff could at-tend train-ing at other DCC if un-able to at-tend							

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					3. Fever	Instructions to children, parents, and DCC staff on usual hand-washing practices and protocol [5]	Classroom activities (stories and songs) about hand hygiene and infection transmission	super-vised and ad-min-istered sani-tiser where indi-cat-ed.	Indi-vid-ual face-to-face su-per-vi-sion of hand sani-tiser use, as in-di-cated	One-off installa-tion of dis-pensers	at own DCC.		
Biswas 2019	Hand sanitiser and respi-ratory hy-giene educa-tion	Pri-mary schools and their stu-dents and staff	Reduce commu-nity-wide influ-enza virus trans-mission by improv-ing hand-washing and respi-ratory hy-giene and use of sani-tiser in school-children as contrib-utors to commu-nity-wide virus trans-mission	Hand sanitiser (63% ethyl alcohol) in colourless, transparent 1.5-litre local plastic bot-tles (manufactured by a local pharmaceutical company and was avail-able commercially in Bangladesh (price: USD 5.75/L)) Video clip on respiratory hy-giene practices Behavioural change ma-terials – 3 colour posters (see Appendix of paper) Curriculum materials for hy-giene classes	Installation of hand sanitiser in wall dispensers in all classrooms and outside all toilets, refilled by field staff as needed Encouragement of use of sanitiser at 5 key times during the day ^[6] Hand and respi-ratory hy-giene education provid-ed. ^[7] Integration of hy-giene messages into school’s hy-giene curriculum Delivery of video clip on respira-	Select-ed teach-ers re-spon-sible for dis-semin-ation of in-ter-ven-tion mes-sages through-out were trained over 2 days	Hand sani-tiser and educa-tion ma-terials pro-vided to schools. Ed-uca-tion pro-vided in class-rooms in groups and	Pri-mary schools (in Bangladesh). Sani-tis-er in each class-room and out-side toi-lets Ed-uca-tion in class-room	10 weeks Interven-tion mes-sages con-veyed in class-rooms 3 times/ week.	Re-fills pro-vid-ed as need-ed.	Not de-scribed	Structured field ob-servation by 2 field staff of 5 hours/ school observ-ing hand-washing and respi-ratory hy-giene be-haviours of children at 2 differ-ent loca-tions in a classroom or outside Every oth-er day, field staff measured the level	Hand-washing observed opportu-nities: IG 604/921 (66%) vs CG 171/802 (21%) Hand sani-tiser used in 91% of observed hand-washing events in inter-vention schools. Average

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

ry hygiene practice	in these messages,	face-to-face.	of hand sanitiser in the morning and in the afternoon to calculate amount of hand sanitiser used/day/school and enrolled children.	consumption of hand sanitiser/child/day: 4.3 mL Observation of proper cough or sneeze etiquette: IG: 33% vs CG: 2%
Behaviour change materials distributed and placed around schools.	Behaviour change communication, sanitiser use, and practices for preventing spread of respiratory secretions.			
Use of sanitiser by classroom teachers after training	Classroom teachers conveyed intervention messages during regular			
Training of selected teachers in consultation with head of school and management committee in key messages				
Communication of key messages by the selected teachers to other teachers				

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

														hy- giene class- es.
														Field staff re- placed sup- plies as need- ed.
Cor- rea 2012	Alco- hol-based hand rubs	Child- care centres and their staff and child- ren	Reduce in- cidence and trans- mission of infection in child- ren by improved hand hy- giene where water is scarce includ- ing pro- vision of ABH and training in hand hygiene teach- ing tech- niques	Dispensers of alco- hol-based hand rubs with ethanol 62.0% (PURELL, GOJO Indus- tries, Akron, OH, USA) Workshop materials ^[8] Visual reminders on ABH techniques in bath- rooms and next to dis- pensers	ABH and training on proper use to staff and children Pre-trial ABH use workshop to teachers that followed rec- ommended HH teaching tech- niques and in- structed teach- ers to add ABH to routine HH and give preference to hand-wash- ing with soap and water if hands visibly soiled Continuous refill- ing of ABH ABH technique refresher work- shops (8/centre) Monitoring of safety, proper use	Local rep- re- sen- ta- tive of GO- JO In- dus- tries Inc. pro- vided dis- pensers and dis- penser in- stal- la- tions free of charge.	Face- to- face train- ing and pro- vi- sion of ma- teri- als; group train- ing	Child- care centres in Colom- bia (cen- tres or com- mu- nity homes) ABH in cen- tres, class- rooms, and com- mon areas de- pend- ing on size	8 months overall 1 ABH dis- penser per centre with < 14 children; 1 per classroom in larger centres; 1 per class- room + 1 for com- mon areas in centres with > 28 children 1 work- shop pre- trial to staff Monthly 30-minute ABH tech- nique re-	Re- filled ABH as need- ed	Not de- scribed	Visual re- minders and monthly refresher training Monitor- ing of safe- ty, prop- er use of ABH, amount of ABH used Se- mi-struc- tured sur- vey on comple- tion of teachers' percep- tions about changes in HH prac- tices and use of	Teachers at 7 interven- tion cen- tres re- ported al- most complete substi- tution of HSW with ABH, and HSW de- creased from 3 times per day to 1 per day, and ABH rose to 6 per day. Teach- ers at re- maining 14 centres report- ed par- tial substi-	

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					of ABH, amount of ABH used	Field-work team delivered other components.		Visual reminders (8 per centre) in bathrooms and next to dispensers	fresher training (Biweekly monitoring)		HSW and ABH. Measurement of consumption of resources and costs related to ABH use and HSW	tution of HSW with ABH. Controls reported HSW 3 times per day. Median number of ABH applications per child rose from 3.5 to 4.5 in preschools and 3.5 to 5.5 in community centres.	
Di-Vita 2011	Household handwashing promotion	Household index patient with ILI	Prevent influenza transmission in households in resource-poor settings through provision of handwashing facilities and use of them at critical	Hand-washing stations with soap	Provision of hand-washing stations	Hand-washing motivation to wash at critical times for pathogen transmission (e.g. after coughing or sneezing)	Not specifically described, presumably the researcher	Face-to-face provision of facilities in households	Household in Bangladesh	Over 2 influenza seasons	Not described	Not described	Not described
										One-off provision of handwashing facilities			
										Frequency of "motivation" not described			

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

			times for pathogen transmission				de-scribed						
Feldman 2016	2 active interventions	Naval ships and their sailors	Reduced infection transmission and improved hand hygiene in sailors who are at increased risk due to closed environments, contact with shared surfaces, and poor HH culture	Septadine solution (Floris, Misgav, Israel) 70% alcohol and 0.5% CHG; inactive materials: purified water, glycerin, propylene glycol, and methylene blue	Installation of CHG disinfection devices on ships alongside regular soap and water	Provision of CHG presumably by study team and funds	CHG sent to ships directly.	Navy fast missile boats and patrol boats of naval base in Israel	4 months	CHG replenished on demand.	Not de-scribed	Total amount of CHG dispensed was tallied.	Mean volume CHG: 8.2 mL per sailor per day (projected yearly cost USD 45 per sailor)
	A. Hand disinfection with chlorhexidine gluconate + hygiene education				Supply and replenishment of CHG (sent to ships regardless of replenishment demands)	Hygiene instruction by a naval physician (to both intervention groups and study control group)	Mode of hygiene instruction not de-scribed.	Dis-pensers installed in key locations on-board (ad-jacent to heads (toi-lets), mess decks (din-ing rooms), com-mon	Unlimit-ed sup-ply of CHG replen-ished on demand for 4 to 5 months.	Automatic amount dispensed: 3 mL			
	B. Hygiene education												

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Gwaltney 1980	<p>A. Virucidal hand preparation</p> <p>B. Placebo (no control)</p>	<p>Healthy young adults</p>	<p>Reduce infection rates by interrupting viral spread by hand or self-inoculation route</p>	<p>A. Virucidal hand preparation: aqueous iodine (2% iodine and 4% potassium iodide)</p> <p>B. Placebo: aqueous solution of food colours (Kroger; Kroger Co., Cincinnati, OH, USA) mixed to resemble the colour of iodine with 0.01% iodine and 0.02% potassium iodide to give an odour of iodine</p> <p>Masks</p>	<p>Immersion of each finger and thumb of both hands to proximal interphalangeal joint (interphalangeal joint of thumb) into designated preparation for 5 seconds then air-dried for 5 to 6 min</p> <p>Exposure of recipients to donors either immediately after treatment or after 2-hour delay by hand contact with donor stroking fingers for 10 s</p> <p>Masks worn by donors and recipients during procedure.</p> <p>Recipients placed in single isolation rooms after second exposure till end of experiment.</p>	<p>Researcher-to-</p>	<p>Face-to-face and individually</p>	<p>US university</p>	<p>Exposure to donors on 3 consecutive days (days 2, 3, and 4) after initial exposure</p>	<p>Not described</p>	<p>Not described</p>	<p>Reported knowledge of hand preparation use as active, placebo, or don't know</p>	<p>Active (n = 24):</p> <p>6 active</p> <p>2 placebo</p> <p>16 don't know</p> <p>Placebo (n = 22):</p> <p>6 active</p> <p>7 placebo</p> <p>9 don't know</p>												
Hubner 2010	<p>Alcoholic hand disin-</p>	<p>Employees (admin-</p>	<p>Reduce absenteeism and spread of</p>	<p>2 alcohol-based hand rubs (500 mL bottles) for desktop use to ensure minimal effort for use:</p>	<p>Provision of hand rub and instruction on use as needed at work only and in ac-</p>	<p>Pre-</p>	<p>sum-</p>	<p>ably</p>	<p>pro-</p>	<p>vid-</p>	<p>In-</p>	<p>per-</p>	<p>son</p>	<p>to</p>	<p>staff</p>	<p>Ad-</p>	<p>min-</p>	<p>istra-</p>	<p>tion</p>	<p>of-</p>	<p>12 months overall</p> <p>Hand rub used as</p>	<p>Hand rub use especially</p>	<p>Not described</p>	<p>Self-reported compliance with hand hy-</p>	<p>Reported mean hand disinfection frequency</p>

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	fection	istrative officers)	infection in administration employees with frequent customer contact and work with paper documents through improved hand hygiene	1. Amphisep E (Bode Chemie, Hamburg, Germany) ethanol (80% w/w) based formula with antibacterial, antifungal, and limited virus inactivating activity. 2. For participants with skin problems: Sterillium (Bode Chemie, Hamburg, Germany) 2-propanol (45% w/w), 1-propanol (30% w/w), and mecetronium etilsulfate (0.2% w/w), with a refatting effect and has activity against bacteria, fungi and enveloped viruses. Hand cream: Baktolan balm, water-in-oil emulsion with no non-antibacterial properties (Bode Chemie, Hamburg, Germany)	cordance with prevailing standard[7]: at least 5 times per day, especially after toileting, blowing nose, before eating, and after contact with ill colleagues, customers, and archive material	ed or arranged by study team	ices in Germany Hand rubs used at desk or work (not outside of work).	much needed for complete wetting of the hands (at least 3 mL or a palmful)[8] at least 5 times per day.	after toiletting, blowing nose, before eating, and after contact with ill colleagues, customers, and archive material	giene measures			times per day: > 5: 19% 3 to 5: 59.8% 1 to 2: 20.5% < 1: 0.7%
Ladegaard 1999 (translated from Danish)	Hand hygiene programme	Day-care centres and their staff, children, and parents of children	Reduce risk of infection in child care through increased hygienic education of day-care professionals, motivation of daycare facilities for regular	Personnel guide on recommendations for: hygiene, ventilation, out-of-stay care, stricter hygienic regulations in cases with selected diseases Fairy tale and poster "The Princess Who Won't Wash Hands" Colouring in drawings "Wash hands" song and rhymes	Staff meeting in each DCC and training in microbiological cause of infection spread guided by National Board of Health and Hygiene Education of children in hand-washing (about bacteria and why and when to wash hands)	Research team presumably provided training.	Face-to-face training and activities by group with staff and children	On-site in DCCs	2-month intervention period 1-hour training of children	None described.	None described.	None described.	None reported.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

			hand hygiene, and informing parents about hand hygiene	T-shirt for children with the inscription “Clean hands - yes thank you” Diploma for children and book “The Princess Who Won't Wash Hands” to also be used by parents with their child Informational leaflet for parents in envelope	Practical hand-washing classes with 4 to 5 children at a time Provision of t-shirt, book, and diploma to children Provision of leaflet for parents					Information sent home to parents via children.			
Little 2015	Web-based hand-washing intervention	Householders (over 18) who were general practice patients	Prevent transmission of respiratory tract infections through improved hand hygiene to reduce spread via close contact (via droplets) and hand-to-face contact	Website-based programme: provided information about the importance of influenza and role of hand-washing; developed a plan to maximise intention formation for hand-washing; reinforced helpful attitudes and norms; addressed negative beliefs (URL provided for demonstration version no longer active; see www.lifeguideonline.org)	Provision of link to website for direct log in Automated emails prompted participants to use sessions and complete monthly questionnaires and maintain hand-washing.	Researcher-delivered web-based programme and emails.	Online individual	Households in England	4 months overall 4 weekly web-based sessions Monthly email questions to maintain hand-washing over 4 months	Tailored feedback provided within web programme	None described.	Emailed questions monthly to maintain hand-washing	None reported.
Luby 2005	Hand-washing promotion neighbourhood	Neighbourhoods and their households	Improve hand-washing and bathing with soap in settings where communitable dis-	Slide shows, videotapes, and pamphlets illustrating health problems from contaminated hands and specific hand-washing instructions Soaps: 90-gram white bars without brand	Hand-washing promotion to neighbourhoods: Neighbourhood meetings of 10 to 15 householders (mothers) from nearby homes	Research team collaboration with Health	Face-to-face in small groups and individually	Neighbourhoods and homes in Karachi, Pakistan	1-year weekly household visits 30- to 45-minute neighbourhood meet-	Soap replaced regularly.	None described.	None described, though soap use measured.	Households' mean use of study soap per week: 3.3 bars Average use per

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	level with 2 interventions at household level		eases are leading causes of childhood morbidity and mortality	names or symbols, same smell with identical generic white wrappers with serial numbers matched to households	and monthly meetings for men	Soap to households	Fieldworker home visits: discussed importance of and correct hand-washing (wet hands, lather them completely with soap, rub them together for 45 seconds, and rinse off completely) technique and promote regular hand-washing habits ^[11]	Orientation: HOPE ^[12]	Fieldworkers were trained in interviewing and hand-washing promotion.	ings 2 to 3 times/week first 2 months then weekly for months 2 to 9, then monthly			resident per day: 4.4 g
	A. Antibacterial soap			A. Households: 2 to 4 white bars of 90-gram antibacterial soap containing 1.2% triclocarban (Safeguard Bar Soap: Procter & Gamble Company (Cincinnati, OH, USA))						Monthly men's meetings first 3 months			
	B. Plain soap			B. Households: plain soap (no triclocarban)		Soap packets				Weekly household visits			
Milar 2016 and Ellis 2010	Skin and soft-tissue infection prevention intervention in addition to	Military trainees	Improve personal hygiene practices to prevent infection, especially acute respiratory infection in military trainees who are at increased risk	A. Enhanced standard: supplemental materials (a pocket card and posters in the barracks) B. CHG: CHG-based body wash (Hibiclens, Mölnlycke Heath Care, Norcross, GA, USA)	Provision of education and hygiene-based measures in addition to standard SSTI prevention brief upon entry: Enhanced standard: supplemental materials CHG: as for enhanced standard	Not described, presumably the researcher	Face-to-face and individually for body wash and pocket card	US military training base	One-off education on entry to training CHG: use of wash 1 per week for entire training period (14 weeks)	None described.	None described.	None described.	None described.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

	SSTI brief on entry also pro- vid- ed to con- trol			group, plus a CHG-based body wash and instruc- tions for use						ed- uca- tion not de- scribed.		
	A. En- hanced stan- dard											
	B. Chlorhex- idine											
Mor- ton 2004	Healthy Ele- hands men- 2004 tary (alco- schools hol gel as and hand- their wash- child- ing dren ad- and junct staff)	Prevent infections in ele- mentary school-age children who are particu- larly vul- nerable through adjunct use of al- cohol gel and ed- ucation based on Health Be- lief Mod- el (HBM) (Kirscht 1974)	Alcohol gel and dis- pensers: AlcoSCRUB (60% eth- yl alcohol) supplied by Erie Scientific Company, Portsmouth, NH, USA “Healthy Hands Rules” protocol ^[13] (Figure 3 in paper) Healthy Hand Resource Manual for school nurse, available for parents Monthly newsletters to parents “Healthy Hands” refrig- erator magnet for fam- ilies (see Figure 2 in pa- per) Informational letter to local primary care	Healthy hands protocol in- troduced after “Germ unit” edu- cation in classes Daily reminders to children on public address system (in first week) then week- ly reminders Review of proto- col in each class- room after vaca- tion by school nurse 2 classroom vis- its from school nurse “Healthy Hands” magnet provid-	Gel pro- vid- ed by sup- pli- ers. Re- search team pro- vid- ed uca- tion- al as- pects. Class- room teach- ers re- spon- sible	Face- to- face train- ing in class- es and indi- vid- ual in- for- ma- tion giv- ing and mon- itor- ing	Ele- men- tary schools in USA Wall- mount- ed near door en- trance of each class- room at age- ap- pro- pri- ate height	46 days 0.5 mL dis- pensed per appli- cation. Use of “special soap” ac- cording to “Healthy Hands Proto- col” (Fig- ure 3 in paper)	Rein- force- ment was con- cerned gel was mak- ing her sick, so school nurse pro- vid- ed addi- tion- al class- room vis- it to al- lay con- cerns.	1 stu- dent was con- cerned gel was mak- ing her sick, so school nurse pro- vid- ed addi- tion- al class- room vis- it to al- lay con- cerns.	Usage of gel calcu- lated.	5 gel ap- plications per day 1 dis- penser lasted 1 month.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

				providers, paediatricians, family practitioners, and advanced practice nurses	“Germ Unit” curriculum and materials including Germ models and Glo Germ	ed to parents and guardians. “Hand Checks on Wednesdays” to identify adverse effects of gel	for encouraging use of gel and reinforcing protocol					each grade level.	
							School nurse assisted in monitoring and hand checks for adverse effects.						
Nicholson 2014	Hand-washing with soap	Households with 5-year-olds and their mothers	Targeted 5-year-old children and their mothers as change agents to reduce incidence of respiratory infections (and diarrhoeal disease) through	Initial supply of 5 bars of free soap (90-gram Lifebuoy bars) replenished on submission of empty wrappers. Environmental cue reminders (wall hangers, danglers) Rewards (e.g. stickers, coins, toy animals)	Provision of soap and social marketing programme (Sidibe 2009) (Lifebuoy branding) to educate, motivate, and reward children for HWWS at key times Weeks 1 to 17: hand-washing occasions, germ education, soap’s	Dedicated team of “promoters” delivered education and home	Face-to-face in groups Individually by mother to child	“Classrooms” held in community buildings Home visits of households	41 weeks Weekly “classrooms” after school and home visits HWWS encouraged 5 key occasions: after defecation, be-	Mothers were asked to provide and share hand-washing tips with other	Technical difficulties with “soap acceleration sensors” to measure	Registers for “classrooms” and home visits where 3-week gaps in attendance triggered supervisors to ask participants to resume	Soap consumption: IG vs CG: 235 g vs 45 g

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

<p>hand-washing using behaviour change principles (Claessen 2008), including social norms for child and mother (Perkins 2003), using fear of contamination and disgust (Curtis 2001), peer pressure (Sidibe 2003), morale boosting, and networking support</p>	<p>importance in germ removal</p> <p>Week 18 onward: encouragement of HWWS on 5 key occasions supported by environmental cues</p> <p>"Classrooms" for children</p> <p>Home visits for mothers</p> <p>Parents' evenings to boost morale, build networks, and run competition for compliance, assignment completion, and folder decoration</p> <p>Establishment of a "Good Mums" club for sharing HWWS tips</p> <p>Rewards provided by mothers.</p> <p>Children encouraged to advocate HWWS within families before meals.</p> <p>Establishment of social norms for child and mother with pledges in front of peers</p>	<p>visits.</p> <p>Mothers provided supplied rewards.</p>	<p>in Mumbai, India</p>	<p>fore each of 3 meals, and during bathing.</p> <p>Week 18 onward: hand-washing on 5 occasions for 10 consecutive days</p> <p>6 weekly parents' meetings</p>	<p>mothers, competitions held for mothers.</p>	<p>HWWS behaviours prevented successful use.</p>	<p>or be withdrawn</p> <p>Monitoring of soap resale on open market by use of unique identifiers on soap wrappers and twice weekly checks in local shops</p> <p>Collection of used soap wrappers as soap consumption measure</p>
--	--	--	-------------------------	---	--	--	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Pan- de- jpong 2019	3 ac- tive inter- ven- tions (no con- trol) dif- fer- ent time- inter- val ap- plica- tions of al- cohol hand gel A. Every 60 min B. Every 120 min C. Once be- fore lunch	Preschool class- es (stu- dents and teach- ers) and their par- ents	Targeted preschool children who can have high infection rates in ILI; have close interac- tion so at risk of airborne, droplet, and con- tact trans- mission; and are of in- creasing- ly younger ages through hand gel as a single strategy of conve- nient and effective disinfec- tion	1 container of alcohol hand gel per classroom (active ingredients: ethyl alcohol, 70%; chlorhex- idine gluconate, 1%; Iri- gasan (triclosan), 0.3%) Cost of hand gel every 60 minutes was USD 6.39 per child per 12-week period Leaflet describing risk factors for ILI for each family	Teachers in- structed to: assist each child with dispensing hand gel at re- quired time interval, store hand gel properly, and re- fill gel as needed. Monitoring of hand gel use at specified times	Teach- ers su- per- vised, stored, and re- filled hand gel. In- struc- tions to teach- ers pre- sum- ably pro- vided by re- searchers. Leaflets dis- trib- uted through school. Mon- itor- ing of use by 2 re- search assis- tants	Face- to- face to schools, teach- ers and child- ren Indi- vid- ual assis- tance to child- ren with hand gel Leaflets given to each fami- ly.	Kinder- garten school in Bangkok, Thai- land	12 weeks overall 1 pump of gel per child per disinfec- tion round at 1 of 3 time in- tervals of school day: A. every 60 min B. every 120 min C. once only be- fore lunch, the school standard for hand hygiene	None de- scribed.	Stu- dents whose fam- ilies de- clined to partici- pate were not asked to use alco- hol hand gel. These stu- dents re- mained in their class- rooms and con- tin- ued to fol- low the school stan- dard for hand	2 research assistants monitored hand gel use every 60 or 120 minutes for the du- ration of study. Classroom teachers were re- quired to co-sign af- ter each disinfec- tion round.	Reported that com- pliance was en- sured for each inter- vention group Cost of hand gel every 60 minutes was USD 6.39 per child per 12-week period.
---------------------------	--	--	--	---	--	--	--	--	--	-------------------------	--	--	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

										hy- giene.			
Priest 2014	Hand sanitiser pro- vi- sion (in addi- tion to hand hy- giene edu- uca- tion ses- sion also pro- vid- ed to con- trol group)	Pri- mary schools and their stu- dents, teach- ers, and ad- min- istra- tive staff	Reduce person-to- person commu- nity trans- mission of infectious disease by targeting improved and ad- ditional hand hy- giene of school children through supervised hand sani- tiser provi- sion as an alternative to improv- ing and maintain- ing bath- room facil- ities	“No touch” dispensers (> 60% ethanol) for each classroom that dispensed dose when hands were placed un- der an infrared sensor Supply of top-up sanitiser as needed	Dispensers in- stalled into each classroom. Teachers asked to ensure that the children used sanitiser at particular times and to over- see general use (McKenzie 2010). Weekly class- room visits to top-up of sanitiser and measure quantity used 30-minute in- class hand hy- giene education session provided (also to control group) plus in- struction in hand sanitiser use.	School liai- son re- search assis- tants topped- up sani- tiser. Teach- ers	In- stal- la- tion of dis- pensers to class- rooms Su- per- vi- sion of child- ren by teach- ers deliv- ered face- to- face indi- vid- ually and as a class.	City schools in New Zealand	20 weeks (2 school terms) Sanitiser to be used by students at least af- ter cough- ing/sneez- ing, blow- ing their nose, and as they leave for morning break and for lunch break. Approx- imately 0.45 mL of sanitiser dispensed per wash. Weekly top-up of sanitiser	Child- ren were able to use the sani- tiser at any time they wished as well as at key times (McKen- zie 2010). due to child- ren tast- ing it when eat- ing, af- fect- ing use.	Change of sani- tiser after week 10 to flavour- less type of the same % ethanol in 41 of 396 class- rooms (10%) (in 9 of 34 schools) due to child- ren tast- ing it when eat- ing, af- fect- ing use.	Weekly classroom visits by school li- aison re- search assis- tants who recorded quantity of sanitiser used Total amount of sanitiser per class- room was measured. Compli- ance de- fined as dispensing a volume equivalent to at least 45 mL per child of hand sani- tiser solu- tion over the trial period.	100% dis- pensing 45 mL per child Average hand sani- tiser dis- pensed/child for 34 schools: 94 mL Median classroom difference in sanitiser usage between first 10 weeks and second 10 weeks amongst classes that switched products was 220 mL.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Ram 2015	Soap and intensive hand-washing promotion	Household compounds and its holders (adults and children) that had a household with ILI	Reduce transmission of ILI and influenza by promoting hand-washing in households with ILI as other householders who are at highest risk of exposure due to crowded and poorly ventilated homes. Followed constructs of Social Cognitive Theory and the Health Belief Model (Glanz 2008) and behaviour change communication using social	Hand-washing station in central location of each compound using: large water container with a tap; plastic case for soap; bar of soap. Cue cards depicting critical times for hand-washing: after coughing or sneezing; after cleaning one's nose or child's nose, after defecation; after clearing a child who has defecated; before food preparation or serving; before eating.	Hand-washing station in each compound Didactic and interactive group-level education and skills training describing influenza symptoms, transmission, and prevention, promoting health and non-health benefits of hand-washing with soap and identification of barriers and proposed solutions to hand-washing with soap Daily surveillance including weighing of soap and replacing if ≥ 20 g and resupply of water in container if needed Posting of cue cards Asking householders to demonstrate hand-washing with soap technique	Intervention staff arranged provision of hand-washing station and presumably provided education. Intervention staff conducted daily surveillance and reinforcement visits.	All elements delivered face-to-face but at compound (facilities), group (education), and individual levels (reinforcement).	Household compounds in a rural area of Bangladesh consisting of several households with common courtyard, shared latrine, water source, and cooking facilities	Initiation of intervention within 18 hours of enrolment, then daily until 10 days following resolution of patient's symptoms Day 1 set up of hand-washing station	Daily surveillance included observation of individual hand-washing reinforcement and modelling as needed.	None described.	Daily surveillance of facilities and reinforcement and modelling of hand-washing behaviours including observed hand-washing Cue cards in common areas of courtyard Presence or absence of soap during each of first 10 days of surveillance from 180 household compounds Patterns and amount of soap use measured. ^[14]	Soap present for at least 7 days in all compounds and on all 10 days in 133 compounds (74%). Soap and water together were present 7 or more of first 10 days in 99% of compounds, with water and soap observed together on all 10 days in 99 compounds (55%) Soap consumption per capita: median: 2.3 g maximal: 5 g (on Day 7)
----------	---	---	---	---	---	--	---	--	---	---	-----------------	--	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

				marketing concepts									
Roberts 2000	Education about infection control measures, hand-washing, and aseptic nose wiping	Child-care centres and their staff and children	Reduce transmission of respiratory infections in childcare centres through improved infection control procedures	<p>GloGerm (GloGerm, Moab, UT, USA)</p> <p>Newsletters to staff</p> <p>Songs and rhymes on hand-washing</p> <p>Plastic bags (sandwich bags available at supermarkets) to cover hand for nose wiping</p>	<p>Staff training in good health (developed by Kendrick 1994) and practical exercise of hand-washing with GloGerm</p> <p>Fortnightly visits and newsletter to reinforce training and to communicate techniques</p> <p>Recommended hand-washing technique as per guidelines of the time^[15] and after toileting, before eating, after changing diaper (staff and child), and after wiping nose unless barrier used</p> <p>Teaching of technique to children and wash hands for infants</p>	<p>Training and reinforcement activities provided by 1 of the researchers</p> <p>Teachers delivered training to children based on their training.</p>	<p>Face-to-face in groups for training and classes individually as needed to children or staff</p>	<p>Child-care centres in Canberra, Australia</p>	<p>8 months overall</p> <p>3-hour training in evening or 1-hour lunch for new staff after study start</p> <p>Duration of hand-washing: "count to 10" to wash and "count to 10" to rinse</p>	<p>Training for new staff provided as needed.</p>	<p>None described.</p>	<p>6-weekly compliance measured by recorded observation of recommended practice for 3 hours in morning in each centre, graded by quantiles of frequency of recommended hand-washing by children.</p>	<p>Compliance was reported only in relation to analysis of outcomes.</p> <p>High compliance reported for nose wiping and child hand-washing.</p>
San-dora 2005	Healthy Families with an index child out-of-	Families	Reduce illness transmission in the home through multifac-torial campaign centred on	<p>Alcohol-based hand sanitiser: active ingredi-ent: 62% ethyl alcohol (PURELL Instant Hand Sanitizer; GOJO Industries, Inc, Akron, OH, USA)</p>	<p>Supply of hand sanitiser and hand hygiene materials</p> <p>Biweekly tele-phone calls</p>	<p>Study in-vesti-gator</p>	<p>Not stat-ed whether ma-teri-als mailed or</p>	<p>Homes in USA</p> <p>Sani-tiser use in home</p>	<p>5 months overall</p> <p>Biweekly education-al materi-als</p>	<p>None de-scribed.</p>	<p>None de-scribed.</p>	<p>Recorded amount of hand sani-tiser used (as report-ed by the primary caregiver)</p>	<p>Median frequency of report-ed times of hand sanitiser use: 5.2 per day</p>

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	home child-care	hand hygiene education and hand sanitiser	Hand hygiene educational materials at home (fact sheets, toys, games)	Biweekly educational materials		delivered in person			Sanitiser dispensed 1 mL each pump.			38% used > 2 ounces of hand sanitiser per fortnight = 4 to 5 uses per day
Savolainen-Kopra 2012	STOPFLUO- Enhanced hygiene interventions IR1. Soap and water wash IR2. Alcohol-based hand rub	Office workers of work units Prevent transmission of respiratory infections in workplaces through enhanced hand hygiene with behavioural recommendations to reduce transmission by droplets during coughing or sneezing	IR1: Liquid hand soap ("Erisan Nonsid" by Far-mos Inc., Turku, Finland) IR2: in addition: Alcohol-based hand rub, 80% ethanol ("LV" by Berner Inc., Helsinki, Finland) Bottles of hand hygiene product (free of charge) to be used at home and in the office (IR2). Written instructions on hygiene for further refer-ence	Toilets equipped with liquid hand soap (all groups) or alcohol-based hand rub (IR2). Guidance on other ways to limit transmission of infections, e.g. frequent hand-washing in of-fice and at home, coughing, sneez-ing into dispos-able handker-chief or sleeve, avoiding hand-shaking Visits to work clusters and monitoring of materials avail-ability Monthly electron-ic "information spot" about vi-ral diseases for motivation to maintain hygiene habits Adherence activi-ties	In col-labora-tion with occu-pation-al health clin-ics serv-icing the cor-pora-tion Specially trained re-search nurse pro-vided guid-ance and vis-ited work-er	In-person pro-vi-sion of soap or hand rub Guid-ance and writ-ten in-struc-tions given per-son-ally. Face-to-face vis-its by study nurse	Of-fice work units in cor-pora-tions in Helsin-ki, Fin-land	15 to 16 months overall Month-ly visits by nurse through-out	Nurs-es as-sist-ed with any prac-tical prob-lems with inter-ven-tion as they arose. New em-ploy-ees re-ceived guid-ance on hand hy-giene and habits.	None de-scribed. Adherence assessed by an elec-tronic self-report survey of trans-mis-sion-limit-ing habits 3 times (more de-tails in protocol). Use of soap (IR1) and alco-hol-based disinfec-tant (IR2) for personal use was recorded. Study nurse checked availabil-ity of soap and alco-hol rub.	Avoid-ing hand-shaking became more com-mon and re-mained high in both groups. Record-ed use for personal use small-er than pre-dicted use based on hand hygiene instruc-tions. Soap or disinfec-tant usage per partic-ipant: IR1: 6.1 IR2: 6.9	

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

													clusters through-out intervention period.
Stebbins 2011	“WHACK the Flu” (hand sanitiser and training in hand and respiratory hygiene)	Elementary schools and their students and home-room teachers	Targeted school-aged children as important sources of influenza transmission through improved cough etiquette and hand hygiene in schools including sanitiser as potential inexpensive non-pharmaceutical interventions	Hand sanitiser dispensers with 62% alcohol-based hand sanitiser from PURELL (GOJO Industries, Inc, Akron, OH, USA) automatically dispensing 1 dose	Delivery of grade-specific presentations on “WHACK the Flu” concepts and proper hand-washing technique and sanitiser use: (W)ash or sanitise your hands often; (H)ome is where you stay when you are sick; (A)void touching your eyes, nose and mouth; (C)over your coughs and sneezes; and (K)keep your distance from sick people (provided URL no longer active) Desired frequency of hand wash use taught to student (see When and how much)	Project staff provided education. Home room teachers reinforced message and monitored proper use of sanitiser.	Face-to-face at schools, presumably as a group in classes	Elementary schools (Pittsburgh, USA) Dispersers installed in each classroom and all major common areas.	Whole intervention over 1 influenza season One-off installation of hand sanitiser dispensers One-off 45-minute education presentation and one-off refresher training at onset of influenza season Goal of use of 1 dose (0.6 mL) of sanitiser 4 times per day ^[16]	Encouraged to wash hands or use additional doses of hand sanitiser, or both, as needed	None reported.	Monthly teacher surveys of observed NPI-related behaviour in their students before, during, and after influenza season Measurement of hand sanitiser use at 2-week intervals throughout the intervention period	Teacher surveys of observed classroom NPI behaviour indicated successful adoption and maintenance of behaviours throughout influenza season. Average sanitiser use: 2.4 times per day

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					Installation of hand sanitiser dispensers												
					Refresher training at each school												
					Reinforcement of message and monitoring of sanitiser												
Ta- laat 2011	In- ten- sive hand hy- giene cam- paign	Schools and their stu- dents, teach- ers, and par- ents	Reduce or prevent transmis- sion of in- fluenza viruses amongst children through intensive hand hy- giene in- tervention campaign	Soap supplied as needed. Grade-specific student booklets each including a set of 12 games and fun activities that promoted hand-washing Hand hygiene activities materials including: games (e.g. how to escape from the germs); puzzles; soap activities (e.g. soap drawing); song specially developed to promote hand hygiene Teachers' guidebook including detailed description of the students' activities and methods to encourage students to practice these activities. Posters with messages to wash hands with soap and water upon arriv-	Establishment of a hand hygiene team in each school Provision of hand hygiene activities: weekly exercises (e.g. games, aerobics, songs, experiments); school activities, (e.g. obligatory hand-washing under supervision, morning broadcast, parent meetings, students-parents information transfer); specific school initiatives: (e.g. competitions and awards, hand-washing committee, school trips to soap factory	Hand hy- giene team (3 teach- ers from so- cial stud- ies, arts, and sports and the school nurse) en- sured that all pre- de- signed activ- ities for the hand hy-	De- liv- ered face- to- face in groups and indi- vidually	Ele- men- tary schools (grades 1 to 3) in Cairo, Egypt In school envi- ron- ment and class- rooms Poster near sinks in class- rooms and on play- ground	12 weeks overall Week- ly hand hygiene campaign activities Week- ly visits by social workers Twice-daily oblig- atory su- pervised hand- washing required by stu- dents for about 45 seconds, followed by prop- er rinsing and dry- ing with a clean	Soap and hand- dry- ing ma- terial pro- vid- ed by school ad- min- istra- tion if child- ren did not bring their own as was the cus- tom or fam- ilies could	None de- scribed.	Observ- ation by social workers of hand hygiene activities, availabili- ty of soap and dry- ing mate- rial, and students' hand- washing during the day Schools created own ac- tivities to improve compli- ance.	About 93% of the stu- dents had soap and drying material available. All but 2 inter- vention schools "had a rig- orous sys- tem of en- suring that school- child- ren were wash- ing their hands at least twice daily".				

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

<p>ing at school, before and after meals, after using the bathroom, and after coughing or sneezing.</p>	<p>and water purification plant) More details in Table 1 of paper</p>	<p>giene campaign were implemented.</p>	<p>cloth towel.</p>	<p>not afford it.</p>
<p>Informational flyers for parents reinforcing the messages delivered at the schools.</p>	<p>Song played regularly.</p>	<p>6 independent social workers visited the schools.</p>		<p>Schools could create own motivating activities such as selecting a weekly hand hygiene champion, developing the-atre plays, and launching school contests for drawings and songs.</p>
	<p>Social worker weekly visits</p>			
	<p>Distribution of flyers to parents</p>			



Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Temime 2018	Multifaceted hand hygiene programme (including alcohol-based hand rub)	Nursing home staff, residents, visitors, and external providers, including outside care providers	Nursing homes and their staff, and visitors and external providers have an increased risk of person-to-person transmission of pathogens, and HH is a simple and cost-effective tool for infection control; however, compliance with HH is poor in nursing homes.	Dispensers and pocket-sized containers of hand rub solution Posters promoting hand hygiene Developed local HH guidelines eLearning module on infection control and HH training with online quizzes requiring sufficient performance	Facilitated access to hand rub solution Campaign to promote HH with posters and event organisation Formation of local work groups in each NH Development of local HH guidelines Staff education using eLearning Monitoring of quantity of hand rub solution used	Same nurse provided HH training for all NHs. Provision of hand rub by NH Local work group developed guideline. eLearning module and posters presumably developed by research team.	Provision of materials for face-to-face Education and quizzes via eLearning	Nursing homes in France	1 year overall One-off provision of hand rub One-off eLearning repeated if unsatisfactory performance.	If staff did not score sufficiently on online quiz, they were invited to repeat the eLearning.	None described.	Estimated mean amount of hand rub solution used per resident per day assessed as proxy for HH frequency, based on quantity of hand rub solution bought by NH (which was routinely monitored in all the NHs).	Hand rub solution used: baseline quantity of consumed hand rub solution was 4.5 mL per resident per day. Over the 1 year, mean quantity consumed was significantly higher in intervention NH (7.9 mL per resident per day) than control (5.7 per resident per day).
-------------	--	---	--	--	---	---	---	-------------------------	--	--	-----------------	--	--

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

Turner 2004a	3 active interventions (no control)	Healthy volunteers	Assess the residual virucidal activity of organic acids used in currently available over-the-counter skin products for the prevention of experimental rhinovirus colds	1.7 mL of hand products: A. 62% ethanol, 1% ammonium lauryl sulphate, and 1% Klucel) B. 3.5% salicylic acid, or vehicle containing C. 1% salicylic acid and 3.5% pyroglutamic acid	Disinfection of hands then application of test product then allowed to dry. 15 min later, fingertips of each hand contaminated with 155 TCID ₅₀ of rhinovirus type 39 in a volume of 100 µL. Hands air-dried for 10 min. Intentional attempted inoculation with virus by contact with fingers, conjunctiva, and nasal mucosa with fingers of right hand. Left hand eluted in 2 mL of virus-collecting broth.	Researcher	Face-to-face individually	Communities in Manitoba, Canada	1.7 mL of product applied. See What for timing	Not described	Not described	Not described	Not described
Turner 2004b	2 active interventions (no control)	Healthy volunteers	Assess the residual virucidal activity of organic acids used in currently available over-the-counter skin products for	Skin cleanser wipe containing: A. 4% pyroglutamic acid formulated with 0.1% benzalkonium chloride B. 62% ethanol	Application of product to hands with towelette then allowed to dry. 15 min later, fingertips of each hand contaminated with 106 TCID ₅₀	Researcher	Face-to-face individually	Communities in Manitoba, Canada	Dose not reported; see What for timing Additional group challenged 1 h after application; final group chal-	Not described	Not described	Not described	Not described

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

	product:	the prevention of experimental rhinovirus colds			of rhinovirus type 39 in a volume of 100 µL.					lenged 3 h after application (remained at study site and not allowed to use or wash hands between).			
	A. Pyrog-lutam acid				Intentional attempted inoculation with virus by contact with fingers, conjunctiva, and nasal mucosa with fingers of right hand.								
	B. Ethanol				Left hand eluted in 2 mL of virus-collecting broth.								
Turner 2012	Anti-viral hand lotion	Healthy adults	Reduce rhinovirus infection and illness through hand disinfection with ethanol and organic acid sanitiser	Lotion containing 62% ethanol, 2% citric acid, and 2% malic acid Daily diary	Provision of lotion and instructions for use Meetings with participants to check compliance	Staff of study site presumably supplied lotion. Study site staff met with participants.	Face-to-face and presumably individually, but not specified	Study site at university community in the USA	9 weeks Every 3 hours whilst awake and after hand-washing for 9 weeks	None reported.	None reported.	Self-reported daily diary of time of each product application Twice weekly for 5 weeks then weekly meetings with participants to reinforce compliance with treatment	“All subjects ... applied at least 90% of the expected amount of hand treatment” (p.1424)

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Yeung 2011	Multifaceted hand hygiene programme (including alcohol-based hand rub)	Long-term care facilities and their health-care workers	Promote use of alcohol-based hand rub by staff in LTCFs as an effective, timely, and low-irritant method of hand hygiene in a high-risk environment	Free supply of pocket-sized containers of alcohol-based antiseptic hand rub (either WHO formulation I (80% ethanol) or II (80% propanol) carried by each HCW (supplier: Vickmans Laboratories)	Replacement hand rub as required	Hand hygiene seminar content	Reminder materials (3 to 5 posters and specially designed ballpoint pens)	Provision of materials	Provision of hand hygiene seminars to HCWs covering: indications, proper method, and importance of antiseptic hand rubbing and washing according to WHO 2006a) guidelines	Provision of feedback session	Direct, unobtrusive observation of hand hygiene adherence	Training of observation staff	Study team delivered the materials, seminars, and observer training.	Administrative staff of LTCF provided replacement hand rub and communicated with HCWs.	6 registered nurses	Delivered face-to-face and individually for hand rub and pens; not described if education was individually or by group, but seminar implies as a group	LTCFs in Hong Kong	Posters posted in common areas.	Adherence observations occurred in common rooms and resident rooms but not bathing or toilet areas.	7 months overall	Initial 2-week intervention period, then 7 months of hand rub provision and reminders	3 identical seminars at start of intervention; each staff member to attend once	Feedback session 3 months after start of intervention	2-hour training of observers	Adherence observations either 9 am to 12 pm or 3 pm to 6 pm, 1 LTCF at a time	Replacement of hand rub as required	As adherence dropped off in the middle months, the feedback session was delivered.	Direct observation of HCW adherence to hand washing and antiseptic hand rubbing (recorded separately and anonymously) during bedside procedures or physical contact with residents	3300 hand hygiene opportunities during 248.5 hours of observation on 92 days	90% attendance of seminars	Hand rubbing with gel increased significantly from 1.5% to 15.9%.	Hand-washing decreased significantly from 24.3% to 17.4%.	Control: 30%	Overall hand-washing adherence increased from 25.8% to 33.3%.
------------	--	---	---	--	----------------------------------	------------------------------	---	------------------------	---	-------------------------------	---	-------------------------------	--	--	---------------------	--	--------------------	---------------------------------	---	------------------	---	---	---	------------------------------	---	-------------------------------------	--	--	--	----------------------------	---	---	--------------	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

																				conducted direct observation of adherence after 2-hour training (100% interrater reliability).
Zomer 2015	Hand hygiene products and training	Day-care centres and their caregivers (staff)	Reduce infections in children attending DCCs through improved access to HH materials (Zomer 2013a) and compliance of their DCC caregivers to hand hygiene guidelines based on	HH products: dispensers for paper towels, soap, alcohol-based hand sanitizer, and hand cream, with refills for 6 months Reminder posters and stickers for children and DCC caregivers Training materials including booklet	Provision of free HH products sponsored by SCA Hygiene Products, Sweden. Provision of posters and stickers for children and staff Provision of training about RIVM 2011 for mandatory HH ^[18] Distribution of training booklet	Study team arranged supply of HH products and presumably provided training.	Products provided to DCCs in person for staff use. Mode of training not specified.	DCCs in regions of the Netherlands	6 months overall Initial one-off supply of products 3 training sessions with 1-month interval 2 team training sessions	Replacement of hand hygiene provided as required.	None described.	6-month follow-up observation of whether intervention dispensers and posters/stickers in use Survey of DCC caregivers HH guidelines compliance observed	2 DCCs did not use any HH products. Sanitiser products used in at least 1 of 2 groups in 94%, 89%, 86%, and 45% of intervention DCCs. Posters used in 86%, stickers in 74%.							

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	socio-cognitive and environmental determinants of caregivers' HH behaviour ^[17] (Zomer 2013b)	Team training sessions aimed at goal-setting and formulating HH improvement activities (Erasmus 2011; Huis 2013)										at 1, 3, and 6 months' follow-up:	DCC survey results:
												no. of HH actions/no. of opportunities	79% attended at least 1 training session; 77% received HH guidelines booklet.
													HH compliance at 6 months:
													IG: 59% vs CG: 44% (Zomer TP, et al, unpublished data)
													All intervention DCCs received guidelines training; all but 2 received at least 1 team training.

Hang hygiene and masks

Ae-lami 2015	Hygienic education and	Religious pilgrims	Prevent influenza-like illness by reduced infection	Hygiene package of: alcohol-based hand rub (gel or spray) surgical masks	Not clearly described, but it appears that packages may have been distributed by trained physi-	Not specifically described	Not described, but it appears	Not described, if before de-	One-off during Hajj season	Not described	Not described	Not described	None described
--------------	------------------------	--------------------	---	--	---	----------------------------	-------------------------------	------------------------------	----------------------------	---------------	---------------	---------------	----------------

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	pack- age	trans- mission through personal hygiene measures	soap paper handkerchiefs user instructions	cians before de- parture to or on site of country of pilgrimage	that par- ture (from Iran) or on site (in Sau- di Ara- bia)	dis- trib- uted (in face- to- face and indi- vidually							
Aiello 2010	2 ac- tive inter- ven- tions: A. Face mask (FM) B. Face mask and hand hy- giene (FM + HH)	Stu- dents living in uni- ver- sity resi- dences	Reduce the inci- dence of and mit- igate ILL by use of non-phar- maceuti- cal inter- ventions of person- al protec- tion mea- sures	7 face masks (stan- dard medical proce- dure masks with ear loops TECNOL proce- dure masks; Kimber- ly-Clark) 7 resealable plastic bags for mask storage when not in use (e.g. eating) and for disposal Alcohol-based hand sanitiser (62% ethyl alcohol in a gel base, portable 2- ounce squeeze bottle, 8- ounce pump) Hand hygiene education (proper hand hygiene practices and cough etiquette) via emailed video, study website, written materials detail- ing appropriate hand sanitiser and mask use	Weekly supply of masks through student mailbox- es Provision of basic hand hygiene edu- cation through an email video link, the study website, and written materi- als; instruction to wear mask as much as possi- ble; education in correct mask use, change of masks daily, use of pro- vided resealable bags for mask storage and dis- posal Provision of re- placement sup- plies which stu- dents signed for upon receipt	Not de- scribed, ex- cept ed- uca- tion web- site; pro- vided via study web- site (URL not pro- vid- ed) “Trained staff” for com- pli- ance mon- itor- ing	Ed- uca- tion via email and study web- site; pro- vi- sion of masks and sani- tiser in per- son resi- dences	Uni- ver- sity resi- dence halls in the USA	One-off education, 6 weeks (exclud- ing spring break) of face mask and/ or hand hygiene measures which com- menced at “the be- ginning of the in- fluenza season just after identifica- tion of the first case of influen- za on cam- pus” (p.496).	Mask wear- ing dur- ing sleep op- tion- al and cour- aged out- side of resi- dence.	Uni- ver- sity spring break oc- curred dur- ing weeks 4 and 5 of the study, with most stu- dents leav- ing cam- pus and trav- el- ling; they were not	Week- ly web- based stu- dent sur- vey includ- ed: self- reported average number of times hands washed/ day and average duration of hand- washing to obtain compos- ite “opti- mal hand- washing” score (at least 20 s ≥ 5/day); aver- age no. of mask hours/	Average mask use hours/day: FM + HH 2.99 vs FM 3.92 Average hand- washing times/day: FM + HH 6.11 vs FM 8.18 vs control group 8.75 Daily wash- ing sec- onds/day: FM + HH 20.65 vs FM 23.15 vs control 22.35

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

						Study-affiliated residence hall staff provided replacement supplies.			vided as needed.		re-quired to continue protective measures at that time.	day/week; average hand sanitiser use/day/week and amount used.	Hand sanitiser use times/day: FM + HH: 5.2 vs FM 2.31 vs control 2.02 No. of proper mask wearing participants/hour of observation: FM + HH 2.26 vs FM 1.94
Aiello 2012	2 interventions: A. Face mask (FM) B. Face mask and hand sanitiser (FM + HH)	Students living in university residences	Prevent ILI and laboratory-confirmed influenza by use of non-pharmaceutical interventions of personal protection measures (e.g. face masks and hand hygiene)	Packets of 7 standard medical procedure masks with ear loops (TECNOL procedure masks, Kimberly-Clark, Roswell, GA, USA) and plastic bags for storage during interruptions in mask use (e.g. whilst eating, sleeping) and for daily disposal Hand sanitiser (2-ounce squeeze bottle, 8-ounce pump bottle with 62% ethyl alcohol in a gel base)	Intervention materials and educational video provided. Supply of masks and instructions on wearing Provision of replacement masks or sanitisers as needed on site	Trained study staff available at tables in each residence hall for surplus masks and sanitiser	Hygiene packs delivered to student mailboxes; face-to-face supply also available	University residence halls in the USA	One-off educational video at start Weekly supply of hygiene packs Masks to be worn at least 6 hours/day Study staff available onsite with re-	Students encouraged but not obliged to wear masks outside of residence hall.	1-week university spring break during the study when majority of students left	Weekly student survey including compliance (e.g. masks hours/day, frequency and amount of sanitiser use, number of hand washes/day, duration of hand-	Self-reported mask wearing: no significant difference Sanitiser use: significantly more in FM + HH than FM or control groups

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

				Replacement face masks and hand sanitiser		and for observing compliance			placement supplies as needed for duration of intervention (6 weeks, excluding spring break)	campus	washing (seconds)	More results in S1 of paper.	
				Educational video: proper hand hygiene and use of standard medical procedure face masks							Observed compliance completed by trained study staff who daily and anonymously observed mask wearing in public areas of residences.	Staff observed an average of 0.0007 participants properly wearing a mask for each hour of observation.	
Cowling 2009	2 active interventions in addition to control of lifestyle education:	Householders with index patient with influenza	Reduce transmission of influenza in households through personal protective measures	A. and B. Liquid soap for each kitchen and bathroom: 221 mL Ivory liquid hand soap (Proctor & Gamble, Cincinnati, OH, USA) Alcohol hand rub in individual small bottles (100 mL) WHO recommended formulation I, 80% ethanol, 1.45% glycerol, and 0.125% hydrogen peroxide (Vickmans Laboratories, Hong Kong, China)	Home visits Provision of soap, hand rub, and masks as applicable and when to use them HH: education about efficacy of hand hygiene Demonstration of proper hand-washing and antiseptics techniques + FM: education about efficacy of surgical face masks in reducing disease spread to household contacts if	Trained study nurse provided interventions.	Face-to-face to householders	Households in Hong Kong	Initial home visit scheduled within 2 days (ideally 12 h) of index case identification. Further home visits day 3 and 6, 7-day follow-up HH: use of liquid soap after every wash-room visit, sneezing or	Not described	Not described	Monitoring of adherence during home visits Evaluation of adherence on final visit by interview or self-reported practices and counting of amount of soap and rub left in bottles and remaining masks for FM group	Most initial visits completed within 12 h. Intervention groups “reported higher adherence ... than the control group. Self-reported data were consistent with measurements of amount of soap,

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR)

checklist		(Continued)													
B.	Face masks and enhanced hygiene (FM + HH)	to 7: box of 75 paediatric masks	all parties wear masks	Demonstration of proper wearing and hygienic disposal	All groups: provision of education about the importance of a healthy diet and lifestyle, both in terms of illness prevention (for household contacts) and symptom alleviation (for the index case)	coughing, when their hands were soiled. Use rub when first returning home and immediately after touching any potentially contaminated surfaces	alcohol hand rub, and face masks used" (p.443) (see Table 6 in paper).	"Adherence to the hand hygiene intervention was	slightly higher in the hand hygiene group than the face mask	plus hand hygiene group."	Median masks used:	Index: 9	Contact: 4	More details in paper and Appendices	
Larson 2010	2 active inter-	Hispanic house-	Reduce incidence and sec-	A. and B.	Provision of materials and instructions for	4	Face-to-face	Households in	19-month follow-up	Change masks be-	None de-scribed.	RA home visits for adherence	Sanitiser use (mean		

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

<p>ven- tions in ad- di- tion to con- trol of URI ed- uca- tion:</p> <p>A. Al- co- hol-based hand sani- tiser (HS)</p> <p>B. Face masks and hand sani- tiser (FM + HS)</p>	<p>hold- ers at least 1 preschool or ele- men- tary school child</p>	<p>ondary transmis- ion of URIs and influenza through school-phar- maceuti- cal house- hold level interven- tions</p>	<p>2-month supply of hand sanitiser in 8-, 4-, and 1- ounce containers:</p> <p>PURELL (Johnson & Johnson, Morris Plains, NJ, USA)</p> <p>B. 2-month supply of masks:</p> <p>Procedure</p> <p>Face Masks for adults and children (Kimber- ly-Clark, Roswell, GA, USA)</p> <p>Replacement supplies at least once every 2 months</p> <p>Disposable thermome- ters</p> <p>Educational materials about URI prevention, treatment, and vaccina- tion (written in Spanish or English language)</p>	<p>when to use in- cluding demon- stration of use and observa- tion of return demonstration by householder</p> <p>A. Mask worn when household- er had: “temper- ature of $\geq 37.8^{\circ}\text{C}$ and cough and/ or sore throat in the absence of a known cause oth- er than influen- za” (CDC defin- ition of influen- za-like illness at the time).</p> <p>Home visits to reinforce adher- ence, replen- ish supplies and record use, an- swer questions</p> <p>B. Telephone calls to reinforce mask use</p> <p>All groups re- ceived URI educa- tional materials.</p>	<p>gual re- search assis- tants (RAs) with mini- mum bac- calau- reate de- gree and ex- peri- ence in com- mu- ni- ty-based re- search; pro- ce- dures were prac- tised with each other until demon- strat- ed profi- cien- cy</p>	<p>to house- hold- ers</p>	<p>New York, USA</p>	<p>Initial home vis- it, then at least every 2 months</p> <p>Sanitis- er for use at home, work, and school</p> <p>B. Tele- phone calls days 1, 3, 6</p> <p>Masks worn for 7 days when within 3 feet of person with ILL or no symp- toms.</p>	<p>tween inter- ac- tions with per- son with ILL</p> <p>House- hold- ers' ques- tions and mis- con- cep- tions ad- dressed on home vis- its.</p>	<p>with ran- dom ac- compani- ment by project manager, who also made ran- dom calls to house- holders</p> <p>Telephone calls to reinforce mask use</p> <p>Used bot- tles or face masks, or both, monitored for usage.</p>	<p>ounces/ month)</p> <p>HH: 12.1</p> <p>FM + HH: 11.6</p> <p>Mask compli- ance was “poor”: 22/44 (50%) used within 48 hours of onset.</p> <p>Mask users re- ported mean mask use of 2.</p>
--	--	---	---	--	--	--	------------------------------	--	--	---	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Simmerman 2011	2 active interventions: A. Hand-washing education and hand-washing kit (HW) B. Hand-washing education, hand-washing kit, and face masks (HW + FM)	Households with a febrile, influenza-positive child	Decrease influenza transmission in household with a febrile influenza-positive child through promoted use of hand-washing or hand-washing with face mask use	A. and B. Hand-washing kit per household including graduated dispenser with standard unscented liquid hand soap (Teepol brand. Active ingredients: linear alkyl benzene sulfonate, potassium salt, and sodium lauryl ether sulphate) Replacement soap as needed Written materials from education including pamphlets and posters attached near sinks in household. B. Box of 50 standard paper surgical face masks and 20 paediatric face masks (Med-con company, Thailand #14IN-20AMB-30IN)	A. and B. Provision of intensive hand-washing education on initial home visit to household members with 5 approaches: discussion, individual hand-washing training, self-monitoring diary, provision of soap, and provision of written materials (Kaewchana 2012) Individual hand-washing training ("why to wash", "when to wash", and "how to wash" in 7 hand-washing steps described in Thailand Ministry of Public Health guidelines) B. Provision of education of benefits of and appropriate face mask wearing Soap replaced as needed. More details (Kaewchana 2012)	Study nurse conducted home visits, provided education and monitoring activities.	Education provided to household member and individually for hand-washing training.	In homes (in Bangkok at initial home visit)	One-off provision of kits at initial home visit conducted within 24 hours of enrolment Subsequent home visits on days 3, 7, and 21 90-day supply of hand-washing supplies 30-minute education provided at initial home visit	B. No face masks whilst eating or sleeping as impractical and could hinder breathing in ill child Impromptu education and training provided by nurses as questions arose.	None described.	Self-monitoring diary recording hand-washing frequency > 20 s and face mask use for that group Reinforcement of messages by nurses on subsequent home visits Amount of household liquid soap and number of face masks used	Reported average hand-washing episodes/day: HW: 4.7 HW + FM: 4.9 Parents had highest frequency (5.7), others (4.8), siblings (4.3), index cases (4.1). Average soap used/week: HW: 54 mL/person HW + FM: 58.1 mL/person B. Mask use: 12/person/week Mask wearing median min-
----------------	---	---	--	--	---	--	--	---	---	--	-----------------	--	---

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

													utes/day: 211
													Parents 153,
													other rela- tions
													59, index patients 35, sib- lings 17
Suess 2012	2 ac- tive in- terven- tions in ad- dition to written infor- mation: A. Mask/ hygiene (MH) B. Mask (M)	House- holds with influen- za-posi- tive index case in the absence of further respiro- tory ill- ness with- in the pre- ceding 14 days	Prevent influenza transmis- ion in house- holds through easily ap- plicable and acces- sible non- pharma- ceutical inter- ven- tions such as face masks or hand hygiene measures	A. Alcohol-based hand rub (Sterilium, Bode Chemie, Germany) A. and B. Surgical face masks in 2 different sizes: children < 14 years (Child’s Face Mask, Kimberly-Clark, USA) and adults (Aérokyn Masques, LCH Medical Products, France) Written information pro- vided on correct use of intervention and on infection prevention (Seuss 2011) (Tips and information on the new flu A/H1N1) (URL provided is no longer active) Digital tympanic ther- mometer General written informa- tion on infection preven- tion	A. Provision of hand rub and masks A. and B. Provi- sion of masks on- ly Provision of ther- mometer and how to use it Mask fit assessed (at first house- hold visit) Information pro- vided by tele- phone and writ- ten instructions at home visit on proper use of in- terventions and recommendations to sleep in a different room than the index patient, not to take meals with the index patient, etc. (Seuss 2011)	Study per- son- nel of arranged pro- vi- sion of ma- teri- als, rang the par- tici- pants, vis- ited the homes, demon- strated and as- sessed fit of masks.	Pro- vi- sion of ma- teri- als in per- son to house- holds Initial tele- phone deliv- ery of infor- mation Face- to- face home visits	House- holds in Berlin, Ger- many	Over 2 consec- utive flu seasons Day 1 house- holds re- ceived all necessary material in- struc- tions. Household visits no later than 2 days af- ter symp- tom onset of the in- dex case, then days 2, 3, 4, 6, 8 (5 times) or on days 3, 4, 6, 8 (4 times) de- pending on the day	Adult masks worn if masks for un- der 14- year- olds did not fit prop- erly. If other house- hold mem- bers de- veloped fever (> 38.0 °C),	In the season 2010/11 partici- pants also re- corded num- ber of masks used per day. Partici- pants of the MH house- holds ad- ditionally noted the number of hand dis- infections per day. Exit ques- tionnaire about	Self-re- ported daily ad- herence with face masks, i.e. if they wore masks “al- ways”, “mostly”, “some- times”, or “never” as instructed. Partici- pants of the MH house- holds ad- ditionally noted the number of hand dis- infections per day. Exit ques- tionnaire about	Face mask use (me- dian/indi- vidual): MH: 12.6 M: 12.9 Daily ad- herence was good, reaching a plateau of over 50% in nearly all groups from the third day on. MH hand rub use (median): 87 mL (Seuss 2011) MH mean frequen- cy of daily

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

<p>In-person demonstration of interventions at first home visit</p>	<p>of recruitment or</p>	<p>cough, or sore throat,</p>	<p>(preventive) behaviour during the past 8 days, general attitudes towards NPI, the actual amount of used intervention materials, and, if applicable, problems with wearing</p>	<p>hand disinfection: 7.6 (SD 6.4) times per day</p>
<p>All participating households received general written information on infection prevention.</p>	<p>Hand rub use: after direct contact</p>	<p>they were asked to</p>	<p>adopt the same preventive behaviour as the index patient.</p>	<p>See paper and Suess 2011 for more results.</p>
	<p>with the index patient (or other symptomatic household members), after at-risk activities or contact^[19]</p>	<p>to adopt the same preventive behaviour as the index patient.</p>	<p>face masks.</p>	
	<p>Mask use: at all times when index patient and/or any other household member with respiratory symptoms were together in 1 room</p>		<p>Used intervention material per household member was calculated by dividing the amount used per household by the number of household members.</p>	
	<p>Regular change of face masks, not worn during the night or</p>		<p>See paper and Suess 2011 for</p>	

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

													outside the household	more details.
Hand hygiene and surface/object disinfection														
Ban 2015	Hand hygiene and surface cleaning or disinfection	Kindergartens and the families of their students	Reduce transmission of infection in young children from contaminated surfaces or hands through hand hygiene and surface cleaning or disinfection	Antibacterial products for hand hygiene and surface cleaning or disinfection: liquid antimicrobial soap for hand-washing (0.2% to 0.3% parachlorometaxilenol). Instant hand sanitiser for hand disinfecting (72% to 75% ethanol), antiseptic germicide (4.5% to 5.5% parachlorometaxilenol, diluting before use). Bleach (4.5% to 5.0% sodium hypochlorite, diluting before use) for surface disinfecting. Produced by Wheathfields Lohmann (Guangzhou) Company Ltd.	Provision of products to kindergartens and families Instruction of parents or guardians and teachers in hand hygiene techniques and use of antibacterial products Daily cleaning of kindergartens with products At least twice/week cleaning of homes and weekly cleaning or disinfecting of items such as children's toys, house furnishings, frequently touched objects (doorknobs, tables or desks), kitchen surfaces (utensils, cutlery, countertops, chopping boards, sinks, floors, etc.), bathroom surfaces (toilet, sink, floor, etc.)	Research team provided products and instructions and monitoring.	Materials provided to kindergartens and families in person to families and staff.	In kindergartens (hard surfaces) and families' homes (Xi-antao, China)	1 year overall Daily hand-washing with soap before eating, after using bathroom, nose blowing, and outdoor activities Hand sanitiser carried daily. Kindergarten cleaning daily Home cleaning at least twice/week	Families and teachers could contact study management at any time as needed. Exchange of empty bottles for new ones at any time	Not described	Close contact with teachers and families for monitoring, e.g. unscheduled parents' meetings, quarterly home visits, phone interviews, and monthly cell phone messages Monthly survey of consumption of products by volume, total usage, person usage	Consumption of products by person (mL/person/day). Liquid soap: 7.7 Sanitiser: 1.4 Bleach: 25.0 Antiseptic-germicide: 12.5	

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					Monitoring activities								
Cara-bin 1999	Hygiene programme	Day-care centres and their staff and children	Reduce infections in at-risk children (under 3 years old) in DCCs with inexpensive, easily implementable and practical interventions	Hygiene materials and documents, e.g. colouring books, hand-washing posters, hygiene videotapes Materials for training Reimbursement of equivalent of 1 full-time educator's salary Bleach (diluted 1:10) for toy and play area cleaning	Provision of comprehensive hygiene training session to entire DCC staff, especially the educators of participating classrooms Training in recommendations for hygiene practices: i. toy cleaning ii. hand-washing technique and schedule iii. use of creative reminder cues for hand-washing iv. open window for daily period v. sandbox and play area cleaning Payment of salary of educator for the day to encourage participation DCC meetings to discuss training session with all staff	Training appears to have been provided by study team.	Appears staff trained as a group, i.e. "entire DCC staff"	Day-care centres in Canada Location of training not described, except may have been off-site from DCCs since 1 DCC did not "send" staff to training.	15-month trial One-off 1-day training Toy cleaning at least every 2 days Hand-washing at least after DCC arrival, after outside play, after bathroom, before lunch Open windows at least 30 min/day Biweekly cleaning of sandbox/play area	Teachers to use creative reminder cues for hand-washing with children	Not described	Follow-up telephone questionnaire for DCC directors about following training recommendations	Use of materials: colouring book: 22/24 poster: 23/24 video-tapes: 18/24 staff meetings: 19/24 Increased frequency of toy cleaning: 6/24 Use of rake and shovel for sandpit: 17/24 Frequency of cleaning sandbox: 14/24

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

Kotch 1994	Hygiene	Caregivers at child day-care centres (CDCCs)	Develop feasible, multicomponent hygienic intervention to reduce infections in children at CDCCs who are at increased risk	Hygiene curriculum for caregivers Availability of soap, running water, and disposable towels Waterless disinfectant scrub (Cal Stat) used only if alternative was not washing at all. Handouts posted in CDCC.	Delivery of hygiene curriculum to caregivers through initial training session which required demonstration of participants' hand-washing and diapering skills Local procedures: Hand-washing of children and staff Disinfection of toilet and diapering areas Physical separation of diapering areas from food preparation and serving areas Hygienic diaper disposal Daily washing and disinfection of toys, sinks, kitchen and bathroom floors Daily laundering of blankets, sheets, dress-up clothes Hygienic preparation, serving, and clean up of food	Research team delivered training. Scrub donated by Calgon Vetal Laboratories.	Face-to-face training and follow-up group and individually	Classrooms of child day-care centres in the USA	8 months overall 3-hour initial training session Cleaning schedules as described in column What (procedures) Onsite follow-up training 1 week and 5 weeks later	Follow-up sessions addressed questions and local adaptations to procedures. As-required induction training	During intervention, research team encouraged directors to address physical barrier to hygiene practice, such as distance between sink and diapering areas and sink access in rooms.	Follow-up sessions reinforced training. Meeting with directors 5 weekly unobtrusive record-observation by training staff	Rate of compliance to barrier modification was better in younger centres, which were more likely to have written guidelines.
---------------	---------	--	--	---	---	--	--	---	--	---	--	--	--

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					Separate training of food handlers								
					As-required induction training for new staff								
					Onsite follow-up training reinforcing adaptations, demonstrations and discussion of hygiene techniques, responding to questions, and review of handouts								
					Monthly meeting with centre directors to encourage leadership and support								
Mc-Coneghy 2017	Multifaceted hand-washing and surface-cleaning intervention	Nursing homes and their staff	Reduce exposure to pathogens and person-person transmission in high-risk facility of close environment and potentially contaminated surfaces through multifaceted inter-	Education and launch materials Online module for certified nursing assistants about: infection prevention, product, and monitoring "Essential bundle" of hygiene products supplied at no cost: - hand sanitiser gel and foam - antiviral facial tissues - disinfecting spray - hand and face wipes Plus additional:	Pre-intervention: NH administrators required to: - identify a "Heroes In Prevention" champion and team - allow all staff participation in education - iPad use for staff in each floor or community - ask staff to incorporate intervention into workflow	Study personnel equipped staff with knowledge and tools and support. NH staff (e.g. champion, hy-	Face-to-face interaction with staff for planning and some aspects and delivery of products	Nursing homes in the USA On-site and at unit/team levels On-line training	6 months overall: training period: 3 months 1-hour launch event 1 or 2 hygiene monitors/site 1 champion/site 1-hour on-line module for selected	Sites could use existing comparable products from other vendor and fill in any gaps with study	2 sites re-trained due to low training participation rate.	Cloud-based audit and feedback system via secure login to web browsers on NHs' existing computers or via iPads included weekly product consumption to get measure:	Online training participation rates: > 90% for 3/5 sites, 13% and 23% for 2/5 Administrators demonstrated high fidelity in reporting measures of

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

vention equipping staff to protect residents from infection within the “culture” of care	- 4 skin cream and wipe products iPads for compliance audits Newsletters for support during intervention	Delivery of 3 components: - education - cleaning products - compliance audit and feedback Education: Launch event for all staff to publicise programme and explain roles Intensive training of "hygiene monitors" for data collection and compliance audit and feedback tool Training of site champion Training of select group of certified nursing assistants (online module) Audit and feedback activities Ongoing support during intervention: - newsletter with best practices - teleconferences with each NH	giene monitors, nursing assistants) delivered as-pects of interventions after specific training.	Some aspects delivered online (e.g. nursing modules, compliance auditing)	nursing assistants iPads for each community or floor Weekly teleconferences initially decreased in frequency over time. Weekly measurement of product consumption	products. New staff provided with education, as needed and came on-board. Re-training of sites with low training participation rates	weekly count of product units consumed x no. of hand hygiene occasions	hand-washing (> 80% of time). Hand-washing rates in Figure 1B in paper reported as “relatively constant” and “not ideal in the first few months”, but improved significantly over time.
--	--	--	--	---	---	--	--	--

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist *(Continued)*

					- "onboarding" education of new staff								
San-dora 2008	Multifactorial intervention, including alcohol-based hand sanitiser and surface disinfection	Elementary school and students	Reduce transmission of infections in school children through improved hand hygiene and environmental disinfection	1 container of disinfecting wipes (Clorox Disinfecting Wipes (The Clorox Company, Oakland, CA, USA); active ingredient, 0.29% quaternary ammonium chloride compound) Pre-labeled 1.7-ounce containers of alcohol-based hand sanitiser (AeroFirst non-aerosol alcohol-based foaming hand sanitiser (DEB SBS Inc, Stanley, NC, USA, for The Clorox Company); active ingredient, 70% ethyl alcohol) Receptacle in classrooms for empty containers	Sanitiser and wipes provided to classroom/teacher with instructions for use. Teachers disinfect desks once daily. Hand sanitiser to be used: before and after lunch, after use of the restroom (on return to the classroom; hand hygiene with soap and water occurred in the restroom, because sanitisers were not placed there), after any contact with potentially infectious secretions (e.g. after exposure to other ill children or shared toys that had been mouthed)	Research team arranged supply of materials and instructed teachers on use. Teachers instructed in use of materials and in collecting empty containers and distrib-	Products provided to schools. In-struction provided to teachers and children.	Elementary schools and classrooms in the USA	8-week period Desks disinfected once a day.	Products replenished as needed.	None described.	Individually labelled containers collected every 3 weeks from the classroom to assess adherence.	Product usage: average wiper used/week: 897 (128 wiper/classroom/week) Average bottles of hand sanitiser used per week: 8.75 (1.25 bottles/classroom/week)

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

													uting new prod- uct.
Quarantine													
Miya- ki 2011	Quar- an- tine from work (stay- at- home or- der)	Em- ploy- ees	Prevent spread of influenza in work- places by quaranti- ning work- ers who had a co- habitat- ing family member with an ILI	Full wages to employee	Non-compulso- ry asking of work- ers whose fami- ly members de- veloped an ILI to stay at home vol- untarily on full wages. Daily measuring of temperature before leaving work. Where symptoms were doubtful, in- dustrial physician made judgement. Company doctors provided input on cancelling of stay-at-home or- ders as required.	Health man- age- ment de- part- ment over- saw the pro- ce- dures and decis- ions.	Mode of ad- vice to em- ploy- ees not de- scribed.	Car in- dus- tries in Japan	Stay-at- home or- der for 5 days af- ter reso- lution of ILI symp- toms or 2 days after alleviation of fever over 7.5 months	Strict stan- dard for can- celling of stay- at- home or- ders de- scribed.	None de- scribed.	Recording of compli- ance with stay-at- home re- quest	100% compli- ance to stay at home re- ported.
Other (miscellaneous) interventions													
Farr 1988a 1	2 ac- tive inter- ven- tions in ad- di- tion to con- trol	Fam- ilies	Reduce transmis- ion of viruses from hand contam- ination via hand- to-hand contact or large-	3-ply tissues with: A. 5.1 mg/inch ² (2.54 cm ²) of the virucidal mixture (58.8% citric acid, 29.4% malic acid, 11.8% sodium lauryl sul- phate) B. 3 mg/inch ² (2.54 cm ²) of saccharin	Family visits to distribute tissues Weekly contact of mother Families instruct- ed to only use supplied tissues.	Nurse epi- demi- olo- gist vis- ited fami- lies.	Face- to- face vis- its to fami- lies and indi- vidu- als in	Com- mu- nities in the USA	6 months overall Monthly family vis- its Week- ly con- tact with mother	Not de- scribed	Not de- scribed	Family vis- its and week- ly con- tact with mother to encour- age com- pliance	Not de- scribed

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	of no tis- sues: A. Viru- cidal nasal tis- sues B. Place- bo tis- sues	particle aerosol through tissues for nose blow- ing and coughs and sneezes		applied uniformly to all 3 plies of the tissue Tissues prepared by Kimberly-Clark Corpora- tion, Neenah, WI, USA.			fam- ilies (es- pe- cially moth- ers)						
Farr 1988b 2	2 ac- tive inter- ven- tions (no con- trol): A. Viru- cidal nasal tis- sues B. Place- bo tis- sues	Fam- ilies	Reduce transmis- sion of viruses from hand contam- ination via hand- to-hand contact or large- particle aerosol through tissues for nose blow- ing and coughs and sneezes	2-ply tissues containing: A. 4.0 mg/inch ² (2.54 cm ²) of antiviral mixture (53.3% citric acid, 26.7% malic acid, 20% sodium lauryl sulphate) B. 3 mg/inch ² (2.54 cm ²) of succinic acid, malic acid, sodium hy- droxide, and polyethyl- ene glycol Tissues prepared by Kimberly-Clark Corpora- tion, Neenah, WI, USA.	Family visits to distribute tissues and encourage compliance Weekly contact of mother Families instruct- ed to only use supplied tissues.	Nurse epi- demi- olo- gist vis- ited fam- ilies month- ly. Study mon- itor vis- ited bi- month- ly.	Face- to- face vis- its to fam- ilies and indi- viduals in fam- ilies (es- pe- cially moth- ers)	Com- mu- nities in the USA	6 months overall Monthly family vis- its Week- ly con- tact with mother Bimonth- ly study monitor visit	None de- scribed.	None de- scribed.	Bimonth- ly study monitor visits to encour- age com- pliance as well as monthly and week- ly contact by nurse	In 124/222 families, 1 or more family members report- ed not us- ing the tis- sues regu- larly and/ or report- ed hav- ing side ef- fects from the tis- sues.
Longini 1988	2 ac- tive inter- ven- tions (no	House- holds and their fami- lies	Prevent intrafamil- ial trans- mission of viral agents in a	Treated tissues of 3-ply material identified with no specific identifiers (Kimberly-Clark Corpora- tion)	Tissues deliv- ered to house- holds with specif- ic instructions on use (all purpos- es, when blowing	Tis- sues as- signed by study	Sup- ply of tis- sues through- out	House- holds in the USA	5 months' overall supply	Re- sup- ply of tis- sues	None de- scribed.	Report- ed use of tissues “not at all, some of the time,	Reported use “all of the time”: A. vs B.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

	control):	communi-ty setting	with inside layer con-taining:	nose, coughing or sneezing) and to discard after use and to help young children use tissues if de-velop a cold.	spon-sor (Kim-ber-ly-Clark Cor-pora-tion).	5-month peri-od			as re-quired		most of the time, or all of the time”	82% vs 71%	
	A. Viru-cidal nasal tis-sues		A. citric and malic acid plus sodium lauryl sul-phate; B. succinic acid.										
	B. Place-bo tis-sues												
Chard 2019 (ad-di-tion-al de-tails from Chard 2018)	Wa-ter, San-ita-tion, and Hy-giene for Health and Ed-uca-tion in Laot-ian Pri-mary Schools (WASH HELPS)	Pri-mary schools and stu-dents	Pre-vent the spread of pathogens within schools through improved water sup-ply and hygiene facilities and im-proved WASH habits in children at home and through-out the life course	For each school: Water supply for school compound: (borehole, protected dug well with pump, or gravity-fed system) Water tank to supply toi-let and hand-washing station School sanitation facil-ities (3 toilet compart-ments) Hand-washing facilities: 2 sinks with tapped wa-ter and supply of soap available (1 bar of soap/pupil) 3 group hand-washing tables with soap and wa-ter At least 1 drinking water filter per classroom	Provision of school: Water supply, sanitation fa-cilities, hand-washing facilities (individual and group), drinking water filters Behaviour change education and promotion including daily group hygiene ac-tivities Daily hand-wash-ing and cleaning schedules	UNICEF Fac-il-ities pro-vided with-in class-rooms (in Laos) School and teach-ers con-duct-ed daily hand-wash-ing activ-ities with chil-dren. Stu-dents par-tici-pat-	Fac-il-ities pro-vided with-in class-rooms (in Laos) Chil-dren par-tici-pat-ed in group hand-wash-ing and clean-ing.	Pri-mary schools and their class-rooms	One-off provision of water and hy-giene facil-ities Daily hand-washing activities and clean-ing for 1 school year Cleaning schedules posted in at least 1 classroom near toilet.	Wa-ter sup-ply tai-lored to the school re-quire-ments/en-viron-ment. San-ita-tion facil-ities pro-vid-ed as need-ed and des-ig-	Rain wa-ter tank pro-vi-sion af-fect-ed by rain wa-ter sup-ply, so changed to tanks with mo-torised hand pumps or grav-i-ty-fed wa-	Unan-nounced visits every 6 to 8 weeks for struc-tured ob-servations to mea-sure fide-lity and ad-herence Fidelity In-dex score (0 to 20): for hard-ware pro-vided see Table 1 in paper and protocol Adherence index: stu-dent re-port of be-haviour-al out-	Fidelity: 30.9% across all schools and visits Adher-ence: 29.4% Hardware provision: 87.8% of schools School-level ad-herence: 61.4% Group com-pound cleaning: 94.8%, toi-let use: 75.5%, group toi-let clean-ing: 68.3%,

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

				Schedules of daily group hand-washing, compound and toilet cleaning		ed in daily group cleaning activities.			nated for boys, girls, and students with disabilities.	ter supply systems. Theft and animal consumption of supplied soap reduced supply.	comes index score (0 to 4)	group hand-washing: 48.7%, individual hand-washing with soap after toilet use: 23.9%. Further details (Chard 2018)	
				Cost per school: USD 13,000 to 17,500									
Harting 2016	Integrated environmental home-based intervention package (IHIP)	Households and their householders including children	Reduce infections and improve child growth in households in rural communities with limited facilities through a multicomponent, low-cost environmental intervention to improve drinking water,	Per household: "OPTIMA-improved stove": improved ventilated solid-fuel stove Kitchen sink with in-kitchen water connection providing piped water Point-of-use water quality intervention applying solar disinfection to drinking water	Community engagement with local and regional stakeholders in design and development Provision of stoves, kitchen sinks, and plastic bottles for solar water treatment, and hygiene education Training of mothers/caretakers in: - solar drinking-water disinfection (SODIS) ^[20] ac-	Health promoters hired local elementary school teachers and implemented and promoted the inter-	Face-to-face and individual household; mode of delivery of training as individual or group not	Households in rural communities in Peru	Stoves and sinks installed over initial 3 months. Monthly reinforcement over 12 months of SODIS, child and kitchen hygiene Weekly spot checks of compliance	Tailored to particular household facilities and environments as needed and to local beliefs and	Not described	Weekly spot-check observations of household hygiene and environmental health conditions (e.g. presence of SODIS bottles on the roof or kitchen) using a checklist Monthly self-report by	SODIS use: 60% initially and 10% at end of study Self-reported use by mothers: 90% with slight decrease at end Self-reported stove use: 90% daily Sink use: 66% daily

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

			sanitation, personal hygiene, and household air quality developed in pilot (Hartinger 2011; Hartinger 2012) using a participatory approach that addressed local beliefs and cultural views		cording to standard procedures - hand hygiene (washing own and children's hands with soap at critical times ^[21]) - advice to separate animals and their excreta from the kitchen environment Project-initiated repairs	ventions. 4 teams of field staff conducted spot-check observations.	de-scribed		Repairs after 9 months Environmental samples test middle and end of 12-month surveillance.	cultural customs Repairs to stoves as needed and checked at 9 months	mothers of stove and sink use	35% of stoves needed minor repairs, 1% needed major repairs. Best-functioning stoves achieved mean 45% and 27% reduction of PM _{2.5} and CO, respectively, in mothers' personal exposure.	
Huda 2012	Sanitation and Hygiene Education and Water Supply in Bangladesh (SHE-WA-B)	Villages and their households with a child < 5 years old	Reduce illness in children < 5 years by improving hygiene practices, sanitation and water supply and treatment in their household	Materials for training of community hygiene promoters and promotion activities including flip charts and flash cards with messages alerting participants to presence of unobservable "germs" and practices to minimise germs See Box 1 in paper for 11 key messages. ^[22]	Engaging local residents under guidance of local NGOs to develop community action plans addressing: Latrine coverage and usage Access to and use of arsenic-free water Improved hygiene practices, especially hand-washing with soap	Community hygiene promoters (local residents with at least 10 years' schooling trained for 10	Face-to-face delivery to groups (villages and households) and individuals	Villages and households in districts of Bangladesh	18 months overall Expected household visit and courtyard meeting every 2 months Hand-washing opportunities: after own or child's defecation,	Community action plans developed for and by local residents.	Not described	Structured observation of hand-washing and child faeces disposal behaviour in households and spot checks of type of household water and sanitation facilities	HW: Food-related: No significant difference from baseline to 18 months; IG versus CG After anus cleaning: 36% versus 27%

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					Recruitment and appointment of community hygiene promoters	days on behaviour change	Meetings held in courtyards of groups of households.	prior to preparing and serving food, prior to eating and feeding a child					Defecation: 30% versus 23%
					Household visits, courtyard meetings, and social mobilisation activities (e.g. water, sanitation and hygiene fairs, village theatre, group discussions in tea stalls (the social meeting point for village men)) by community promoters	change communication in water, sanitation, and hygiene)	Household visits						No access to latrine decreased from 10.3% to 6.8%.
					Structured observation in households								No significant improvement in access to improved latrines, solid waste disposal, drainage systems, and covered containers for water storage
Ibfelt 2015	Disinfection of toys	Day-care nurseries	Reduce transmission of pathogens via shared toys in daycare environment through regular disinfection treatment	Disinfectants: Turbo Oxysan (Ecolab, Valby, Denmark) for washing machines Sirafan M, Ecolab (1% to 3% benzalkonium chloride, 1% to 3% didecyl-dimethylammonium chloride, and 5% to 7% alcohol ethoxylates) for immersion or wiping	Collection and commercial cleaning of toys from nurseries: - linen and toys suitable for washing machines were washed at 46 °C and subsequently disinfected - toys not suitable for washing machines immersed	Commercial cleaning company: Berendsen A/S, Søborg, Denmark	Cleaning companies collected the toys and linen and cleaned them	Day-care nurseries in Denmark	2 to 3 months overall Cleaning every 2 weeks	Staggered cleaning to ensure children had toys to play with whilst oth-	None de-scribed.	None de-scribed.	None de-scribed.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

					in disinfectant or wiped with microfibre cloth	off-site, then returned them.	ing facility	ers were being cleaned				
Najnin 2019 (see also Qadri 2015 for further details)	2 active interventions: A. Combined cholera vaccine and 'behaviour change communication' intervention	Low-income households and compounds	Prevent or reduce transmission of respiratory illness based on the Integrated Behavioural Model for Water Sanitation and Hygiene (IBM-WASH) theoretical framework (Dreibelbis 2013; Hurland 2013)	A. and B. Cholera vaccine ShanChol™ (Shantha Biotechnics-Sanofi, India) A. Following hardware per compound: a. Hand-washing hardware: (i) Bucket with a tap (provided free of charge) (ii) Soapy water bottle (mixture of a commercially available sachet of powdered detergent (~USD 0.03) with 1.5 L of water in a plastic bottle with a hole punched in the cap) supplied by participating compounds (iii) Bowl to collect rinse water after washing hands (see photo in text or in Najnin 2017 doi.org/10.1093/ije/dyx187) b. Water treatment hardware:	A. and B. Provision of cholera vaccine (2 doses at least 14 days apart) Provision of hand-washing hardware and behaviour change communication activities Encouragement of hand-washing after defecation, after cleaning child's anus, and before preparing food Encouragement to add chlorine to own water vessels Benefits were again explained. Follow-up visits by health promoters	Dushtha Hand-Shasthyawashing Kendra (DSK), an NGO, delivered the hardware and behavioural intervention (throughel in community health promoters). Separate data collectors observed soap available at	Households and compounds (where several households share a common water source, kitchen, and toilets) in Bangladesh	Behaviour change communication messages delivered first (within 3 months of cholera vaccination). Point-of-use water hardware provided 3 months later. Follow-up health promoter visits 3 times in 2 months after hardware installation, then 2 times/month (over nearly 2 years).	Hardware-related problems (breakage/leakage) were addressed on health promoter follow-up visits.	None described.	Unannounced home visits by data collectors who observed presence of soap/soapy water in most convenient place for hand-washing (either reserved in a container or available at the tap) Residual chlorine was measured indicating uptake of chlorine dispenser.	Presence of soap / soapy water and water: A. Hand-washing group compounds: 45% (1,729 / 3,886); B. Vaccine-only group compound: 22% (438 / 1,965); C. Control: 28% (556 / 1991) Residual chlorine present in stored drinking water of 4% (160/3886) of households in the vac-

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

				Dispenser containing liquid sodium hypochlorite			abili- ty.	com- pound and house- hold lev- els.					cine-plus- behaviour- change com- pound and none in the oth- er 2 com- pounds.
				See Figure 2 in Najnin 2017 for photos of both doi.org/10.1093/ije/dyx187									
				and more details.									
				Participants own water vessels for water treatment									
				Print materials for behaviour change to compounds and households									
Gargling													
Goodall 2014	2 ac- tive inter- ven- tions:	Uni- ver- sity stu- dents	Decrease the in- cidence of URTI through increased vitamin D levels (associ- ated with greater frequen- cy and severity of URTIs) and gargling (as pre- ventative measure against URTIs)	A. Vitamin D ₃ : container of 8 capsules of 10,000 IU (purchased from Euro-Pharm International Canada Inc.) Weekly email reminder B. Gargling: 30 mL of tap water 2/day	A. Vitamin D: in- structed to take 1 pill weekly B. Gargling: in- structed to gargle twice daily for 30 seconds All participants received general lifestyle and health advice on sleep, nutrition, hand hygiene, and exercise.	Not spec- ified, pre- sum- ably the re- searchers in- clud- ing a study phar- ma- cist	Vita- min D ₃ sup- plied indi- vidu- ally, but no fur- ther de- tails. Method of lifestyle and health ad- vice pro- vi- sion	In uni- ver- sity stu- dent hous- ing (in resi- dences or off- cam- pus) in Cana- da	2 months overall Vitamin D ₃ : week- ly supple- mentation and email reminder Gargling: 30 mL of water for 30 sec- onds twice daily	None de- scribed.	None de- scribed.	None de- scribed.	None de- scribed.

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

														also not described.
Ide 2014	2 active interventions (no control): A. Green tea gargling B. Water gargling	High school students	Prevent influenza and infection in high school students who are at increased risk from close interaction through gargling as a non-pharmaceutical intervention, specifically green tea containing highly bioactive catechin (-)-epigallocatechin gallate, with possible anti-influenza virus properties	A. Bottled green tea (500 mL) containing a catechin concentration of 37 ± 0.2 mg/dL, including approximately 18% (-)-epigallocatechin gallate (manufactured by the Kakegawa Tea Merchants Association). Concentration measured by high-performance liquid chromatography based on the average concentration in 10 bottles from the same production lot (September 2011) used for gargling in the study. B. Tap water	A. Provision of green tea B. Advice to gargle with tap water and not to gargle green tea during study A. and B. Advice to gargle at least 3 times/day (after arriving at school, after lunch, and after school) Consumption of green tea and other tea was not restricted for either group. Safety monitoring carried out throughout the study (not further described).	Materials supplied by researchers	Green tea supplied individually to students. Mode of gargling advice not described.	High schools in Japan	Gargling 3 times/day for 90 days	None described.	None described.	Daily questionnaire included questions about daily adherence to gargling regimen. Adherence rate of gargling at or above 75%, and absence of green tea gargling when in the water gargling group.	Gargling adherence rate: green tea group: 73.7%; water group: 67.2%	
Sato-mura 2005	2 active inter-	Healthy adults	Prevent URTIs through	A. Water B. 15 to 30 times diluted 7% povidone-iodine (as	Local administrators instructed participants to:	Local project ad-	Not specified,	18 health-care	60 days overall	If diluted povi-	3 partici-	Completion of gargling	9 participants did not com-	

Table 1. Description of interventions in included studies, using the items from the Template for Intervention Description and Replication (TIDieR) checklist (Continued)

ven- tions:	gargling water alone, A. Wa- ter gar- gling B. Povi- done-io- dine gar- gling	indicated by manufac- turer)	- gargle dose of water or povi- done-iodine 3 times/day; - maintain hand- washing routine; - not change oth- er hygiene habits; - not take any cold remedies; - complete gar- gling diary. Weekly monitor- ing of hygienic actions and encourage- ment to keep up assigned inter- vention every week	min- istra- tors (18 health- care pro- fes- sion- als) pro- vid- ed in- struc- tions and mon- itor- ing and en- cour- age- ment.	but like- ly to have been face- to- face and indi- vid- ual- ly, at least ini- tial- ly for in- struc- tions	sites in Japan (4 in north- ern re- gion, 9 in cen- tral re- gion, 5 in west- ern re- gion)	1. Water gargling: 20 mL for 15 s at least 3 times/day 2. Povi- done-io- dine gar- gling: 20 mL of dilution 3 times/day	done-io-pants dine as- signed seri- ous dis- com- fort or was not avail- able, par- tici- pants were al- lowed to gar- gle with wa- ter in- stead.	diary: fre- quency of gargling and hand- washing Weekly monitor- ing and encour- agement by local adminis- trators	plete di- ary. Average frequen- cy of gar- gling / per- son / day: With wa- ter: A: 3.6 B: 0.8 Control: 0.9 With povi- done-io- dine: A.: <0.1 B: 2.9 Control: 0.2
----------------	--	---------------------------------	---	---	---	--	---	--	--	---

ABH: alcohol-based rub
ARI: acute respiratory infection
CDC: Centers for Disease Control and Prevention
CG: control group
CHG: chlorhexidine gluconate
CO: carbon monoxide
DCCs: daycare centres
FM: face masks
HCP: healthcare personnel
HCW: healthcare worker
HH: hand hygiene
HSG: hand sanitiser group
HSW: hand-washing with soap and water
HWWS: hand-washing with soap
IG: intervention group
IHIP: integrated environmental home-based intervention package

ILI: influenza-like illness
IU: international units
LTCFs: long-term care facilities
NGOs: non-governmental organisations
NH: nursing home
no.: number
NPIs: non-pharmaceutical interventions
PM2.5: particulate matter of less than 2.5 microns
RAs: research assistants
RIs: respiratory infections
RTIs: respiratory tract infections
SD: standard deviation
SSTI: skin and soft-tissue infection
SWG: soap-and-water group
TCID: tissue-culture infectious dose
URTI: upper respiratory tract infection
WHO: World Health Organization
wk: week

w/w: weight for weight

[1]: Occupational Safety and Health Administration (OSHA). OSHA technical manual: section VIII: chapter 2: respiratory protection. US Department of Labor. www.osha.gov/dts/osta/otm/otm_viii/otm_viii_2.html (accessed 21 April 2020).

[2]: Ministry of Health and Long-Term Care, Public Health Division, Provincial Infectious Diseases Advisory Committee. Preventing respiratory illnesses: protecting patient and staff: infection control and surveillance standards for febrile respiratory illness (FRI) in non-outbreak conditions in acute care hospitals [September 2005] http://www.health.gov.on.ca/english/providers/program/infectious/diseases/best_prac/bp_fri_080406.pdf (accessed September 11 2009). [URL inactive]

[3]: Before eating, after sneezing, coughing, handling money, using restroom, returning to desk and interacting with others who may be sick

[4]: after coming into classroom, before and after lunch, after break, after physical education, when they went home and after coughing, sneezing or blowing their noses

[5]: after toileting and when visibly dirty plus a protocol for particular circumstances: after coming into the classroom; before and after lunch; after playing outside; when they went home; after coughing, sneezing, or blowing their noses; and after diapering

[6]: 1) when entering into the classroom; 2) after sneezing, coughing, or blowing their nose; 3) after using the toilet/washroom; 4) before eating any food; and 5) when leaving the school at the end of the day

[7]: what to do if hands were dirty, why students should wash their hands, benefits of washing hands and using hand sanitizer, procedure for washing hands using hand sanitizer, to cover mouth and nose with upper part of sleeve while coughing and/or sneezing

[8]: Boyce JM, Pittet D, Healthcare Infection Control Practices Advisory Committee, HICPAC/ SHEA/APIC/IDSA Hand Hygiene Task Force. Guideline for hand hygiene in healthcare settings. Recommendations of the Healthcare Infection Control Practices Advisory Committee and the HICPAC/SHEA/APIC/ IDSA Hand Hygiene Task Force. *MMWR Recommendations and Reports* 2002;51(RR-16):1–45. www.cdc.gov/mmwr/preview/mmwrhtml/rr5116a1.htm (accessed 21 April 2020). International Bank for Reconstruction and Development/ World Bank, Bank-Netherlands Water Partnership, Water and Sanitation Program. Hand washing manual: a guide for developing a hygiene promotion program to increase handwashing with soap. <http://go.worldbank.org/PJTS4A53C0> (Accessed 16 May 2007). [URL inactive] California State Department of Education. *Techniques for Preventing the Spread of Infectious Diseases*. Sacramento (CA): California State Department of Education, 1983. Geiger BF, Artz L, Petri CJ, Winnail SD, Mason JW. *Fun with Handwashing Education*. Birmingham (AL): University of Alabama, 2000. Roberts A, Pareja R, Shaw W, Boyd B, Booth E, Mata JI. A tool box for building health communication capacity. www.globalhealthcommunication.org/tools/29 (Accessed 10 October 2007). [URL inactive] Stark P. *Handwashing Technique. Instructor's Packet. Learning Activity Package*. Sacramento (CA): California State Department of Education, 1982.

[9]: DIN EN 1500: Chemische Desinfektionsmittel und Antiseptika, Hygienische Händedesinfektion, Prüfverfahren und Anforderungen (Phase 2/Stufe 2). Brüssel (Belgium): CEN, European Committee for Standardization 1997;1-20.

[10]: DIN EN 12791: Chemische Desinfektionsmittel und Antiseptika, Chirurgische Händedesinfektionsmittel - Prüfverfahren und Anforderungen (Phase 2/Stufe 2). Brüssel (Belgium): CEN, European Committee for Standardization 2005;1-31.

[11]: after defaecation, after cleaning an infant who had defaecated, before preparing food, before eating, and before feeding infants

[12]: non-governmental organisation that supports community-based health and development initiatives

- [13]: “Healthy Hands” Rules (from Figure 3 in paper): Do use “special soap” when arrive to school, before lunch, after go to bathroom (only if soap and water not available), if rub nose or eyes or if fingers in mouth, if teacher asks. Do not: use “special soap” if hand dirt on them, put “special soap” on another student, play with ‘special soap”, put hands near eyes after using “special soap”.
- [14]: Calculated by subtracting each day’s soap weight from the previous day’s weight. Maximum number of grams of soap consumed for each compound was identified and the day on which the maximum soap consumption was recorded. A per capita estimate of daily soap consumption was calculated
- [15]: National Health and Medical Research Council. Staying Healthy in Child Care. Canberra (Australia): Australian Government Publishing Service, 1994
- [16]: upon arrival, before and after lunch, and prior to departure
- [17]: knowledge and awareness of HH guidelines, perceived importance of performing HH, perceived behavioural control (i.e. perceived ease or difficulty of performing the behaviour), and habit
- [18]: “According to the Dutch national guidelines, HH is mandatory for caregivers before touching/preparing food, before caregivers themselves ate or assisted children with eating, and before wound care; and after diapering, after toilet use/wiping buttocks, after caregivers themselves coughed/sneezed/wiped their own nose, after contact with body fluids (e.g. saliva, vomit, urine, blood, or mucus when wiping children’s noses), after wound care, and after hands were visibly soiled.” (p. 2495)
- [19]: having touched household items being used by the index patients and/or other symptomatic household contacts, and after coughing/sneezing, before meals, before preparing meals and when returning home
- [20]: SODIS: www.sodis.ch/index_EN.html
- [21]: after defecation, after changing diapers, before food preparation and before eating
- [22]: 1. Wash both hands with water and soap before eating/ handling food 2. Wash both hands with water and soap/ash after defecation 3. Wash both hands with water and soap/ash after cleaning baby’s bottom 4. Use hygienic latrine by all family members including Children 5. Dispose of children’s faeces into hygienic latrines 6. Clean and maintain latrine 7. Construct a new latrine if the existing one is full and fill the pit with soil/ash. 8. Safe collection and storage of drinking water 9. Draw drinking water from arsenic safe water point 10. Wash raw fruits and vegetables with safe water before eating and cover food properly 11. Manage menstruation period safely (p.605)

Table 2. Results from trials of hand hygiene compared to control

Study	Comparison (see Table 1 for details of interventions)	Reported outcomes	Results
Alzaher 2018 cluster-RCT Saudi Arabia	Hand-washing workshop and posters vs usual practice	% absence days due to URI	0.39% and 0.72% in intervention group schools; 0.86% and 1.39% in control schools
Arbogast 2016 cluster-RCT USA	Hand sanitiser + wipes + hand foam vs none Both groups received education + signage about hand-washing	1. Health insurance claims for preventable illnesses per employee 2. Absences per employee	1. 0.30 claims in intervention; 0.37 in control (27% relative reduction; P = 0.03) 2. 1.45 in intervention; 1.53 in control (5.0% relative reduction in intervention; P = 0.30)
Azor-Martinez 2016 RCT Spain	Hand-washing with soap and water plus hand sanitiser vs usual hand-washing practices	% absence days due to URI	1.15% in intervention; 1.68% in control. Significantly lower in intervention (P < 0.001)
Azor-Martinez 2018 cluster-RCT Spain	Education and hand hygiene with soap and water vs hand hygiene with sanitiser vs usual hand-washing procedures	1. URI incidence rate ratio (primary) 2. Percentage difference in absenteeism days	1. HH soap vs control 0.94 (95% CI 0.82 to 1.08); HH sanitiser vs control 0.77 (95% CI 0.68 to 0.88); HH soap vs HH sanitiser 1.21 (95% CI 1.06 to 1.39) 2. HH soap 3.9% vs control 4.2% (P < 0.001); HH sanitiser 3.25% vs control 4.2% (P = 0.026); HH soap 3.9% vs HH sanitiser 3.25% (P < 0.001)
Biswas 2019 cluster-RCT Bangladesh	Hand sanitiser and respiratory hygiene education and cough/sneeze hygiene vs no intervention	1. ILI incidence rate (at least 1 episode) 2. Laboratory-confirmed influenza	1. 22 per 1000 student-weeks in intervention; 27 per 1000 student-weeks in control, not statistically significantly different 2. 3 per 1000 student-weeks in intervention; 6 per 1000 student-weeks in control, P = 0.01
Correa 2012 cluster-RCT Colombia	Alcohol-based hand sanitiser in addition to hand-washing vs usual hand-washing practice	ARIs in 3rd trimester of follow-up	Hazard ratio for intervention to control 0.69 (95% CI 0.57 to 0.83)
Cowling 2008 cluster-RCT Hong Kong	Hand hygiene (36 households) vs face mask (mask) vs education (control)	Secondary attack rate for: 1. laboratory-confirmed influenza; 2. ILI definition 1; 3. ILI definition 2; 4. ILI definition 3.	1. HH 0.06; mask 0.07; control 0.06 2. HH 0.18; mask 0.18; control 0.18 3. HH 0.11; mask 0.10; control 0.11 4. HH 0.04; mask 0.08; control 0.04
Cowling 2009 cluster-RCT Hong Kong	Hand hygiene (HH) vs face mask + hand hygiene (HH + mask) vs education (control)	Secondary attack rate for: 1. laboratory-confirmed influenza;	1. HH 5; HH + mask 7; control 10 2. HH 16; HH + mask 21; control 19 3. HH 4; HH + mask 7; control 5

Table 2. Results from trials of hand hygiene compared to control (Continued)

		2. ILI definition 1;	
		3. ILI definition 2.	
DiVita 2011 (conference abstract)	Hand-washing stations with soap and motivation vs none	1. SAR for laboratory-confirmed influenza	1. SAR higher in intervention group (11.0% vs 7.5%)
RCT		2. SAR for ILI	2. SAR higher in intervention group (14.2% vs 11.9%)
Bangladesh			
Feldman 2016 cluster-RCT	Hand disinfection + soap and water installed vs none	1. Number of respiratory infections	1. 11 in each group
Israel		2. Number of off-duty days	2. 112 in intervention; 104 in control
Gwaltney 1980 RCT	Virucidal hand wash vs placebo	1. Number with illness after immediate exposure	1. 0 of 8 in intervention; 7 of 7 in control
USA		2. Number with illness after 2-hour delay in exposure	2. 1 of 10 in intervention; 6 of 10 in control
Hubner 2010 RCT	Hand disinfection provided vs none	Odds ratios (95% CI) (intervention:control)	1. 1.02 (0.20 to 5.23)
Germany		1. Influenza	2. 0.35 (0.17 to 0.71)
		2. Common cold	3. 1.87 (0.52 to 6.74)
		3. Sinusitis	4. 0.62 (0.31 to 1.25)
		4. Sore throat	5. 0.38 (0.14 to 0.99)
		5. Fever	6. 0.45 (0.22 to 0.91)
		6. Cough	
Ladegaard 1999 RCT	Hand hygiene and education vs none	Sick days during the "effect period"	22 days/child in the intervention group vs 36 days/child in the control group
Denmark			
Larson 2010 cluster-RCT	Education vs education with alcohol-based hand sanitiser vs education with hand sanitiser and face masks	Incidence rate ratios (episodes per 1000 person-weeks) for:	1. HS 29; HS + masks 39; control 35
USA		1. URI;	2. HS 1.9; HS + masks 1.6; control 2.3
		2. ILI;	3. HS 0.6; HS + masks 0.5; control 2.3
		3. influenza;	4. HS 0.14; HS + masks 0.12; control 0.14
		Secondary attack rates for:	5. HS 0.02; HS + masks 0.02; control 0.02
		4. URI/ILI/influenza;	
		5. ILI/influenza.	
Little 2015 RCT	Bespoke automated web-based hand hygiene motivational intervention with tailored feedback vs none	Number of participants with 1 or more episodes of URI	Risk ratio for intervention to control 0.86 (95% CI 0.83 to 0.89; P < 0.001)
England			

Table 2. Results from trials of hand hygiene compared to control (Continued)

Luby 2005 RCT Pakistan	Antibacterial soap and education about hand-washing vs plain soap and education vs none	1. Cough or difficulty breathing in children < 15 yrs (episodes/100 person-weeks) 2. Congestion or coryza in children < 15 yrs (episodes/100 person-weeks) 3. Pneumonia in children < 5 yrs (episodes/100 person-weeks)	All outcomes significantly lower than control 1. 4.21 in antibacterial soap group; 4.16 in plain soap group; 8.50 in control group 2. 7.32 in antibacterial soap group; 6.87 in plain soap group; 14.78 in control group 3. 2.42 in antibacterial soap group; 2.20 in plain soap group; 4.40 in control group
Millar 2016 cluster-RCT USA	Standard educational promotion of hand-washing vs enhanced promotion vs promotion plus a once-weekly application of chlorhexidine-based body wash	Incidence rates of ARI over 20 months	37.7 enhanced + body wash; 29.3 enhanced; 35.3 standard; RR for enhanced + body wash to standard 1.07 (95% CI 1.03 to 1.11); RR for enhanced to enhanced + body wash 0.78 (95% CI 0.75 to 0.81)
Morton 2004 cluster-RCT Cross-over study USA	Alcohol gel plus education vs regular hand-washing	Absence due to infectious illness	Results not stated numerically
Nicholson 2014 cluster-RCT India	Combination hand-washing promotion with provision of free soap vs none	Target children: 1. Episodes of ARI (per 100 person-weeks) 2. School absence episodes (per 100 person-days) Families: 3. Episodes of ARI	1. 16 in intervention; 19 in control 2. 1.2 in intervention; 1.7 in control 3. 10 in intervention; 11 in control
Priest 2014 cluster-RCT New Zealand	Hand hygiene education and hand sanitiser vs education alone	1. % absence days due to respiratory illness 2. % absence days due to any illness	1. 0.84% in intervention group; 0.80% in control (P = 0.44) 2. 1.21% in intervention group; 1.16% in control (P = 0.35)
Ram 2015 RCT Bangladesh	Education to promote intensive hand-washing in households plus soap provision vs none	1. Secondary attack ratio for intervention to control for ILI 2. Laboratory-confirmed influenza	1. 1.24 (95% CI 0.93 to 1.65) 2. 2.40 (95% CI 0.68 to 8.47)
Roberts 2000 cluster-RCT Australia	Hand-washing programme with training for staff and children vs none	Incidence rate ratio for ARI	IRR 0.92 for intervention to control (95% CI 0.86 to 0.99)
Sandora 2008 cluster-RCT USA	Hand sanitiser and education vs none	Incidence rates for ARI (episodes per person-month)	0.43 in intervention; 0.42 in control

Table 2. Results from trials of hand hygiene compared to control (Continued)

Savolainen-Kopra 2012 cluster-RCT Finland	Hand hygiene with soap and water (IR1 group) vs with alcohol-based hand rub (IR2 group) vs control (none); intervention groups also received education	1. Number of respiratory infection episodes/week 2. Number of reported infection episodes/week 3. Number of reported sick leave episodes/week	1. 0.076 in IR1; 0.085 in IR2; 0.080 in control, NS 2. 0.097 in IR1; 0.107 in IR2; 0.104 in control, NS 3. 0.042 in IR1; 0.035 in IR2; 0.035 in control. Significantly higher in IR1 compared with control
Simmerman 2011 cluster-RCT Thailand	Hand-washing (HW) vs hand-washing plus paper surgical face masks (HW + FM) vs control (none)	Odds ratios for secondary attack rates for influenza	OR for HW: control 1.20 (95% CI 0.76 to 1.88) OR for HW + masks: control 1.16 (95% CI 0.74 to 1.82) OR for HW + masks: HW 0.72 (95% CI 0.21 to 2.48)
Stebbins 2011 cluster-RCT USA	Training in hand and respiratory (cough) hygiene + hand sanitizer vs none	Incidence rate ratios for intervention to control for: 1. laboratory-confirmed influenza (RT-PCR); 2. influenza-A; 3. absence.	1. IRR 0.81 (95% CI 0.54 to 1.23) 2. IRR 0.48 (95% CI 0.26 to 0.87) 3. IRR 0.74 (95% CI 0.56 to 0.97)
Talaat 2011 cluster-RCT Egypt	Mandatory hand-washing intervention + education vs none	1. Number of absence days due to ILI 2. Number of absence days	1. 917 in intervention; 1671 in control (P < 0.001) 2. 13,247 in intervention; 19,094 in control (P < 0.001)
Temime 2018 cluster-RCT France	Hand hygiene with alcohol-based hand rub, promotion, staff education, and local work groups vs none	Incidence rate of ARI clusters (5 or more people in same nursing home)	2 ARI clusters in intervention; 1 in control
Turner 2012 RCT USA	Antiviral hand treatment vs no treatment	1. Number of rhinovirus infections 2. Common cold infections 3. Rhinovirus-associated illnesses	1. 49 in intervention; 49 in control, NS 2. 56 in intervention; 72 in control, NS 3. 26 in intervention; 24 in control, NS
White 2001 DB-RCT USA	Hand rub with benzalkonium chloride (hand sanitiser) vs placebo	ARI symptoms Laboratory: testing of virucidal and bactericidal activity of the product	30% to 38% decrease of illness and absenteeism (RR for illness absence incidence 0.69; RR for absence duration 0.71)
Yeung 2011 cluster-RCT Hong Kong	Alcohol-based hand gel + materials + education vs control (basic life support workshop)	Difference between pre-study period and poststudy in pneumonia infections recorded in residents	0.63/1000 reduction in intervention group; 0.16/1000 increase in control
Zomer 2015 cluster-RCT Netherlands	4 components: 1. Hand hygiene products, paper towel dispensers, soap, alcohol-based hand sanitiser,	Incidence rate ratio for intervention to control for common cold	IRR 1.07 (95% CI 0.97 to 1.19) 8.2 episodes per child-year in intervention; 7.4 episodes per child-year in control

Table 2. Results from trials of hand hygiene compared to control (Continued)

and hand cream provided for 6 months

2. Training and booklet

3. 2 team training sessions aimed at hand hygiene improvement

4. Posters and stickers for caregivers and children as reminders

Combination vs usual practice

ARI: acute respiratory infection

CI: confidence interval

cluster-RCT: cluster-randomised controlled trial

DB-RCT: double-blind randomised controlled trial

HH: hand hygiene

HS: hand sanitiser

HW: hand-washing

ILI: influenza-like illness

IRR: incidence rate ratio

NS: non-significant

OR: odds ratio

RCT: randomised controlled trial

RR: risk ratio

RT-PCR: reverse-transcriptase polymerase chain reaction

SAR: secondary attack rate

URI: upper respiratory infection

yrs: years

Table 3. Results from trials of hand hygiene + medical/surgical masks compared to control

Study	Comparison (see Table 1 for details of interventions)	Reported outcomes	Results
Aelami 2015 (conference abstract) RCT Saudi Arabia	Hand hygiene education + alcohol-based hand rub + soap + surgical masks vs none	Proportion with ILI (defined as presence of ≥ 2 of the following during their stay: fever, cough, and sore throat)	52% in intervention; 55.3% in control ($P < 0.001$)
Aiello 2010 cluster-RCT USA	Face mask use (FM) vs face masks + hand hygiene (FM + HH) vs control Note that this study is not included in meta-analysis as each treatment group included only 1 cluster.	1. ILI 2. Laboratory-confirmed influenza A or B	Significant reduction in ILI cases in both intervention groups compared with control over weeks 3 to 6 No significant differences between FM and FM + HH
Aiello 2012 cluster-RCT USA	Face mask use (FM) vs face masks + hand hygiene (FM + HH) vs control	1. Clinical ILI 2. Laboratory-confirmed influenza A or B	1. Non-significant reductions in FM group compared with control over all weeks. Significant reduction in FM + HH group compared with control in weeks 3 to 6

Table 3. Results from trials of hand hygiene + medical/surgical masks compared to control (Continued)

			2. Non-significant reductions in both intervention groups compared with control
Cowling 2009 cluster-RCT Hong Kong	Hand hygiene (HH) vs hand hygiene plus face masks (HH + mask) vs control	Secondary attack ratio for: 1. laboratory-confirmed influenza; 2. ILI definition 1; 3. ILI definition 2.	1. HH 5; HH + mask 7; control 10 2. HH 16; HH + mask 21; control 19 3. HH 4; HH + mask 7; control 5
Larson 2010 cluster-RCT USA	Education (control) vs education with alcohol-based hand sanitiser (HS) vs education + HS + face masks (HS + mask)	Incidence rate ratios (episodes per 1000 person-weeks) for: 1. URI; 2. ILI; 3. influenza. Secondary attack rates for: 4. URI/ILI/influenza; 5. ILI/influenza.	1. HS 29; HS + mask 39; control 35 2. HS 1.9; HS + mask 1.6; control 2.3 3. HS 0.6; HS + mask 0.5; control 2.3 4. HS 0.14; HS + mask 0.12; control 0.14 5. HS 0.02; HS + mask 0.02; control 0.02
Simmerman 2011 cluster-RCT Thailand	Control vs hand-washing (HW) vs hand-washing + paper surgical face masks (HW + mask)	Odds ratio for secondary attack rates for influenza	OR for HW: control 1.20 (95% CI 0.76 to 1.88) OR for HW + mask: control 1.16 (95% CI 0.74 to 1.82) OR for HW + mask: HW 0.72 (95% CI 0.21 to 2.48)
Suess 2012 cluster-RCT Germany	Face mask + hand hygiene (mask + HH) vs face masks only (mask) vs none (control)	Secondary attack rates in household contacts: 1. Laboratory-confirmed influenza 2. ILI	1. Mask 9; mask + HH 15; control 23 2. Mask 9; mask + HH 9; control 17

CI: confidence interval

cluster-RCT: cluster-randomised controlled trial

FM: face mask

HH: hand hygiene

HS: hand sanitiser

HW: hand-washing

ILI: influenza-like illness

OR: odds ratio

RCT: randomised controlled trial

URI: upper respiratory infection

Table 4. Results from trials of soap + water compared to hand sanitisers

Study	Comparison (see Table 1 for details of interventions)	Reported outcomes	Results
Azor-Martinez 2018 cluster-RCT Spain	Education and hand hygiene with soap and water (HH soap) vs hand hygiene with sanitiser (HH sanitiser) vs usual hand-washing procedures	1. URI incidence rate ratio (primary) 2. Percentage difference in absenteeism days	1: HH soap vs control 0.94 (95% CI 0.82 to 1.08); HH sanitiser vs control 0.77 (95% CI 0.68 to 0.88); HH soap vs HH sanitiser 1.21 (95% CI 1.06 to 1.39) 2: HH soap 3.9% vs control 4.2% (P < 0.001); HH sanitiser 3.25% vs control 4.2% (P = 0.026); HH soap 3.9% vs HH sanitiser 3.25% (P < 0.001)
Pandepjpong 2012	Alcohol hand gel applied every 60 minutes vs every 120 minutes vs once before lunch (3 groups).	Absent days due to confirmed ILI/present days	0.017 in every hour group; 0.025 in every 2 hours group; 0.026 in before lunch group. Statistically significant difference between every hour group and

Table 4. Results from trials of soap + water compared to hand sanitisers (Continued)

cluster-RCT			before lunch group, and between every hour and every 2 hours groups
Thailand			
Savolainen-Ko- pra 2012	Hand hygiene with soap and water (IR1 group) vs with alcohol-based hand rub (IR2 group) vs control (none); intervention groups also received education	1. Number of respiratory infection episodes/week 2. Number of reported infection episodes/week 3. Number of reported sick leave episodes/week	1. 0.076 in IR1; 0.085 in IR2; 0.080 in control, NS 2: 0.097 in IR1; 0.107 in IR2; 0.104 in control, NS 3: 0.042 in IR1; 0.035 in IR2; 0.035 in control. Significantly higher in IR1 compared with control
cluster-RCT			
Finland			
Turner 2004a and- Turner 2004b	Study 1. Ethanol vs salicylic acid 3.5% vs salicylic acid 1% and pyrog-lutamic acid 3.5% Study 2. Skin cleanser wipe vs ethanol (control)	% of volunteers infected with rhinovirus	7% in each intervention group; 32% in control (study 1) 22% in intervention, 30% in control (study 2)
RCT			
Canada			

CI: confidence interval
cluster-RCT: cluster-randomised controlled trial
HH: hand hygiene
ILI: influenza-like illness
NS: non-significant
RCT: randomised controlled trial
URI: upper respiratory infection

Table 5. Results from trials of surface/object disinfection (with or without hand hygiene) compared to control

Study	Comparison (see Table 1 for details of interventions)	Reported outcomes	Results
Ban 2015	Hand hygiene products, surface cleaning and disinfection provided to families and kindergartens vs none	1. Respiratory illness 2. Cough and expectoration	1. OR 0.47 for intervention to control (95% CI 0.38 to 0.59) 2. OR 0.56 (95% CI 0.48 to 0.65)
cluster-RCT			
China			
Carabin 1999	One-off hygiene education and disinfection of toys with bleach vs none	Difference in incidence rate for URTI (cluster-level result)	0.28 episodes per 100 child-days lower in intervention group (95% CI 1.65 lower to 1.08 higher); URTI incidence rate IRR 0.80 (95% CI 0.68 to 0.93)
cluster-RCT			
Canada			
Ibfelt 2015	Disinfectant washing of linen and toys by commercial company every 2 weeks vs usual care	Presence of respiratory viruses on surfaces	Statistically significant reduction in intervention group in adenovirus, rhinovirus, RSV, metapneumovirus, but not other viruses including coronavirus
cluster-RCT			
Denmark			
Kotch 1994	Training in hand-washing and diapering and disinfection of surfaces vs none	Respiratory illness incidence rate in: 1. children < 24 months; 2. children ≥ 24 months.	1. 14.78 episodes per child-year in intervention; 15.66 in control 2. 12.87 in intervention; 11.77 in control
RCT			
USA			

Table 5. Results from trials of surface/object disinfection (with or without hand hygiene) compared to control

Study	Comparison	Reported outcomes	Results
McConeghy 2017 RCT USA	Staff education, cleaning products, and audit of compliance and feedback vs none	Infection rates	Upper respiratory infections not reliably recorded or reported.
Sandora 2008 cluster-RCT USA	Hand sanitiser and disinfection of classroom surfaces vs materials about good nutrition (control)	Absence due to respiratory illness (multi-variable analysis)	Rate ratio 1.10 for intervention to control (95% CI 0.97 to 1.24)

CI: confidence interval
 cluster-RCT: cluster-randomised controlled trial
 IRR: incident rate ratio
 OR: odds ratio
 RCT: randomised controlled trial
 RSV: respiratory syncytial virus
 URTI: upper respiratory tract infection

Table 6. Results from trials of complex interventions compared to control

Study	Comparison (see Table 1 for details of interventions)	Reported outcomes	Results
Complex hygiene and sanitation interventions compared to control			
Chard 2019 cluster-RCT Laos	Complex sanitation intervention and education vs none	Pupil-reported symptoms of respiratory infection over 1 week	NS difference between groups. 29% of intervention group; 32% control group; adjusted risk ratio 1.08 (95% CI 0.95 to 1.23)
Hartinger 2016 cluster-RCT Peru	Cooking and sanitation provision and education vs none	Number of ARI episodes per child-year	NS difference between groups. Risk ratio for intervention to control 0.95 (95% CI 0.82 to 1.10)
Huda 2012 cluster-RCT Bangladesh	Sanitation provision and education vs none	Respiratory illness	12.6% in intervention group; 13.0% in control group. Not adjusted for multiple outcome measurements. No CIs reported.
Najnin 2019 cluster-RCT Bangladesh	Sanitation and behaviour change intervention (plus cholera vaccine) vs none	Respiratory illness in past 2 days	2.8% in intervention group; 2.9% in control group

ARI: acute respiratory infection
 CI: confidence interval
 cluster-RCT: cluster-randomised controlled trial
 NS: non-significant
 RCT: randomised controlled trial

Table 7. Results from trials of virucidal tissues compared to control

Study	Comparison	Reported outcomes	Results
Virucidal tissues compared with placebo or no tissues			
Farr 1988a and Farr 1988b cluster-RCT USA Trial 1 and Trial 2	Trial 1. Virucidal nasal tissues vs placebo vs none Trial 2. Virucidal nasal tissues vs placebo	Respiratory illnesses per person over 24 weeks Trial 1 Trial 2	Trial 1: 3.4 in tissues group; 3.9 in placebo group; 3.6 in no-tissues group Trial 2: 3.4 in tissues group; 3.6 in placebo group NS
Longini 1988 DB-PC RCT USA	Virucidal nasal tissues vs placebo	Secondary attack rate of viral infections (number of infections in household members of index case)	10.0 in intervention; 14.3 in placebo; NS

cluster-RCT: cluster-randomised controlled trial

DB-PC: double-blind, placebo-controlled

NS: non-significant

RCT: randomised controlled trial

vs: versus

Table 8. Summary of main results of the review for the primary outcomes

Interventions	RCT/cluster-RCT (N = 67)
Medical/surgical masks	<p>Masks (medical/surgical) compared to no masks 9 trials no effect on ILI (RR 0.99, 0.82 to 1.18) (Aiello 2010; Barasheed 2014; Canini 2010; Cowling 2008; Jacobs 2009; MacIntyre 2009; MacIntyre 2015; MacIntyre 2016; Suess 2012); 6 trials no effect on laboratory-confirmed influenza 95% CI RR 0.84 (0.61 to 1.17) (Aiello 2012; Cowling 2008; MacIntyre 2009; MacIntyre 2015; MacIntyre 2016; Suess 2012); 2 trials in HCWs no effect on ILI (RR 0.37, 0.05 to 2.50) (Jacobs 2009; MacIntyre 2015).</p> <p>Medical/surgical masks vs other (non-N95) masks: 1 trial more ILI with cloth mask (RR 13.25, 1.74 to 100.97) (MacIntyre 2015); 1 trial no effect of catechin-treated masks on influenza (adjusted OR 2.35, 0.40 to 13.72) (Ide 2016).</p>
N95 respirator	<p>N95 respirators compared to medical/surgical masks</p> <p>3 trials no difference for clinical respiratory illness (RR 0.70, 0.45 to 1.10) (MacIntyre 2011; MacIntyre 2013; Radonovich 2019);</p> <p>4 trials no difference for ILI (95% CI RR 0.81, 0.62 to 1.05) (Loeb 2009; MacIntyre 2009; MacIntyre 2011; Radonovich 2019); 4 trials no difference for laboratory-confirmed influenza (95% CI RR 1.06, 0.81 to 1.38) (Loeb 2009; MacIntyre 2009; MacIntyre 2011; Radonovich 2019).</p> <p>4 studies conducted in HCWs, 3 trials no difference for clinical respiratory illness (RR 0.70, 0.45 to 1.10) (MacIntyre 2011; MacIntyre 2013; Radonovich 2019); 3 trials no difference for ILI (RR 0.64, 0.32 to 1.31) (Loeb 2009; MacIntyre 2011; Radonovich 2019); 3 trials no difference for laboratory-confirmed ILI (RR 1.02, 0.73 to 1.43) (Loeb 2009; MacIntyre 2011; Radonovich 2019).</p>
Hand hygiene	<p>Hand hygiene compared to control 16 trials found effect on combined outcome (ARI or ILI or influenza) (RR 0.89, 0.84 to 0.95) (Azor-Martinez 2018; Biswas 2019; Correa 2012; Cowling 2008; Cowling 2009; Hubner 2010; Larson 2010; Little 2015; Millar 2016; Nicholson 2014; Ram 2015; Roberts 2000; Sandora 2005; Simmerman 2011; Stebbins 2011; Zomer 2015); 7 trials effect on ARI (RR 0.84, 0.82 to 0.86) (Azor-Martinez 2018; Correa 2012; Larson 2010; Little 2015; Millar 2016; Nicholson 2014; Sandora 2005); 10 trials no effect on ILI</p>

Table 8. Summary of main results of the review for the primary outcomes (Continued)

	(RR 0.98, 0.85 to 1.13) (Biswas 2019; Cowling 2008; Cowling 2009; Hubner 2010; Larson 2010; Little 2015; Ram 2015; Roberts 2000; Simmerman 2011; Zomer 2015); 8 trials no effect on laboratory-confirmed influenza (RR 0.91, 95% CI 0.63 to 1.30) (Biswas 2019; Cowling 2008; Cowling 2009; Hubner 2010; Larson 2010; Ram 2015; Simmerman 2011; Stebbins 2011)
Hand hygiene + medical/surgical masks	<p>Hand hygiene + medical/surgical masks compared to control</p> <p>7 trials no effect on ILI (95% CI RR 0.97, 0.80 to 1.19) (Aelami 2015; Aiello 2010; Aiello 2012; Cowling 2009; Larson 2010; Simmerman 2011; Suess 2012); 4 trials no effect on laboratory-confirmed influenza (RR 0.97, 0.69 to 1.36) (Cowling 2009; Larson 2010; Simmerman 2011; Suess 2012).</p> <p>Hand hygiene + medical/surgical masks compared to hand hygiene</p> <p>3 trials no effect on ILI (RR 1.03, 0.69 to 1.53) or laboratory-confirmed influenza (RR 0.99, 0.69 to 1.44) (Cowling 2009; Larson 2010; Simmerman 2011).</p>
Soap + water compared to sanitiser, and comparisons of different types of sanitiser	<p>Soap + water compared to sanitiser, and comparisons of different types of sanitiser</p> <p>1 trial hand sanitiser was more effective than soap and water (Azor-Martinez 2018); 1 trial there was no difference (Savolainen-Kopra 2012).</p> <p>2 trials in children antiseptic was more effective (Morton 2004; White 2001); 1 trial in children antiseptic = soap (Luby 2005).</p> <p>1 trial hand sanitisers were better than placebo, but no difference between sanitisers (Turner 2004a); 1 trial no difference between different wipes (Turner 2004b).</p>
Surface/object disinfection (with or without hand hygiene) compared to control	<p>Surface/object disinfection compared to control</p> <p>2 trials were effective on ARI (Ban 2015; Carabin 1999); 1 trial was effective for viruses detected on surface (Ibfelt 2015); 2 trials showed no difference in ARIs (Kotch 1994; McConeghy 2017).</p>
Disinfection of living quarters	-
Complex interventions	<p>Complex interventions compared to control</p> <p>4 trials in low-income countries found no effect on respiratory viral illness (Chard 2019; Hartinger 2016; Huda 2012; Najnin 2019).</p>
Physical interventions (masks, gloves, gowns combined)	-
Gloves	-
Gowns	-
Physical distancing	-
Quarantine in the community	<p>Quarantine compared to control</p> <p>1 trial effective for influenza (Cox hazard ratio 0.799, 95% CI 0.66 to 0.97) (Miyaki 2011).</p>
Eye protection	-
Gargling	<p>Gargling compared to control</p> <p>1 trial gargling with tap water was effective, povidone-iodine was not effective (Satomura 2005); 1 trial gargling with green tea was not more effective than tap water (Ide 2014); 1 trial gargling with water was not effective (Goodall 2014); pooling of 2 trials no effect of gargling (RR 0.91, 95% CI 0.63 to 1.31) (Goodall 2014; Satomura 2005).</p>
Virucidal tissues	<p>Virucidal tissues compared to control</p>

Table 8. Summary of main results of the review for the primary outcomes (Continued)

1 trial had a small effect (Farr 1988a) ("The study authors conclude that virucidal tissues have only a small impact upon the overall rate of natural acute respiratory illnesses"); 2 trials non-significant difference (Farr 1988b; Longini 1988).

Nose wash	-
-----------	---

ARI: acute respiratory infection
 CI: confidence interval
 HCW: healthcare worker
 ILI: influenza-like illness
 OR: odds ratio
 RCT: randomised controlled trial
 RR: risk ratio

Table 9. Trial authors' outcome definitions

	Study	Outcomes definitions
Masks (n = 13)		
1.	<p>Cowling 2008</p> <p>cluster-RCT</p> <p>Hong Kong</p>	<p>Laboratory:</p> <p>QuickVue Influenza A+B rapid test</p> <p>Viral culture on MDCK (Madin-Darby canine kidney cells)</p> <p>Samples were harvested using NTS, but the text refers to a second procedure from June 2007 onwards with testing for influenza viruses on index participants with a negative QuickVue result but a fever $\geq 38^\circ\text{C}$ who were also randomised and further followed up. Data on clinical signs and symptoms were collected for all participants, and an additional NTS was collected for later confirmation of influenza infection by viral culture. It is noteworthy that dropout was higher in households of index participants who had a negative result on the rapid influenza test (25/44, 57%) compared to those who had a positive result (45/154, 29%).</p> <p>Effectiveness: secondary attack ratios (SAR): SAR is the proportion of household contacts of an index case who subsequently were ill with influenza (symptomatic contact individuals with at least 1 NTS positive for influenza by viral culture or PCR)</p> <p>3 clinical definitions were used for secondary analysis:</p> <ol style="list-style-type: none"> 1. fever $\geq 38^\circ\text{C}$ or at least 2 of the following symptoms: headache, coryza, sore throat, muscle aches and pains; 2. at least 2 of the following S/S: fever $\geq 37.8^\circ\text{C}$, cough, headache, sore throat and muscle aches and pains; and 3. fever of $\geq 37.8^\circ\text{C}$ plus cough or sore throat. <p>Safety: harms were not mentioned as an outcome in the methods, but it was reported in the results that there were no adverse events.</p>
2.	<p>Jacobs 2009</p> <p>RCT</p> <p>Japan</p>	<p>Laboratory-confirmation not reported.</p> <p>Effectiveness: URTI is defined on the basis of a symptom score with a score > 14 being a URTI according to Jackson's 1958 criteria ("Jackson score"). These are not explained in text, although the symptoms are listed in Table 3 (any, sore throat, runny nose, stuffy nose, sneeze, cough, headache, earache, feel bad) together with their mean and scores (SD) by intervention arm.</p> <p>Safety: the text does not mention or report harms. These appear to be indistinguishable from URTI symptoms (e.g. headache, which is reported as of significantly longer duration in the intervention arm). Compliance is self-reported as high (84.3% of participants).</p>

Table 9. Trial authors' outcome definitions (Continued)

3.	Loeb 2009 cluster-RCT HCW Canada	<p>Clinical respiratory illness, influenza-like illness, and laboratory-confirmed respiratory virus infection.</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom. 2. Influenza-like illness, defined as fever $\geq 38^\circ\text{C}$ plus 1 respiratory symptom. 3. Laboratory-confirmed viral respiratory infection. Laboratory confirmation was by nucleic acid detection using multiplex RT-PCR for 17 respiratory viruses. <p>Safety: harms were not mentioned as an outcome in the methods, but it is stated in the results that no adverse events were reported by participants.</p>
4.	MacIntyre 2009 cluster-RCT Australia	<p>Eligibility criteria were stipulated as follows:</p> <ol style="list-style-type: none"> 1. the household contained > 2 adults > 16 years of age and 1 child 0 to 15 years of age; 2. the index child had fever (temperature > 37.8 °C) and either a cough or sore throat; 3. the child was the first and only person to become ill in the family in the previous 2 weeks; 4. adult caregivers consented to participate in the study; and 5. the index child was not admitted to the hospital. <p>Definitions used for outcomes:</p> <ol style="list-style-type: none"> 1. ILI defined by the presence of fever (temperature > 37.8 °C), feeling feverish or a history of fever, > 2 symptoms (sore throat, cough, sneezing, runny nose, nasal congestion, headache), or 1 of the symptoms listed plus laboratory confirmation of respiratory viral infection. 2. Laboratory confirmation: multiplex RT-PCR tests to detect influenza A and B and RSV, PIV types 1–3, picornaviruses (enteroviruses or rhinoviruses), adenoviruses, coronaviruses 229E and OC43, and hMPV plus ≥ 1 sym <p>Effectiveness: presence of ILI or a laboratory diagnosis of respiratory virus infection within 1 week of enrolment.</p> <p>Safety: harms not mentioned as an outcome in the methods, but it is reported in the results that more than 50% of participants reported concerns with mask wearing, mainly that wearing a face mask was uncomfortable, but there were no significant differences between the P2 (N95) and surgical mask groups. Other concerns were that the child did not want the parent wearing a mask.</p>
5.	Aiello 2010 cluster-RCT USA	<p>Laboratory details are described in appendix.</p> <p>Effectiveness: ILI, defined as cough and at least 1 constitutional symptom (fever/feverishness, chills, headache, myalgia). ILI cases were given contact nurses phone numbers to record the illness and paid USD 25 to provide a throat swab. 368 participants had ILI, 94 of which had a throat swab analysed by PCR. 10 of these were positive for influenza (7 for A and 3 for B), respectively by arm 2, 5 and 3 using PCR, 7 using cell culture.</p> <p>Safety: no outcomes on harms planned or reported.</p>
6.	Canini 2010 cluster-RCT USA	<p>The primary endpoint was the proportion of household contacts who developed an ILI during the 7 days following inclusion. Exploratory cluster-level efficacy outcome, the proportion of households with 1 or more secondary illness in household contacts.</p> <p>A temperature over 37.8 °C with cough or sore throat was used as primary clinical case definition.</p> <p>The authors also used a more sensitive case definition based on a temperature over 37.8 °C or at least 2 of the following: sore throat, cough, runny nose, or fatigue.</p>

Table 9. Trial authors' outcome definitions (Continued)

		<p>Safety: adverse reactions due to mask wearing were reported, with 38 (75%) participants in the intervention arm experiencing discomfort with mask use due to warmth (45%), respiratory difficulties (33%), and humidity (33%). Children wearing children face masks reported feeling pain more frequently than other participants wearing adult face masks ($P = 0.036$).</p>
7.	<p>Aiello 2012</p> <p>cluster-RCT in halls of residence in the USA</p>	<p>Clinically verified ILI - case definition (presence of cough and at least 1 or more of fever/feverishness, chills, or body aches)</p> <p>Laboratory-confirmed influenza A or B. Throat swab specimens were tested for influenza A or B using real-time PCR.</p> <p>Safety: no outcomes on harms planned or reported.</p>
8.	<p>Barasheed 2014</p> <p>cluster-RCT Saudi Arabia</p>	<p>Laboratory: 2 nasal swabs from all ILI cases and contacts. 1 for influenza POCT using the QuickVue Influenza (A+B) assay (Quidel Corporation, San Diego, USA) and 1 for later NAT for influenza and other respiratory viruses. However, there was a problem with getting POCT on time during Hajj.</p> <p>Effectiveness: to assess the effectiveness of face masks in the prevention of transmission of ILI. ILI was defined as subjective (or proven) fever plus 1 respiratory symptom (e.g. dry or productive cough, runny nose, sore throat, shortness of breath).</p> <p>Safety: no outcomes on harms planned or reported.</p>
9.	<p>MacIntyre 2011</p> <p>cluster-RCT China</p>	<p>Clinical respiratory illness</p> <p>Influenza-like illness</p> <p>Laboratory-confirmed viral respiratory infection</p> <p>Laboratory-confirmed influenza A or B</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory or 1 respiratory symptom and a systemic symptom. 2. Influenza-like illness, defined as fever ≥ 38 °C plus 1 respiratory symptom (i.e. cough, runny nose, etc.). 3. Laboratory-confirmed viral respiratory infection (detection of adenoviruses, human metapneumovirus, coronavirus 229E/NL63, parainfluenza viruses 1, 2, and 3, influenza viruses A and B, respiratory syncytial virus A and B, rhinovirus A/B and coronavirus OC43/HKU1 by multiplex PCR). 4. Laboratory-confirmed influenza A or B. 5. Adherence with mask/respirator use. <p>Safety: adherence and adverse effects of mask wearing were collected at exit interviews 4 weeks' poststudy. Significantly higher adverse events with N95 respirator compared to medical mask for discomfort, headache, difficulty breathing, nose pressure, trouble communicating, not wearing, and unspecified "other" side effects. Over 50% of those wearing N95 respirators reported adverse events. Of those wearing medical masks versus N95 respirators, 85.5% (420/491) versus 47.4% (447/943) reported no adverse events ($P < 0.001$), respectively.</p>
10.	<p>MacIntyre 2013</p> <p>cluster-RCT China</p>	<p>Laboratory:</p> <ol style="list-style-type: none"> 1. Laboratory-confirmed viral respiratory infection in symptomatic participants, defined as detection of adenoviruses; human metapneumovirus; coronaviruses 229E/NL63 and OC43/HKU1; parainfluenza viruses 1, 2, and 3; influenza viruses A and B; respiratory syncytial viruses A and B; or rhinoviruses A/B by NAT using a commercial multiplex PCR (Seegen, Inc., Seoul, Korea). 2. Laboratory-confirmed influenza A or B in symptomatic participants.

Table 9. Trial authors' outcome definitions (Continued)

		<p>3. Laboratory-confirmed bacterial colonisation in symptomatic participants, defined as detection of <i>Streptococcus pneumoniae</i>, <i>Legionella</i>, <i>Bordetella pertussis</i>, <i>Chlamydia</i>, <i>Mycoplasma pneumoniae</i>, or <i>Haemophilus influenzae</i> type B by multiplex PCR (Seegen, Inc.).</p> <p>Effectiveness: clinical respiratory illness defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom. ILI defined as fever (38 °C) plus 1 respiratory symptom.</p> <p>Safety: adverse effects measured using a semi-structured questionnaire. Investigators stated that there was higher reported adverse effects and discomfort of N95 respirators compared with the other 2 arms. In terms of comfort, 52% (297 of 571) of the medical mask arm reported no problems, compared with 62% (317 of 512) of the targeted arm and 38% (217 of 574) of the N95 arm (P < 0.001).</p>
11.	MacIntyre 2015 cluster-RCT Vietnam	<p>Clinical respiratory illness, influenza-like illness, and laboratory-confirmed respiratory virus infection.</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms or 1 respiratory symptom and a systemic symptom. 2. Influenza-like illness, defined as fever ≥ 38 °C plus 1 respiratory symptom. 3. Laboratory-confirmed viral respiratory infection. Laboratory confirmation was by nucleic acid detection using multiplex RT-PCR for 17 respiratory viruses. <p>Safety: adverse events associated with face mask use were reported in 40.4% (227/562) of HCWs in the medical/surgical mask arm and 42.6% (242/568) in the cloth mask arm (P = 0.45). The most frequently reported adverse events were: general discomfort (35.1%; 397/1130) and breathing problems (18.3%; 207/1130). The rate of ILI was higher in the cloth mask arm compared to medical/surgical masks (RR 13.25, 95% CI 1.74 to 100.97).</p>
12.	MacIntyre 2016 cluster-RCT China	<p>Clinical respiratory illness, influenza-like illness, and laboratory-confirmed viral respiratory infection.</p> <ol style="list-style-type: none"> 1. Clinical respiratory illness, defined as 2 or more respiratory symptoms (cough, nasal congestion, runny nose, sore throat, or sneezes) or 1 respiratory symptom and a systemic symptom (chill, lethargy, loss of appetite, abdominal pain, muscle or joint aches). 2. Influenza-like illness, defined as fever ≥ 38 °C plus 1 respiratory symptom. 3. Laboratory-confirmed viral respiratory infection, defined as detection of adenoviruses, human metapneumovirus, coronaviruses 229E/NL63 and OC43/HKU1, parainfluenza viruses 1, 2, and 3, influenza viruses A and B, respiratory syncytial virus A and B, or rhinovirus A/B by NAT using a commercial multiplex PCR. <p>Safety: no outcomes on harms planned or reported.</p>
13.	Radonovich 2019 cluster-RCT USA	<p>Laboratory. Primary outcome: incidence of laboratory-confirmed influenza, defined as:</p> <ol style="list-style-type: none"> 1. detection of influenza A or B virus by RT-PCR in an upper respiratory specimen collected within 7 days of symptom onset; 2. detection of influenza from a randomly obtained swab from an asymptomatic participant; and 3. influenza seroconversion (symptomatic or asymptomatic), defined as at least a 4-fold rise in haemagglutination inhibition antibody titres to influenza A or B virus between pre-season and postseason serological samples deemed not attributable to vaccination. <p>Effectiveness. Secondary outcomes: incidence of 4 measures of viral respiratory illness or infection as follows:</p>

Table 9. Trial authors' outcome definitions (Continued)

1. acute respiratory illness with or without laboratory confirmation;
2. laboratory-detected respiratory infection, defined as detection of a respiratory pathogen by PCR or serological evidence of infection with a respiratory pathogen during the study surveillance period(s), which was added to the protocol prior to data analysis; and
3. laboratory-confirmed respiratory illness, identified as previously described (defined as self-reported acute respiratory illness plus the presence of at least PCR-confirmed viral pathogen in a specimen collected from the upper respiratory tract within 7 days of the reported symptoms and/or at least a 4-fold rise from pre-intervention to postintervention serum antibody titres to influenza A or B virus).

Influenza-like illness, defined as temperature of at least 100 °F (37.8 °C) plus cough and/or a sore throat, with or without laboratory confirmation.

Safety: 19 participants reported skin irritation or worsening acne during years 3 and 4 at 1 site in the N95 respirator group.

Hand and hygiene (n = 32)

14.	Alzaher 2018 cluster-RCT Saudi Arabia	Episode of URI was defined as having 2 of the following symptoms for a day or 1 of the symptoms for 2 or more consecutive days: 1) a runny nose, 2) a stuffy or blocked nose or noisy breathing, 3) sneezing, 4) a cough, 5) a sore throat, and 6) feeling hot, having a fever or a chill.
15.	Arbogast 2016 cluster-RCT USA	ICD-9 used: 46611: acute bronchiolitis due to respiratory syncytial virus, 46619: acute bronchiolitis due to other infectious organisms, 4800: pneumonia due to adenovirus, 4809: viral pneumonia, unspecified, 4870: influenza with pneumonia, 07999: unspecified viral infection, 4658: acute upper respiratory infections of other multiple sites, 4659: acute upper respiratory infections of unspecified site, 4871: influenza with other respiratory manifestations.
16.	Azor-Martinez 2016 RCT Spain	Upper respiratory illness was defined as 2 of the following symptoms during 1 day, or 1 of the symptoms for 2 consecutive days: (1) runny nose; (2) stuffy or blocked nose or noisy breathing; (3) cough; (4) feeling hot or feverish or having chills; (5) sore throat; or (6) sneezing.
17.	Azor-Martinez 2018 RCT Spain	Respiratory illness (RI) was defined as the presence of 2 of the following symptoms during 1 day or the presence of 1 of the symptoms for 2 consecutive days: (1) runny nose, (2) stuffy or blocked nose or noisy breathing, (3) cough, (4) feeling hot or feverish or having chills, (5) sore throat, or (6) sneezing. ICD-10 and ICD-9 diagnosis codes used: nonspecific upper respiratory tract infection (465.9), otitis media (382.9), pharyngotonsillitis (463), lower respiratory tract infections (485 and 486), acute bronchitis (490), and bronchiolitis (466.19). Study authors combined the bronchopneumonia code (485) and pneumonia code (486) under the label "lower respiratory tract infections." If > 1 antibiotic was prescribed during an episode, they used the first prescription for analysis. The final diagnosis was done by the medical researchers on the basis of the symptoms described above and a review of the medical history of children with RIs.
18.	Biswas 2019 cluster-RCT Bangladesh	Influenza-like illness: an ILI episode was defined as measured fever > 38 °C or subjective fever and cough. Laboratory-confirmed influenza Nasal swabs for real-time RT-PCR.
19.	Correa 2012	Acute respiratory infection was defined as 2 or more of the following symptoms for at least 24 hours, lasting at least 2 days: runny, stuffy, or blocked nose or noisy

Table 9. Trial authors' outcome definitions (Continued)

	cluster-RCT Colombia	breathing; cough; fever, hot sensation, or chills; and/or sore throat. Ear pain alone was considered ARI alternately.
20.	Cowling 2009 cluster-RCT Hong Kong	Laboratory-confirmed of influenza virus infection by RT-PCR for influenza A and B virus. Clinical influenza-like illness: used 2 clinical definitions of influenza based on self-reported data from the symptom diaries as secondary analyses. The first definition of clinical influenza was at least 2 of the following signs and symptoms: temperature 37.8 °C or greater, cough, headache, sore throat, and myalgia; the second definition was temperature 37.8 °C or greater plus cough or sore throat.
21.	DiVita 2011 (conference abstract) RCT Bangladesh	Influenza-like illness was defined as fever in children < 5 years old and fever with cough or sore throat in individuals > 5 years old.
22.	Feldman 2016 cluster-RCT Israel	Infectious diseases grouped into diarrhoeal, respiratory, and skin infection. Based on ICD-9, but no supplementary material was accessible for further definition (Supplementary Material C lists all ICD-9 diagnoses tallied in this "outcome").
23.	Gwaltney 1980 RCT USA	Viral cultures and serology if rhinovirus in laboratory-inoculation
24.	Hubner 2010 RCT Germany	Assessing illness rates due to common cold and diarrhoea. Collecting data on illness symptoms (common cold, sinusitis, sore throat, fever, cough, bronchitis, pneumonia, influenza, diarrhoea) and associated absenteeism at the end of every month. Definitions of symptoms were given to the participants as part of the individual information at the beginning of the study. Whilst most symptoms are quite self-explanatory, "influenza" and "pneumonia" are specific diagnoses that were confirmed by professional diagnosis only. Similarly, (self-) diagnosis of "fever" required objective measurement with a thermometer.
25.	Ladegaard 1999 RCT Denmark	Laboratory: serological evidence Effectiveness: influenza-like illness (described as fever, history of fever or feeling feverish in the past week, myalgia, arthralgia, sore throat, cough, sneezing, runny nose, nasal congestion, headache). However, a positive laboratory finding for influenza converts the ILI definition into one of influenza.
26.	Larson 2010 cluster-RCT USA	Study goals: rates of symptoms and secondary transmission of URIs, incidence of virologically confirmed influenza, knowledge of prevention and treatment strategies for influenza and URIs, and rates of influenza vaccination. 1. Laboratory-confirmed influenza: nasal swabs to test for influenza types A and B as well as other common respiratory viruses by rapid culture (R-Mix, Diagnostic Hybrids, Inc., Athens, OH, USA). PCR and subtyping of the samples was done during the second half of the second year of the study. 2. Influenza-like illness: CDC definition of ILI from the Sentinel Physicians' Network was used to determine when masks should be worn: "temperature of $\geq 37.8^{\circ}\text{C}$ and cough and/or sore throat in the absence of a known cause other than influenza".

Table 9. Trial authors' outcome definitions (Continued)

		3. Episodes of URI = upper respiratory infection: not clear, no explicitly stated definition, reported that the most commonly reported URI symptoms are cough or rhinorrhoea.
27.	Little 2015 RCT England	Respiratory tract infections defined as 2 symptoms of an RTI for at least 1 day or 1 symptom for 2 consecutive days. For reported ILI, study authors did not use WHO or CDC definitions because these definitions require measured temperature, and thus were not appropriate (participants were not included after a clinical examination), and they did not use the European Centre for Disease Prevention and Control definition (1 systemic and 1 respiratory symptom) because, according to the international influenza collaboration, this definition does not necessarily differentiate ILI from a common cold. Influenzanet suggests making high temperature a separate element. Their pragmatic definition of ILI therefore required a high temperature (feeling very hot or very cold; or measured temperature > 37.5 °C), a respiratory symptom (sore throat, cough, or runny nose), and a systemic symptom (headache, severe fatigue, severe muscle aches, or severe malaise).
28.	Luby 2005 RCT Pakistan	Defined pneumonia in children according to the WHO clinical case definition: cough or difficulty breathing with a raised respiratory rate (> 60 per minute in individuals younger than 60 days old, > 50 per minute for those aged 60 to 364 days, and > 40 per minute for those aged 1 to 5 years)
29.	Millar 2016 cluster-RCT USA	Medically attended, outpatient cases of acute respiratory infection in the study population. The case definition was any occurrence of the following International Classification of Disease, 9 Revision, Clinical Modification (ICD-9) symptom or disease-specific codes: 460-466, 480-488, and specifically 465.9, 482.9, 486, and 487.1.
		Acute respiratory infections (460 to 466) 460 Acute nasopharyngitis (common cold) 461 Acute sinusitis 462 Acute pharyngitis 463 Acute tonsillitis 464 Acute laryngitis and tracheitis 465 Acute upper respiratory infections of multiple or unspecified sites 466 Acute bronchitis and bronchiolitis
		Pneumonia and influenza (480 to 488) 480 Viral pneumonia 481 Pneumococcal pneumonia (<i>Streptococcus pneumoniae</i> pneumonia) 482 Other bacterial pneumonia 483 Pneumonia due to other specified organism 484 Pneumonia in infectious diseases classified elsewhere 485 Bronchopneumonia, organism unspecified 486 Pneumonia, organism unspecified 487 Influenza 488 Influenza due to identified avian influenza virus 465.9 Acute upper respiratory infections of unspecified site

Table 9. Trial authors' outcome definitions (Continued)

482.9 Bacterial pneumonia NOS		
487.1 Diagnosis of influenza with other respiratory manifestations		
30.	Morton 2004 cluster-RCT Cross-over study USA	Respiratory illnesses defined by symptoms of upper respiratory infections such as nasal congestion, cough, or sore throat, in any combination, with or without fever
31.	Nicholson 2014 cluster-RCT India	Acute respiratory infections Operational definitions for all the illnesses were taken from Black's Medical Dictionary. ARIs defined as "Pneumonia, cough, fever, chest pain and shortness of breath, cold, inflammation of any or all of the airways, that is, nose, sinuses, throat, larynx, trachea and bronchi".
32.	Pandejpong 2012 cluster-RCT Thailand	Influenza-like illness defined if 2 or more symptoms of stuffy nose, cough, fever or chills, sore throat, headache, diarrhoea, presence of hand, foot, or mouth ulcers.
33.	Priest 2014 cluster-RCT New Zealand	Respiratory illness was defined as an episode of illness that included at least 2 of the following caregiver-reported symptoms for 1 day, or 1 of these symptoms for 2 days (but not fever alone): runny nose, stuffy or blocked nose or noisy breathing, cough, fever, sore throat, or sneezing.
34.	Ram 2015 RCT Bangladesh	Influenza-like illness Age-specific definitions of ILI. For individuals ≥ 5 years old, ILI was defined as history of fever with cough or sore throat. For children < 5 years old, ILI was defined as fever; study authors used this relatively liberal case definition in order to include influenza cases with atypical presentations in children. Laboratory-confirmed influenza infection Oropharyngeal swabs from index case patients for laboratory testing for influenza. All swabs were tested by PCR for influenza A and B, with further subtyping of influenza A isolates.
35.	Roberts 2000 cluster-RCT Australia	The symptoms of acute upper respiratory illness elicited from parents were: a runny nose, a blocked nose, and cough. Study authors used a definition of colds based on a community intervention trial of virucidal impregnated tissues. A cold was defined as either 2 symptoms for 1 day or 1 of the respiratory symptoms for at least 2 consecutive days, but not including 2 consecutive days of cough alone. Study authors defined a new episode of a cold as the occurrence of respiratory symptoms after a period of 3 symptom-free days.
36.	Sandora 2005 cluster-RCT USA	The overall rates of secondary respiratory and GI illness. Respiratory illness was defined as 2 of the following symptoms for 1 day or 1 of the symptoms for 2 consecutive days: (1) runny nose; (2) stuffy or blocked nose or noisy breathing; (3) cough; (4) fever, feels hot, or has chills; (5) sore throat; and (6) sneezing. An illness was considered new or separate when a period of at least 2 symptom-free days had elapsed since the previous illness. An illness was defined as a secondary case when it began 2 to 7 days after the onset of the same illness type (respiratory or GI) in another household member.

Table 9. Trial authors' outcome definitions (Continued)

37.	Savolainen-Kopra 2012 cluster-RCT Finland	Nasal and pharyngeal stick samples from participants with respiratory symptoms
38.	Simmerman 2011 cluster-RCT Thailand	Influenza-like illness defined by WHO as fever plus cough or sore throat, based on self-reported symptoms. Laboratory-confirmed secondary influenza virus infections amongst household members described as the secondary attack rate. The secondary influenza virus infection was defined as a positive rRT-PCR result on days 3 or 7 or a four-fold rise in influenza HI antibody titres with the virus type and subtype matching the index case.
39.	Stebbins 2011 cluster-RCT USA	The primary outcome was an absence episode associated with an influenza-like illness that was subsequently laboratory confirmed as influenza A or B. The following CDC definition for ILI was used: fever $\geq 38^{\circ}\text{C}$ with sore throat or cough.
40.	Talaat 2011 cluster-RCT Egypt	Nasal swab for QuickVue test for influenza A and B viruses. Influenza-like illness (defined as fever $> 38^{\circ}\text{C}$ and either cough or sore throat).
41.	Temime 2018 cluster-RCT France	ARIs were defined as the combination of at least 1 respiratory symptom and 1 symptom of systemic infection.
42.	Turner 2004b RCT Canada	Virologic assays
43.	Turner 2012 RCT USA	Laboratory-confirmed rhinovirus infection by PCR assay. Common cold illness was defined as the presence of any of the symptoms of nasal obstruction, rhinorrhoea, sore throat, or cough on at least 3 consecutive days. Illnesses separated by at least 3 symptom-free days were considered as separate illnesses.
44.	Yeung 2011 cluster-RCT Hong Kong	Pneumonia
45.	Zomer 2015 cluster-RCT Netherlands	Incidence of gastrointestinal and respiratory infections in children monitored by parents. The common cold was defined as a blocked or runny nose with at least 1 of the following symptoms: coughing, sneezing, fever, sore throat, or earache.
Hand hygiene and masks (n = 6)		
46.	Aelami 2015 (conference abstract)	Influenza-like illness was defined as the presence of at least 2 of the following during their stay: fever, cough, and sore throat.

Table 9. Trial authors' outcome definitions (Continued)

	RCT	Saudi Arabia	Safety: no outcomes on harms planned or reported.
47.	Aiello 2010	cluster-RCT USA	Influenza-like illness case definition (presence of cough and at least 1 constitutional symptom (fever/feverishness, chills, or body aches). Safety: no outcomes on harms planned or reported.
48.	Cowling 2009	cluster-RCT Hong Kong	2 clinical definitions of influenza. First definition was at least 2 of the following signs and symptoms: temperature 37.8 °C or greater, cough, headache, sore throat, and myalgia. The second was temperature 37.8 °C or greater plus cough or sore throat. Safety: no outcomes on harms planned or reported.
49.	Larson 2010	cluster-RCT USA	Study goals: rates of symptoms and secondary transmission of URIs, incidence of virologically confirmed influenza, knowledge of prevention and treatment strategies for influenza and URIs, and rates of influenza vaccination. <ol style="list-style-type: none"> 1. Laboratory-confirmed influenza: nasal swabs to test for influenza types A and B as well as other common respiratory viruses by rapid culture (R-Mix, Diagnostic Hybrids, Inc., Athens, OH, USA). PCR and subtyping of the samples was done during the second half of the second year of the study. 2. Influenza-like illness: CDC definition of ILI from the Sentinel Physicians' Network was used to determine when masks should be worn: "temperature of $\geq 37.8^{\circ}\text{C}$ and cough and/or sore throat in the absence of a known cause other than influenza". 3. Episodes of URI = upper respiratory infection: not clear, no explicitly stated definition, reported that the most commonly reported URI symptoms are cough or rhinorrhoea. Safety: no outcomes on harms planned or reported.
50.	Simmerman 2011	cluster-RCT Thailand	Laboratory-confirmed secondary influenza virus infections amongst household members described as the secondary attack rate. The secondary influenza virus infection was defined as a positive rRT-PCR result on days 3 or 7 or a four-fold rise in influenza HI antibody titres with the virus type and subtype matching the index case. Influenza-like illness defined by WHO as fever plus cough or sore throat, based on self-reported symptoms. Safety: no outcomes on harms planned or reported.
51.	Suess 2012	cluster-RCT Germany	Quantitative RT-PCR for samples of nasal wash. Influenza virus infection as a laboratory-confirmed influenza infection in a household member who developed fever ($> 38.0^{\circ}\text{C}$), cough, or sore throat during the observation period. Also secondary outcome measure of the occurrence of ILI as defined by WHO as fever plus cough or sore throat. Safety: the study reported that the majority of participants (107/172, 62%) did not report any problems with mask wearing. This proportion was significantly higher in the group of adults (71/100, 71%) compared to the group of children (36/72, 50%) ($P = 0.005$). The main problem stated by participants (adults and children) was "heat/humidity" (18/34, 53% of children; 10/29, 35% of adults) ($P = 0.1$), followed by "pain" and "shortness of breath" when wearing a face mask.

Surface/object disinfection (with or without hand hygiene)(n = 8)

Table 9. Trial authors' outcome definitions (Continued)

52.	Ban 2015 cluster-RCT China	Acute respiratory illness classified as the appearance of 2 or more of the following symptoms: fever, cough and expectoration, runny nose and nasal congestion.
53.	Carabin 1999 cluster-RCT Canada	The presence of nasal discharge (runny nose) accompanied by 1 or several of the following symptoms: fever, sneezing, cough, sore throat, ear pain, malaise, irritability. A URTI was defined as a cold for 2 consecutive days.
54.	Chard 2019 cluster-RCT Laos	Pupils were considered to have symptoms of respiratory infection if they reported cough, runny nose, stuffy nose, or sore throat.
55.	Ibfelt 2015 cluster-RCT Denmark	Laboratory confirmation of 16 respiratory viruses: influenza A; influenza B; coronavirus NL63229E, OC43 and HKU1; parainfluenza virus 1, 2, 3, and 4; rhinovirus; RSV A/B; adenovirus; enterovirus; parechovirus; and bocavirus using quantitative PCR
56.	Kotch 1994 RCT USA	Respiratory symptoms include coughing, runny nose, wheezing or rattling in the chest, sore throat, or earache.
57.	McConeghy 2017 RCT USA	Classified infections as lower respiratory tract infections (i.e. pneumonia, bronchitis, or chronic obstructive pulmonary disease exacerbation) or other.
58.	Sandora 2008 cluster-RCT USA	RI was defined as an acute illness that included > 1 of the following symptoms: runny nose, stuffy or blocked nose, cough, fever or chills, sore throat, or sneezing.
59.	White 2001 DB-RCT USA	RI was defined as: cough, sneezing, sinus trouble, bronchitis, fever alone, pink-eye, headache, mononucleosis, and acute exacerbation of asthma.
Other (miscellaneous) interventions (n = 4)		
60.	Hartinger 2016 cluster-RCT Peru	ARI was defined as a child presenting cough or difficulty breathing, or both. ALRI was defined as a child presenting cough or difficulty breathing, with a raised respiratory rate > 50 per minute in children aged 6 to 11 months and > 40 per minute in children aged > 12 months on 2 consecutive measurements. An episode was defined as beginning on the first day of cough or difficulty breathing and ending with the last day of the same combination, followed by at least 7 days without those symptoms.
61.	Huda 2012 cluster-RCT Bangladesh	Study authors classified acute respiratory illness as having cough and fever or difficulty breathing and fever within 48 h prior to interview.

Table 9. Trial authors' outcome definitions (Continued)

62.	Najnin 2019 cluster-RCT Bangladesh	Classified participants as having respiratory illness if they reported having fever plus either cough or nasal congestion or fever plus breathing difficult.
63.	Satomura 2005 RCT Japan	Upper respiratory tract infection defined as all of the following conditions: <ol style="list-style-type: none"> 1. both nasal and pharyngeal symptoms; 2. severity of at least 1 symptom increased by 2 grades or more; and 3. worsening of a symptom of 1 increment or more for > 3 days. <p>Because of the difference in the mode of transmission, study authors excluded influenza-like diseases featured by moderate or severe fever; anti-influenza vaccination in the pre-season and arthralgia, and treated them separately. The incidence was determined by 1 study physician who was blinded to group assignment.</p>
Virucidal tissues (n = 2)		
64.	Farr 1988a cluster-RCT USA trial 1 and trial 2	RI defined as: occurrence of at least 2 respiratory symptoms on the same day or the occurrence of a single respiratory symptom on 2 consecutive days (except for sneezing). The respiratory symptoms were as follows: sneezing, nasal congestion, nasal discharge, sore throat, scratchy throat, hoarseness, coughing, malaise, headache, feverishness, chilliness and myalgia.
65.	Longini 1988 DB-PC RCT USA	Respiratory illness defined as 1 or more of the following symptoms occurring during the course of acute episode: coryza, sore throat or hoarseness, earache, cough, pain on respiration, wheezy breathing or phlegm from the chest.

ALRI: acute lower respiratory infection

ARIs: acute respiratory infections

CDC: Centers for Disease Control and Prevention

CI: confidence interval

cluster-RCT: cluster-randomised controlled trial

CRI: clinical respiratory illness

DB-PC: double-blind, placebo-controlled

DB-RCT: double-blind randomised controlled trial

GI: gastrointestinal

HCW: healthcare workers

HI: haemagglutinin

hMPV: human metapneumo virus

ICD-9: International Classification of Disease, 9th Revision, Clinical Modification

ICD-10: International Classification of Disease, 10th Revision, Clinical Modification

ILI: influenza-like illness

NAT: nucleic acid testing

NOS: not otherwise specified

NTS: nasal and throat swab

PCR: polymerase chain reaction

PIV: parainfluenza virus

POCT: point-of-care testing

RCT: randomised controlled trial

RI: respiratory infection

RR: risk ratio

rRT-PCR: real-time reverse transcriptase polymerase chain reaction

RSV: respiratory syncytial virus

RTI: respiratory tract infection

RT-PCR: reverse transcriptase polymerase chain reaction

SAR: secondary attack ratios

SD: standard deviation
 S/S: signs and symptoms
 URI: upper respiratory infection
 URTI: upper respiratory tract infection
 WHO: World Health Organization

APPENDICES

Appendix 1. Cochrane Central Register of Controlled Trials (CENTRAL) search string

([mh "Influenza, Human"] OR [mh "Influenzavirus A"] OR [mh "Influenzavirus B"] OR [mh "Influenzavirus C"] OR Influenza:ti,ab OR [mh "Respiratory Tract Diseases"] OR Influenzas:ti,ab OR "Influenza-like":ti,ab OR ILI:ti,ab OR Flu:ti,ab OR Flus:ti,ab OR [mh ^"Common Cold"] OR "common cold":ti,ab OR colds:ti,ab OR coryza:ti,ab OR [mh coronavirus] OR [mh "sars virus"] OR coronavirus:ti,ab OR Coronaviruses:ti,ab OR [mh "coronavirus infections"] OR [mh "severe acute respiratory syndrome"] OR "severe acute respiratory syndrome":ti,ab OR "severe acute respiratory syndromes":ti,ab OR sars:ti,ab OR [mh "respiratory syncytial viruses"] OR [mh "respiratory syncytial virus, human"] OR [mh "Respiratory Syncytial Virus Infections"] OR "respiratory syncytial virus":ti,ab OR "respiratory syncytial viruses":ti,ab OR rsv:ti,ab OR parainfluenza:ti,ab OR "Respiratory illness":ti,ab OR ((Transmission) AND (Coughing OR Sneezing)) OR ((respiratory:ti,ab AND Tract) AND (infection:ti,ab OR Infections:ti,ab OR illness:ti,ab)))

AND

([mh "Hand Hygiene"] OR handwashing:ti,ab OR "hand-washing":ti,ab OR ((Hand:ti,ab OR Alcohol:ti,ab) AND (wash:ti,ab OR Washing:ti,ab OR Cleansing:ti,ab OR Rinses:ti,ab OR hygiene:ti,ab OR rub:ti,ab OR Rubbing:ti,ab OR sanitizer:ti,ab OR sanitiser:ti,ab OR cleanser:ti,ab OR disinfected:ti,ab OR Disinfectant:ti,ab OR Disinfect:ti,ab OR antiseptic:ti,ab OR virucid:ti,ab)) OR [mh "gloves, protective"] OR Glove:ti,ab OR Gloves:ti,ab OR [mh Masks] OR [mh "respiratory protective devices"] OR facemask:ti,ab OR Facemasks:ti,ab OR mask:ti,ab OR Masks:ti,ab OR OR respirator:ti,ab OR respirators:ti,ab OR [mh ^"Protective Clothing"] OR [mh "Protective Devices"] OR "patient isolation":ti,ab OR ((school:ti,ab OR Schools:ti,ab) AND (Closure:ti,ab OR Closures:ti,ab OR Closed:ti,ab)) OR [mh Quarantine] OR quarantine:ti,ab OR "Hygiene intervention":ti,ab OR [mh Mouthwashes] OR gargling:ti,ab OR "nasal tissues":ti,ab OR [mh "Eye Protective Devices"] OR Glasses:ti,ab OR Goggle:ti,ab OR "Eye protection":ti,ab OR Faceshield:ti,ab OR Faceshields:ti,ab OR Goggles:ti,ab OR "Face shield":ti,ab OR "Face shields":ti,ab OR Visors:ti,ab)

AND

([mh "Communicable Disease Control"] OR [mh "Disease Outbreaks"] OR [mh "Disease Transmission, Infectious"] OR [mh "Infection Control"] OR "Communicable Disease Control":ti,ab OR "Secondary transmission":ti,ab OR ((Reduced:ti,ab OR Reduce:ti,ab OR Reduction:ti,ab OR Reducing:ti,ab OR Lower:ti,ab) AND (Incidence:ti,ab OR Occurrence:ti,ab OR Transmission:ti,ab OR Secondary:ti,ab)))

Appendix 2. PubMed search string

("Influenza, Human"[Mesh] OR "Influenzavirus A"[Mesh] OR "Influenzavirus B"[Mesh] OR "Influenzavirus C"[Mesh] OR Influenza[tiab] OR "Respiratory Tract Diseases"[Mesh] OR "Bacterial Infections/transmission"[Mesh] OR Influenzas[tiab] OR "Influenza-like"[tiab] OR ILI[tiab] OR Flu[tiab] OR Flus[tiab] OR "Common Cold"[Mesh:NoExp] OR "common cold"[tiab] OR colds[tiab] OR coryza[tiab] OR coronavirus[Mesh] OR "sars virus"[Mesh] OR coronavirus[tiab] OR Coronaviruses[tiab] OR "coronavirus infections"[Mesh] OR "severe acute respiratory syndrome"[Mesh] OR "severe acute respiratory syndrome"[tiab] OR "severe acute respiratory syndromes"[tiab] OR sars[tiab] OR "respiratory syncytial viruses"[Mesh] OR "respiratory syncytial virus, human"[Mesh] OR "Respiratory Syncytial Virus Infections"[Mesh] OR "respiratory syncytial virus"[tiab] OR "respiratory syncytial viruses"[tiab] OR rsv[tiab] OR parainfluenza[tiab] OR "Respiratory illness"[tiab] OR ((Transmission[tiab]) AND (Coughing[tiab] OR Sneezing[tiab])) OR ((respiratory[tiab] AND Tract[tiab]) AND (infection[tiab] OR Infections[tiab] OR illness[tiab])))

AND

("Hand Hygiene"[Mesh] OR handwashing[tiab] OR hand-washing[tiab] OR ((Hand[tiab] OR Alcohol[tiab]) AND (wash[tiab] OR Washing[tiab] OR Cleansing[tiab] OR Rinses[tiab] OR hygiene[tiab] OR rub[tiab] OR Rubbing[tiab] OR sanitizer[tiab] OR sanitiser[tiab] OR cleanser[tiab] OR disinfected[tiab] OR Disinfectant[tiab] OR Disinfect[tiab] OR antiseptic[tiab] OR virucid[tiab])) OR "gloves, protective"[Mesh] OR Glove[tiab] OR Gloves[tiab] OR Masks[Mesh] OR "respiratory protective devices"[Mesh] OR facemask[tiab] OR Facemasks[tiab] OR mask[tiab] OR Masks[tiab] OR respirator[tiab] OR respirators[tiab] OR "Protective Clothing"[Mesh:NoExp] OR "Protective Devices"[Mesh] OR "patient isolation"[tiab] OR ((school[tiab] OR Schools[tiab]) AND (Closure[tiab] OR Closures[tiab] OR Closed[tiab])) OR Quarantine[Mesh] OR quarantine[tiab] OR "Hygiene intervention"[tiab] OR "Mouthwashes"[Mesh] OR gargling[tiab] OR "nasal tissues"[tiab] OR "Eye Protective Devices"[Mesh] OR Glasses[tiab] OR Goggle[tiab] OR "Eye protection"[tiab] OR Faceshield[tiab] OR Faceshields[tiab] OR Goggles[tiab] OR "Face shield"[tiab] OR "Face shields"[tiab] OR Visors[tiab])

AND

("Communicable Disease Control"[Mesh] OR "Disease Outbreaks"[Mesh] OR "Disease Transmission, Infectious"[Mesh] OR "Infection Control"[Mesh] OR Transmission[sh] OR "Prevention and control"[sh] OR "Communicable Disease Control"[tiab] OR "Secondary transmission"[tiab] OR ((Reduced[tiab] OR Reduce[tiab] OR Reduction[tiab] OR Reducing[tiab] OR Lower[tiab]) AND (Incidence[tiab] OR Occurrence[tiab] OR Transmission[tiab] OR Secondary[tiab])))

AND

(Randomized controlled trial[pt] OR controlled clinical trial[pt] OR randomized[tiab] OR randomised[tiab] OR placebo[tiab] OR "drug therapy"[sh] OR randomly[tiab] OR trial[tiab] OR groups[tiab])

NOT

(Animals[Mesh] not (Animals[Mesh] and Humans[Mesh]))

NOT

("Case Reports"[pt] OR Editorial[pt] OR Letter[pt] OR Meta-Analysis[pt] OR "Observational Study"[pt] OR "Systematic Review"[pt] OR "Case Report"[ti] OR "Case series"[ti] OR Meta-Analysis[ti] OR "Meta Analysis"[ti] OR "Systematic Review"[ti])

Appendix 3. Embase (Elsevier) search string

('influenza'/exp OR Influenza:ti,ab OR 'Respiratory Tract Disease'/exp OR Influenzas:ti,ab OR Influenza-like:ti,ab OR ILI:ti,ab OR Flu:ti,ab OR Flus:ti,ab OR 'Common Cold'/de OR "common cold":ti,ab OR colds:ti,ab OR coryza:ti,ab OR 'coronavirus'/exp OR 'SARS coronavirus'/exp OR coronavirus:ti,ab OR Coronaviruses:ti,ab OR 'coronavirus infection'/exp OR 'severe acute respiratory syndrome'/exp OR "severe acute respiratory syndrome":ti,ab OR "severe acute respiratory syndromes":ti,ab OR sars:ti,ab OR 'Pneumovirus'/exp OR 'Human respiratory syncytial virus'/exp OR "respiratory syncytial virus":ti,ab OR "respiratory syncytial viruses":ti,ab OR rsv:ti,ab OR parainfluenza:ti,ab OR "Respiratory illness":ti,ab OR ((Transmission) AND (Coughing OR Sneezing)) OR ((respiratory:ti,ab AND Tract) AND (infection:ti,ab OR Infections:ti,ab OR illness:ti,ab)))

AND

('hand washing'/exp OR handwashing:ti,ab OR hand-washing:ti,ab OR ((Hand:ti,ab OR Alcohol:ti,ab) AND (wash:ti,ab OR Washing:ti,ab OR Cleansing:ti,ab OR Rinses:ti,ab OR hygiene:ti,ab OR rub:ti,ab OR Rubbing:ti,ab OR sanitizer:ti,ab OR sanitiser:ti,ab OR cleanser:ti,ab OR disinfected:ti,ab OR Disinfectant:ti,ab OR Disinfect:ti,ab OR antiseptic:ti,ab OR virucid:ti,ab)) OR 'protective glove'/exp OR Glove:ti,ab OR Gloves:ti,ab OR 'mask'/exp OR 'gas mask'/exp OR facemask:ti,ab OR Facemasks:ti,ab OR mask:ti,ab OR Masks:ti,ab OR respirator:ti,ab OR respirators:ti,ab OR 'protective clothing'/de OR 'protective equipment'/exp OR "patient isolation":ti,ab OR ((school:ti,ab OR Schools:ti,ab) AND (Closure:ti,ab OR Closures:ti,ab OR Closed:ti,ab)) OR 'Quarantine'/exp OR quarantine:ti,ab OR "Hygiene intervention":ti,ab OR 'mouthwash'/exp OR gargling:ti,ab OR "nasal tissues":ti,ab OR 'eye protective device'/exp OR Glasses:ti,ab OR Goggle:ti,ab OR "Eye protection":ti,ab OR Faceshield:ti,ab OR Faceshields:ti,ab OR Goggles:ti,ab OR "Face shield":ti,ab OR "Face shields":ti,ab OR Visors:ti,ab)

AND

('Communicable Disease Control'/exp OR 'epidemic'/exp OR 'disease transmission'/exp OR 'Infection Control'/exp OR "Communicable Disease Control":ti,ab OR "Secondary transmission":ti,ab OR ((Reduced:ti,ab OR Reduce:ti,ab OR Reduction:ti,ab OR Reducing:ti,ab OR Lower:ti,ab) AND (Incidence:ti,ab OR Occurrence:ti,ab OR Transmission:ti,ab OR Secondary:ti,ab)))

AND

(random* OR factorial OR crossover OR placebo OR blind OR blinded OR assign OR assigned OR allocate OR allocated OR 'crossover procedure'/exp OR 'double-blind procedure'/exp OR 'randomized controlled trial'/exp OR 'single-blind procedure'/exp NOT ('animal'/exp NOT ('animal'/exp AND 'human'/exp)))

Appendix 4. CINAHL (EBSCO) search string

((MH "Influenza, Human+") OR (MH "Orthomyxoviridae+") OR TI Influenza OR AB Influenza OR (MH "Respiratory Tract Diseases+") OR TI Influenzas OR AB Influenzas OR TI Influenza-like OR AB Influenza-like OR TI ILI OR AB ILI OR TI Flu OR AB Flu OR TI Flus OR AB Flus OR (MH "Common Cold+") OR TI "common cold" OR AB "common cold" OR TI colds OR AB colds OR TI coryza OR AB coryza OR (MH "coronavirus+") OR (MH "sars virus+") OR TI coronavirus OR AB coronavirus OR TI Coronaviruses OR AB Coronaviruses OR (MH "coronavirus infections+") OR (MH "severe acute respiratory syndrome+") OR TI "severe acute respiratory syndrome" OR AB "severe acute respiratory syndrome" OR TI "severe acute respiratory syndromes" OR AB "severe acute respiratory syndromes" OR TI sars OR AB sars OR (MH "respiratory syncytial viruses+") OR TI "respiratory syncytial virus" OR AB "respiratory syncytial virus" OR TI "respiratory syncytial viruses" OR AB "respiratory syncytial viruses" OR TI rsv OR AB rsv OR TI parainfluenza OR AB parainfluenza OR TI "Respiratory illness" OR AB "Respiratory illness" OR ((Transmission) AND (Coughing OR Sneezing)) OR ((TI respiratory OR AB respiratory AND Tract) AND (TI infection OR AB infection OR TI Infections OR AB Infections OR TI illness OR AB illness)))

AND

((MH "Handwashing+") OR TI handwashing OR AB handwashing OR TI hand-washing OR AB hand-washing OR ((TI Hand OR AB Hand OR TI Alcohol OR AB Alcohol) AND (TI wash OR AB wash OR TI Washing OR AB Washing OR TI Cleansing OR AB Cleansing OR TI Rinses OR AB Rinses OR TI hygiene OR AB hygiene OR TI rub OR AB rub OR TI Rubbing OR AB Rubbing OR TI sanitizer OR AB sanitiser OR TI sanitizer OR AB sanitiser OR TI cleanser OR AB cleanser OR TI disinfected OR AB disinfected OR TI Disinfectant OR AB Disinfectant OR TI Disinfect OR AB Disinfect OR TI antiseptic OR AB antiseptic OR TI virucid OR AB virucid)) OR (MH "gloves+") OR TI Glove OR AB Glove OR Gloves OR (MH "Masks+") OR (MH "respiratory protective devices+") OR TI facemask OR AB facemask OR TI Facemasks OR AB Facemasks OR TI mask OR AB mask OR TI Masks OR AB Masks OR TI respirator OR AB respirator OR TI respirators OR AB respirators OR (MH "Protective Clothing") OR (MH "Protective Devices+") OR TI "patient isolation" OR AB "patient isolation" OR ((TI school OR AB school OR TI Schools OR AB Schools) AND (TI Closure OR AB Closure OR TI Closures OR AB Closures OR TI Closed OR AB Closed)) OR (MH "Quarantine+") OR TI quarantine OR AB quarantine OR TI "Hygiene intervention" OR AB "Hygiene intervention" OR (MH "Mouthwashes+") OR TI gargling OR AB gargling OR TI "nasal tissues" OR AB "nasal tissues" OR (MH "Eye Protective Devices+") OR TI Glasses OR AB Glasses OR TI Goggle OR AB Goggle OR TI "Eye protection" OR AB "Eye protection" OR TI Faceshield OR AB Faceshield OR TI Faceshields OR AB Faceshields OR TI Goggles OR AB Goggles OR TI "Face shield" OR AB "Face shield" OR TI "Face shields" OR AB "Face shields" OR TI Visors OR AB Visors)

AND

((MH "Infection Control+") OR (MH "Disease Outbreaks+") OR (MH "Infection Control+") OR TI "Communicable Disease Control" OR AB "Communicable Disease Control" OR TI "Secondary transmission" OR AB "Secondary transmission" OR ((TI Reduced OR AB Reduced OR TI Reduce OR AB Reduce OR TI Reduction OR AB Reduction OR TI Reducing OR AB Reducing OR TI Lower OR AB Lower) AND (TI Incidence OR AB Incidence OR TI Occurrence OR AB Occurrence OR TI Transmission OR AB Transmission OR TI Secondary OR AB Secondary))) AND ((MH "Clinical Trials+") OR (MH "Quantitative Studies") OR TI placebo* OR AB placebo* OR (MH "Placebos") OR (MH "Random Assignment") OR TI random* OR AB random* OR TI ((singl* or doubl* or tripl* or trebl*) W1 (blind* or mask*)) OR AB ((singl* or doubl* or tripl* or trebl*) W1 (blind* or mask*)) OR TI clinic* trial* OR AB clinic* trial* OR PT clinical trial)

Appendix 5. Previous search strategies (pre-2010)

Details of the 2010 update and the search strategy used in the original review and the 2009 search strategy updates for MEDLINE, CENTRAL, EMBASE and CINAHL

In the 2010 update we searched, as we have done previously, the Cochrane Central Register of Controlled Trials (CENTRAL) 2010, Issue 3, which includes the Acute Respiratory Infections Group's Specialised Register, MEDLINE (April 2009 to October week 2, 2010), EMBASE (April 2009 to October 2010) and CINAHL (January 2009 to October 2010). Details of previous searches are in Appendix 1. In addition, to include more of the literature of low-income countries in this update, we ran searches in LILACS (2008 to October 2010), Indian MEDLARS (2008 to October 2010) and IMSEAR (2008 to October 2010).

We used the following search strategy (updated to include new and emerging respiratory viruses) to search MEDLINE and CENTRAL. We combined the MEDLINE search strategy with the Cochrane Highly Sensitive Search Strategy for identifying randomised trials in MEDLINE: sensitivity- and precision-maximising version (2008 revision) (Ovid format) (Lefebvre 2011). We also included an additional search strategy based on the work of Fraser, Murray and Burr (Fraser 2006) to identify observational studies.

- 1 Influenza, Human/
- 2 exp Influenzavirus A/
- 3 exp Influenzavirus B/
- 4 Influenzavirus C/
- 5 (influenza* or flu).tw.
- 6 Common Cold/
- 7 common cold*.tw.
- 8 Rhinovirus/
- 9 rhinovir*.tw.
- 10 adenoviridae/ or mastadenovirus/ or adenoviruses, human/
- 11 adenoviridae infections/ or adenovirus infections, human/
- 12 adenovir*.tw.
- 13 coronavirus/ or coronavirus 229e, human/ or coronavirus oc43, human/ or infectious bronchitis virus/ or sars virus/
- 14 coronavir*.tw.
- 15 coronavirus infections/ or severe acute respiratory syndrome/
- 16 (severe acute respiratory syndrome* or sars).tw.
- 17 respiratory syncytial viruses/ or respiratory syncytial virus, human/
- 18 Respiratory Syncytial Virus Infections/
- 19 (respiratory syncytial virus* or rsv).tw.
- 20 Pneumovirus Infections/
- 21 parainfluenza virus 1, human/ or parainfluenza virus 3, human/
- 22 parainfluenza virus 2, human/ or parainfluenza virus 4, human/
- 23 (parainfluenza* or para-influenza* or para influenza).tw.
- 24 enterovirus a, human/ or exp enterovirus b, human/ or enterovirus c, human/ or enterovirus d, human/
- 25 Enterovirus Infections/
- 26 enterovir*.tw.
- 27 Human bocavirus/
- 28 bocavirus*.tw.
- 29 Metapneumovirus/
- 30 metapneumovir*.tw.
- 31 Parvovirus B19, Human/
- 32 parvoviridae infections/ or erythema infectiosum/
- 33 parvovirus*.tw.
- 34 Parechovirus/
- 35 parechovirus*.tw.
- 36 acute respiratory tract infection*.tw.
- 37 acute respiratory infection*.tw.
- 38 or/1-37

39 Handwashing/
 40 (handwashing or hand washing or hand-washing).tw.
 41 hand hygiene.tw.
 42 (sanitizer* or sanitiser*).tw.
 43 (cleanser* or disinfectant*).tw.
 44 gloves, protective/ or gloves, surgical/
 45 glov*.tw.
 46 masks/ or respiratory protective devices/
 47 (mask or masks or respirator or respirators).tw.
 48 Protective Clothing/
 49 Protective Devices/
 50 Patient Isolators/
 51 Patient Isolation/
 52 patient isolat*.tw.
 53 (barrier* or curtain* or partition*).tw.
 54 negative pressure room*.tw.
 55 ((reverse barrier or reverse-barrier) adj3 (nurs* or unit or isolation)).tw.
 56 Cross Infection/pc [Prevention & Control]
 57 (cross infection* adj2 prevent*).tw.
 58 Communicable Disease Control/
 59 Infection Control/
 60 (school* adj3 (clos* or dismissal*)).tw.
 61 temporary closur*.tw.
 62 mass gathering*.tw.
 63 (public adj2 (gathering* or event*)).tw.
 64 (bans or banning or banned or ban).tw.
 65 (outbreak adj3 control*).tw.
 66 distancing*.tw.
 67 Quarantine/
 68 quarantine*.tw.
 69 (protective adj2 (cloth* or garment* or device* or equipment)).tw.
 70 ((protective or preventive) adj2 (procedure* or behaviour* or behavior*)).tw.
 71 personal protect*.tw.
 72 (isolation room* or isolation strateg*).tw.
 73 (distance adj2 patient*).tw.
 74 ((spatial or patient) adj separation).tw.
 75 cohorting.tw.
 76 or/39-75
 77 38 and 76
 78 (animals not (animals and humans)).sh.
 79 77 not 78

Ovid MEDLINE

1 Influenza, Human/
 2 exp Influenzavirus A/
 3 exp Influenzavirus B/
 4 Influenzavirus C/
 5 (influenza* or flu).tw.
 6 Common Cold/
 7 common cold*.tw.
 8 Rhinovirus/
 9 rhinovir*.tw.
 10 adenoviridae/ or mastadenovirus/ or adenoviruses, human/
 11 adenoviridae infections/ or adenovirus infections, human/
 12 adenovir*.tw.
 13 coronavirus/ or coronavirus 229e, human/ or coronavirus oc43, human/ or infectious bronchitis virus/ or sars virus/
 14 coronavir*.tw.
 15 coronavirus infections/ or severe acute respiratory syndrome/
 16 (severe acute respiratory syndrome* or sars).tw.
 17 respiratory syncytial viruses/ or respiratory syncytial virus, human/
 18 Respiratory Syncytial Virus Infections/

- 19 (respiratory syncytial virus* or rsv).tw.
- 20 Pneumovirus Infections/
- 21 parainfluenza virus 1, human/ or parainfluenza virus 3, human/
- 22 parainfluenza virus 2, human/ or parainfluenza virus 4, human/
- 23 (parainfluenza* or para-influenza* or para influenza).tw.
- 24 enterovirus a, human/ or exp enterovirus b, human/ or enterovirus c, human/ or enterovirus d, human/
- 25 Enterovirus Infections/
- 26 enterovir*.tw.
- 27 Human bocavirus/
- 28 bocavirus*.tw.
- 29 Metapneumovirus/
- 30 metapneumovir*.tw.
- 31 Parvovirus B19, Human/
- 32 parvoviridae infections/ or erythema infectiosum/
- 33 parvovirus*.tw.
- 34 Parechovirus/
- 35 parechovirus*.tw.
- 36 acute respiratory tract infection*.tw.
- 37 acute respiratory infection*.tw.
- 38 or/1-37
- 39 Handwashing/
- 40 (handwashing or hand washing or hand-washing).tw.
- 41 hand hygiene.tw.
- 42 (sanitizer* or sanitiser*).tw.
- 43 (cleanser* or disinfectant*).tw.
- 44 gloves, protective/ or gloves, surgical/
- 45 glov*.tw.
- 46 masks/ or respiratory protective devices/
- 47 (mask or masks or respirator or respirators).tw.
- 48 Protective Clothing/
- 49 Protective Devices/
- 50 Patient Isolators/
- 51 Patient Isolation/
- 52 patient isolat*.tw.
- 53 (barrier* or curtain* or partition*).tw.
- 54 negative pressure room*.tw.
- 55 ((reverse barrier or reverse-barrier) adj3 (nurs* or unit or isolation)).tw.
- 56 Cross Infection/pc [Prevention & Control]
- 57 (cross infection* adj2 prevent*).tw.
- 58 Communicable Disease Control/
- 59 Infection Control/
- 60 (school* adj3 (clos* or dismissal*)).tw.
- 61 temporary closur*.tw.
- 62 mass gathering*.tw.
- 63 (public adj2 (gathering* or event*)).tw.
- 64 (bans or banning or banned or ban).tw.
- 65 (outbreak adj3 control*).tw.
- 66 distancing*.tw.
- 67 Quarantine/
- 68 quarantine*.tw.
- 69 (protective adj2 (cloth* or garment* or device* or equipment)).tw.
- 70 ((protective or preventive) adj2 (procedure* or behaviour* or behavior*)).tw.
- 71 personal protect*.tw.
- 72 (isolation room* or isolation strateg*).tw.
- 73 (distance adj2 patient*).tw.
- 74 ((spatial or patient) adj separation).tw.
- 75 cohorting.tw.
- 76 or/39-75
- 77 38 and 76
- 78 (animals not (animals and humans)).sh.
- 79 77 not 78

Embase.com search strategy, October 2010

The search strategy was broadened in 2010 to be more inclusive of new and emerging viruses.

```
#3 #1 AND #25899
#2 766172
#2.8 #2.3 NOT #2.7766172
#2.7 #2.4 NOT #2.6
#2.6 #2.4 AND #2.5
#2.5 'human'/de AND [embase]/lim
#2.4 'animal'/de OR 'nonhuman'/de OR 'animal experiment'/de AND [embase]/lim
#2.3 #2.1 OR #2.2
#2.2 random*:ab,ti OR placebo*:ab,ti OR crossover*:ab,ti OR 'cross over':ab,ti OR allocat*:ab,ti OR trial:ti OR (doubl* NEXT/1 blind*):ab,ti
AND [embase]/lim
#2.1 'randomized controlled trial'/exp OR 'single blind procedure'/exp OR 'double blind procedure'/exp OR 'crossover procedure'/exp AND
[embase]/lim

#1 74545
#1.65 #1.28 AND #1.6474545
#1.64 #1.29 OR #1.30 OR #1.31 OR #1.32 OR #1.33 OR #1.34 OR #1.35 OR
#1.36 OR #1.37 OR #1.38 OR #1.39 OR #1.40 OR #1.41 OR #1.42 OR #1.43
OR #1.44 OR #1.45 OR #1.46 OR #1.47 OR #1.48 OR #1.49 OR #1.50 OR
#1.51 OR #1.52 OR #1.53 OR #1.54 OR #1.55 OR #1.56 OR #1.57 OR #1.58
OR #1.59 OR #1.60 OR #1.61 OR #1.62 OR #1.63
#1.63 cohorting:ab,ti OR 'cohort isolation':ab,ti AND [embase]/lim
#1.62 ((spatial OR patient*) NEAR/2 separation):ab,ti AND [embase]/lim
#1.61 (distance NEAR/2 patient*):ab,ti AND [embase]/lim
#1.60 (isolation NEXT/1 (room* OR strateg*)):ab,ti AND [embase]/lim
#1.59 'personal protection':ab,ti AND [embase]/lim
#1.58 ((protective OR preventive) NEAR/2 (procedure* OR behaviour* OR behavior*)):ab,ti AND [embase]/lim
#1.57 (protective NEAR/2 (cloth* OR garment* OR device* OR equipment)):ab,ti AND [embase]/lim
#1.56 quarantin*:ab,ti AND [embase]/lim
#1.55 distancing:ab,ti AND [embase]/lim
#1.54 ((outbreak* OR transmission OR infection*) NEAR/2 control):ab,ti AND [embase]/lim
#1.53 bans:ab,ti OR banning:ab,ti OR banned:ab,ti OR ban:ab,ti AND [embase]/lim
#1.52 (public NEAR/2 (gathering* OR event*)):ab,ti AND [embase]/lim
#1.51 'mass gathering':ab,ti OR 'mass gatherings':ab,ti AND [embase]/lim
#1.50 (temporar* NEAR/2 closur*):ab,ti AND [embase]/lim
#1.49 (school* NEAR/3 (clos* OR dismissal*)):ab,ti AND [embase]/lim
#1.48 'infection control'/de AND [embase]/lim
#1.47 'epidemic'/dm_pc AND [embase]/lim
#1.46 (('cross infection' OR 'cross infections') NEAR/2 prevent*):ab,ti AND [embase]/lim
#1.45 'cross infection'/dm_pc AND [embase]/lim
#1.44 (('reverse barrier' OR 'reverse-barrier') NEAR/3 (nurs* OR unit OR isolat*)):ab,ti AND [embase]/lim
#1.43 'negative pressure room':ab,ti OR 'negative pressure rooms':ab,ti AND [embase]/lim
#1.42 barrier*:ab,ti OR curtain*:ab,ti OR partition*:ab,ti AND [embase]/lim
#1.41 (patient* NEAR/2 isolat*):ab,ti AND [embase]/lim
#1.40 'patient isolator'/de AND [embase]/lim
#1.39 'protective equipment'/de AND [embase]/lim
#1.38 'protective clothing'/de AND [embase]/lim
#1.37 facemask*:ab,ti OR mask:ab,ti OR masks:ab,ti OR goggles:ab,ti
OR respirator*:ab,ti OR respirators:ab,ti AND [embase]/lim
#1.36 'face mask'/exp OR 'mask'/de OR 'surgical mask'/de AND [embase]/lim
#1.35 glov*:ab,ti AND [embase]/lim
#1.34 'surgical glove'/de AND [embase]/lim
#1.33 cleanser*:ab,ti OR disinfect*:ab,ti OR antiseptic*:ab,ti OR virucid*:ab,ti AND [embase]/lim
#1.32 sanitizer*:ab,ti OR sanitiser*:ab,ti AND [embase]/lim
#1.31 (alcohol NEAR/2 rub*):ab,ti AND [embase]/lim
#1.30 handwash*:ab,ti OR (hand* NEAR/2 (wash* OR cleans* OR hygiene)):ab,ti AND [embase]/lim
#1.29 'hand washing'/de AND [embase]/lim
#1.28 #1.1 OR #1.2 OR #1.3 OR #1.4 OR #1.5 OR #1.6 OR #1.7 OR #1.8 OR #1.9 OR #1.10 OR #1.11 OR #1.12 OR #1.13 OR #1.14 OR #1.15 OR
#1.16 OR #1.17 OR #1.18 OR #1.19 OR #1.20 OR #1.21 OR #1.22 OR #1.23
OR #1.24 OR #1.25 OR #1.26 OR #1.27
```


#1.27 (respiratory NEAR/2 (infect* OR illness* OR virus* OR pathogen* OR acute)):ab,ti AND [embase]/lim
 #1.26 parechovirus*:ab,ti AND [embase]/lim
 #1.25 'parechovirus'/de AND [embase]/lim
 #1.24 parvovirus*:ab,ti AND [embase]/lim
 #1.23 'parvovirus infection'/de OR 'erythema infectiosum'/exp AND [embase]/lim
 #1.22 'parvovirus'/de OR 'human parvovirus b19'/de AND [embase]/lim
 #1.21 'human metapneumovirus'/de OR 'human metapneumovirus infection'/de AND [embase]/lim
 #1.20 'bocavirus'/de OR 'bocavirus infection'/de AND [embase]/lim
 #1.19 enterovir*:ab,ti AND [embase]/lim
 #1.18 'enterovirus infection'/de OR 'coxsackie virus infection'/de OR 'echovirus infection'/de AND [embase]/lim
 #1.17 'enterovirus'/de OR 'coxsackie virus'/exp OR 'echo virus'/de AND [embase]/lim
 #1.16 parainfluenza:ab,ti OR 'para influenza':ab,ti OR 'para-influenza':ab,ti AND [embase]/lim
 #1.15 'parainfluenza virus'/exp AND [embase]/lim
 #1.14 'pneumovirus infection'/de AND [embase]/lim
 #1.13 'respiratory syncytial virus':ab,ti OR 'respiratory syncytial viruses':ab,ti OR rsv:ab,ti AND [embase]/lim
 #1.12 'respiratory syncytial pneumovirus'/de OR 'respiratory syncytial virus infection'/exp AND [embase]/lim
 #1.11 coronavir*:ab,ti OR sars:ab,ti OR 'severe acute respiratory syndrome':ab,ti AND [embase]/lim
 #1.10 'coronavirus infection'/de OR 'severe acute respiratory syndrome'/de AND [embase]/lim
 #1.9 'coronavirus'/de OR 'human coronavirus n163'/de OR 'sars coronavirus'/de OR 'transmissible gastroenteritis virus'/de
 #1.8 adenovir*:ab,ti AND [embase]/lim
 #1.7 'adenovirus infection'/de OR 'human adenovirus infection'/de OR 'human adenovirus'/exp AND [embase]/lim
 #1.6 rhinovir*:ab,ti AND [embase]/lim
 #1.5 'rhinovirus infection'/de OR 'human rhinovirus'/de AND [embase]/lim
 #1.4 'common cold':ab,ti OR 'common colds':ab,ti OR coryza:ab,ti OR colds:ab,ti AND [embase]/lim
 #1.3 'common cold'/de OR 'common cold symptom'/de AND [embase]/lim
 #1.2 influenza*:ab,ti OR flu:ab,ti AND [embase]/lim
 #1.1 'influenza'/exp AND [embase]/lim

CINAHL (EBSCO) search strategy, October 2010

The search strategy was broadened in 2010 to be more inclusive of new and emerging viruses.

S54 S32 and S53
 S53 S44 or S52
 S52 S45 or S46 or S47 or S48 or S49 or S50 or S51
 S51 TI observational stud* or AB observational stud*
 S50 TI cohort stud* or AB cohort stud*
 S49 (MH "Cross Sectional Studies")
 S48 (MH "Nonconcurrent Prospective Studies")
 S47 (MH "Correlational Studies")
 S46 (MH "Case Control Studies+")
 S45 (MH "Prospective Studies")
 S44 S33 or S34 or S35 or S36 or S37 or S38 or S39 or S40 or S41 or S42 or S43
 S43 TI allocat* N1 random* or AB allocat* N1 random*
 S42 (MH "Quantitative Studies")
 S41 TI placebo* or AB placebo*
 S40 (MH "Placebos")
 S39 TI random* allocation* or AB random* allocation*
 S38 (MH "Random Assignment")
 S37 TI (randomised control* trial* or randomized control* trial*) or AB (randomised control* trial* or randomized control* trial*)
 S36 TI ((singl* W1 blind*) or (singl* W1 mask*) or (doubl* W1 blind*) or (doubl* W1 mask*) or (trebl* W1 blind*) or (trebl* W1 mask*) or (tripl* W1 blind*) or (tripl* W1 mask*)) or AB ((singl* W1 blind*) or (singl* W1 mask*) or (doubl* W1 blind*) or (doubl* W1 mask*) or (trebl* W1 blind*) or (trebl* W1 mask*) or (tripl* W1 blind*) or (tripl* W1 mask*))
 S35 TI clinic* W1 trial* or AB clinic* W1 trial*
 S34 PT clinical trial
 S33 (MH "Clinical Trials+")
 S32 S15 and S31
 S31 S16 or S17 or S18 or S19 or S20 or S21 or S22 or S23 or S24 or S25 or S26 or S27 or S28 or S29 or S30
 S30 TI (bans or banning or banned or ban or "outbreak control" or "outbreak controls" or distancing* or quarantine* or "protective clothing" or "protective garment" or "protective garments" or "protective gown" or "protective gowns" or "protective device" or "protective devices" or "protective equipment" or "protective behaviour" or "protective behavior" or "protective behaviours" or "protective behaviors" or "protective procedure" or "protective procedures" or "preventive behaviours" or "preventive behaviour" or "preventive behavior" or "preventive behaviors" or "preventive procedure" or "preventive procedures" or "personal protective" or

"isolation room" or "isolation rooms" or "isolation strategy" or "isolation strategies" or "patient distance" or "patient distancing" or "patient separation" or "spatial separation") or AB (handwashing or "hand washing" or hand-washing or "hand hygiene" or sanitizer or sanitiser or cleanser* or disinfectant* or glov* or mask or masks or respirator or respirators or "patient isolation" or "patient isolators" or barrier* or curtain* or partition* or "negative pressure room" or "negative pressure rooms" or "reverse barrier nursing" or "reverse barrier unit" or "reverse barrier isolation" or "cross infection" or "infection control" or "disease control" or "school closure" or "school closures" or "school dismissal" or "school dismissals" or "temporary closure" or "temporary closures" or "mass gathering" or "mass gatherings" or "public gathering" or "public gatherings" or "public event" or "public events")

S29 TI (handwashing or "hand washing" or hand-washing or "hand hygiene" or sanitizer or sanitiser or cleanser* or disinfectant* or glov* or mask or masks or respirator or respirators or "patient isolation" or "patient isolators" or barrier* or curtain* or partition* or "negative pressure room" or "negative pressure rooms" or "reverse barrier nursing" or "reverse barrier unit" or "reverse barrier isolation" or "cross infection" or "infection control" or "disease control" or "school closure" or "school closures" or "school dismissal" or "school dismissals" or "temporary closure" or "temporary closures" or "mass gathering" or "mass gatherings" or "public gathering" or "public gatherings" or "public event" or "public events") or AB (handwashing or "hand washing" or hand-washing or "hand hygiene" or sanitizer or sanitiser or cleanser* or disinfectant* or glov* or mask or masks or respirator or respirators or "patient isolation" or "patient isolators" or barrier* or curtain* or partition* or "negative pressure room" or "negative pressure rooms" or "reverse barrier nursing" or "reverse barrier unit" or "reverse barrier isolation" or "cross infection" or "infection control" or "disease control" or "school closure" or "school closures" or "school dismissal" or "school dismissals" or "temporary closure" or "temporary closures" or "mass gathering" or "mass gatherings" or "public gathering" or "public gatherings" or "public event" or "public events")

S28 (MH "Sterilization and Disinfection")

S27 (MH "Quarantine")

S26 (MH "Area Restriction (Iowa NIC)") OR (MH "Infection Protection (IowaNIC)")

S25 (MH "Infection Control")

S24 (MH "Cross Infection/PC")

S23 (MH "Isolation, Reverse")

S22 (MH "Patient Isolation")

S21 (MH "Protective Devices")

S20 (MH "Protective Clothing")

S19 (MH "Respiratory Protective Devices")

S18 (MH "Masks")

S17 (MH "Gloves")

S16 (MH "Handwashing+")

S15 S1 or S2 or S3 or S4 or S5 or S6 or S7 or S8 or S9 or S10 or S11 or S12 or S13 or S14

S14 TI ("acute respiratory tract infection" or "acute respiratory tract infections" or "acute respiratory infection" or "acute respiratory infections") or AB (influenza* or flu or "common cold" or "common colds" or rhinovir* or adenovir* or coronavir* or sars or "severe acute respiratory syndrome" or "respiratory syncytial virus" or "respiratory syncytial viruses" or rsv or pneumovir* or parainfluenza* or "para influenza" or para-influenza or enterovir* or bocavir* or metapneumovir* or parvovir* or parechovir*)

S13 TI (influenza* or flu or "common cold" or "common colds" or rhinovir* or adenovir* or coronavir* or sars or "severe acute respiratory syndrome" or "respiratory syncytial virus" or "respiratory syncytial viruses" or rsv or pneumovir* or parainfluenza* or "para influenza" or para-influenza or enterovir* or bocavir* or metapneumovir* or parvovir* or parechovir*) or AB (influenza* or flu or "common cold" or "common colds" or rhinovir* or adenovir* or coronavir* or sars or "severe acute respiratory syndrome" or "respiratory syncytial virus" or "respiratory syncytial viruses" or rsv or pneumovir* or parainfluenza* or "para influenza" or para-influenza or enterovir* or bocavir* or metapneumovir* or parvovir* or parechovir*)

S12 (MH "Respiratory Tract Infections+")

S11 (MH "Parvovirus Infections+")

S10 (MH "Enterovirus Infections+")

S9 (MH "Enteroviruses+")

S8 (MH "Respiratory Syncytial Virus Infections")

S7 (MH "Respiratory Syncytial Viruses")

S6 (MH "SARS Virus")

S5 (MH "Severe Acute Respiratory Syndrome")

S4 (MH "Coronavirus Infections+")

S3 (MH "Coronavirus+") OR (MH "Coronavirus Infections")

S2 (MH "Common Cold")

S1 (MH "Influenza+") OR (MH "Influenza A H5N1") OR (MH "Influenza A

LILACS (Latin America and Caribbean) search strategy

(mh:"Influenza, Human" OR "Gripe Humana" OR "Influenza Humana" OR influenza* OR flu OR gripe OR gripe OR mh:"Influenzavirus A" OR mh:b04.820.545.405* OR mh:b04.909.777.545.405* OR mh:"Influenzavirus B" OR mh:b04.820.545.407* OR mh:b04.909.777.545.407* OR "influenzavirus B" OR mh:"Influenzavirus C" OR "Influenzavirus C" OR mh:"Common Cold" OR "common cold" OR "common colds" OR "Resfriado Común" OR "Resfriado Comum" OR coryza OR "Coriza Aguda") AND (mh:handwashing OR "Lavado de Manos" OR "Lavagem de Mãos" OR "Desinfección de Manos" OR "Desinfecção de Mãos" OR "Higienização de Mãos Pré-Cirúrgica" OR handwash* OR "hand washing" OR "hand hygiene" OR "hand cleaning" OR "hand cleanse" OR "hand cleansing" OR higiene OR sanitizer* OR

sanitiser* OR cleanser* OR disinfect* OR esteriliza* OR desinfectar* OR virucid* OR antiseptic* OR mh:"Gloves, Protective" OR "protective glove" OR "protective gloves" OR "Guantes Protectores" OR "Luvas Protetoras" OR mh:e07.700.600.400* OR mh:j01.637.215.600.400* OR mh:j01.637.708.600.400* OR glove* OR guantes OR luvas OR mh:masks OR mask* OR máscaras OR mascarillas OR facemask* OR goggles OR respirator* OR mh:"Respiratory Protective Devices" OR "Dispositivos de Protección Respiratoria" OR "Dispositivos de Proteção Respiratória" OR mh:"Protective Clothing" OR "Ropa de Protección" OR "Roupa de Proteção" OR mh:e07.700.600* OR mh:j01.637.215.600* OR mh:j01.637.708.600* OR mh:"Protective Devices" OR "Equipos de Seguridad" OR "Equipamentos de Proteção" OR mh:e07.700* OR mh:j01.637.708* OR mh:vs2.006.001.001* OR mh:vs4.002.001.001.007.002.002* OR mh:"Patient Isolation" OR "patient isolation" OR "Aislamiento de Pacientes" OR "Isolamento de Pacientes" OR mh:"Patient Isolators" OR "patient isolators" OR "Aisladores de Pacientes" OR "Isoladores de Pacientes" OR barrier* OR curtain* OR partition* OR barrera OR barreira OR cortina OR tabique OR mh:"Cross Infection" OR "cross infection" OR "Infección Hospitalaria" OR "Infecção Hospitalar" OR "Infecciones en Hospitales" OR "Infecciones Nosocomiales" OR "Infeções Nosocomiais" OR mh:"Infection Control" OR mh:n06.850.780.200.450* OR "Control de Infecciones" OR "Controle de Infecções" OR mh:"Communicable Disease Control" OR "Control de Enfermedades Transmisibles" OR "Controle de Doenças Transmissíveis" OR mh:n06.850.780.200* OR mh:sp8.946.819.811* OR mh:"Disease Outbreaks/prevention & control" OR mh:quarantine OR cuarentena OR quarentena OR "personal protection" OR "isolation room" OR "sala de aislamiento" OR "quarto de isolamento" OR "patient distance" OR "distancia del paciente" OR "spatial separation" OR cohort* OR ban OR bans OR banning OR banned OR prohibici* OR proibi* OR "outbreak control" OR distanc* OR "school closure" OR "school closures" OR "temporary closure" OR "temporary closures" OR "cierre de la escuela" OR "fechamento da escola" OR "public gathering" OR "public gatherings" OR "reunion publica" OR "reverse barrier nursing" OR "reverse barrier unit" OR "reverse barrier isolation" OR "negative pressure room" OR "negative pressure rooms" OR "patient separation") AND db:("LILACS") AND type_of_study:(("clinical_trials" OR "cohort" OR "case_control")

Indian MEDLARS search strategy

(influenza\$ or flu or common cold\$ or rhinovir\$ or coronavir\$ or adenovir\$ or severe acute respiratory syndrome\$ or sars or respiratory syncytial virus\$ or rsv or parainfluenza\$ or enterovir\$ or metapneumovir\$ or parvovir\$ or bocavir\$ or parechovir\$) and (handwashing or hand washing or mask\$ or glove\$ or protect\$ or isolat\$ or barrier\$ or curtain\$ or partition\$ or cross infection\$ or infection control\$ or disease control\$ or school\$ or quarantine\$ or ban\$ or cohort\$ or distanc\$ or spatial separation\$)

IMSEAR (Index Medicus for the South East Asia Region) search strategy

(influenza or flu or common cold or rhinovirus or coronavirus or adenovirus or severe acute respiratory syndrome or sars or respiratory syncytial virus or rsv or parainfluenza or enterovirus or bocavirus or metapneumovirus or parvovirus or parechovirus) and (handwashing or hand washing or hand hygiene or sanitizer or sanitiser or cleanser or disinfectant or gloves or masks or mask or protective clothing or protective devices or patient isolation or barrier or curtain or partition or cross infection or disease control or infection control or school or schools or bans or banning or banned or ban or distancing or quarantine or isolation or spatial separation or cohorting or cohort isolation)

In the first publication of this review we searched the Cochrane Central Register of Controlled Trials (CENTRAL) (The Cochrane Library 2006, issue 4); MEDLINE (1966 to November 2006); OLDMEDLINE (1950 to 1965); EMBASE (1990 to November 2006) and CINAHL (1982 to November 2006). The MEDLINE search terms were modified for OLDMEDLINE, EMBASE and CINAHL.

In this 2009 update we searched the Cochrane Central Register of Controlled Trials (CENTRAL) (The Cochrane Library 2009, issue 2); Ovid MEDLINE (2006 to May Week 1 2009); OLDMEDLINE (1950 to 1965); Ovid EMBASE (2006 to Week 18, 2009) and Ovid CINAHL (2006 to May Week 1 2009).

Ovid MEDLINE

- 1 exp Influenza/
- 2 influenza.tw.
- 3 flu.tw.
- 4 exp Common Cold/
- 5 common cold.tw.
- 6 exp Rhinovirus/
- 7 rhinovirus*.tw.
- 8 exp Adenoviridae/
- 9 adenovirus*.tw.
- 10 exp Coronavirus/
- 11 exp Coronavirus Infections/
- 12 coronavirus*.tw.
- 13 exp Respiratory Syncytial Viruses/
- 14 exp Respiratory Syncytial Virus Infections/
- 15 respiratory syncytial virus*.tw.
- 16 respiratory syncytial virus.tw.
- 17 exp Parainfluenza Virus 1, Human/
- 18 exp Parainfluenza Virus 2, Human/
- 19 exp Parainfluenza Virus 3, Human/
- 20 exp Parainfluenza Virus 4, Human/
- 21 (parainfluenza or para-influenza or para influenza).tw.
- 22 exp Severe Acute Respiratory Syndrome/

Physical interventions to interrupt or reduce the spread of respiratory viruses (Review)

Copyright © 2020 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

23 (severe acute respiratory syndrome or SARS).tw.
 24 acute respiratory infection*.tw.
 25 acute respiratory tract infection*.tw.
 26 or/1-25 (59810)
 27 exp Hand Washing/
 28 (handwashing or hand washing or hand-washing).tw.
 29 hand hygiene.tw.
 30 (sanitizer* or sanitiser*).tw.
 31 (cleanser* or disinfectant*).tw.
 32 exp Gloves, Protective/
 33 exp Gloves, Surgical/
 34 glov*.tw.
 35 exp Masks/
 36 mask*1.tw.
 37 exp Patient Isolators/
 38 exp Patient Isolation/
 39 patient isolat*.tw.
 40 (barrier* or curtain* or partition*).tw.
 41 negative pressure room*.tw.
 42 reverse barrier nursing.tw.
 43 Cross Infection/pc [Prevention]
 44 school closure*.tw.
 45 (clos* adj3 school*).tw.
 46 mass gathering*.tw.
 47 public gathering*.tw.
 48 (ban or bans or banned or banning).tw.
 49 (outbreak* adj3 control*).tw.
 50 distancing.tw.
 51 exp Quarantine/
 52 quarantine*.tw.
 53 or/27-49
 54 26 and 53
 55 (animals not (humans and animals)).sh.
 56 54 not 55

CENTRAL search strategy

#1 MeSH descriptor Influenza, Human explode all trees
 #2 influenza:ti,ab,kw
 #3 flu:ti,ab,kw
 #4 MeSH descriptor Common Cold explode all trees
 #5 "common cold":ti,ab,kw
 #6 MeSH descriptor Rhinovirus explode all trees
 #7 rhinovirus*:ti,ab,kw
 #8 MeSH descriptor Adenoviridae explode all trees
 #9 adenovirus*:ti,ab,kw
 #10 MeSH descriptor Coronavirus explode all trees
 #11 MeSH descriptor Coronavirus Infections explode all trees
 #12 coronavirus*:ti,ab,kw
 #13 MeSH descriptor Respiratory Syncytial Viruses explode all trees
 #14 MeSH descriptor Respiratory Syncytial Virus Infections explode all trees
 #15 respiratory syncytial virus*:ti,ab,kw
 #16 respiratory syncythial virus*:ti,ab,kw
 #17 MeSH descriptor Parainfluenza Virus 1, Human explode all trees
 #18 MeSH descriptor Parainfluenza Virus 2, Human explode all trees
 #19 MeSH descriptor Parainfluenza Virus 3, Human explode all trees
 #20 MeSH descriptor Parainfluenza Virus 4, Human explode all trees
 #21 (parainfluenza or para-influenza or para influenza):ti,ab,kw
 #22 MeSH descriptor Severe Acute Respiratory Syndrome explode all trees
 #23 (severe acute respiratory syndrome or SARS):ti,ab,kw
 #24 acute respiratory infection*:ti,ab,kw
 #25 acute respiratory tract infection*:ti,ab,kw

#26 (#1 OR #2 OR #3 OR #4 OR #5 OR #6 OR #7 OR #8 OR #9 OR #10 OR #11 OR #12 OR #13 OR #14 OR #15 OR #16 OR #17 OR #18 OR #19 OR #20 OR #21 OR #22 OR #23 OR #24 OR #25)
 #27 MeSH descriptor Handwashing explode all trees
 #28 (handwashing or hand washing or hand-washing):ti,ab,kw
 #29 hand hygiene:ti,ab,kw
 #30 (sanitizer* or sanitiser*):ti,ab,kw
 #31 (cleanser* or disinfectant*):ti,ab,kw
 #32 MeSH descriptor Gloves, Protective explode all trees
 #33 MeSH descriptor Gloves, Surgical explode all trees
 #34 glov*:ti,ab,kw
 #35 MeSH descriptor Masks explode all trees
 #36 mask*:ti,ab,kw
 #37 MeSH descriptor Patient Isolators explode all trees
 #38 MeSH descriptor Patient Isolation explode all trees
 #39 (barrier* or curtain* or partition*):ti,ab,kw
 #40 negative NEXT pressure NEXT room*:ti,ab,kw
 #41 "reverse barrier nursing":ti,ab,kw
 #42 MeSH descriptor Cross Infection explode all trees with qualifier: PC
 #43 school NEXT closure*:ti,ab,kw
 #44 (clos* NEAR/3 school*):ti,ab,kw
 #45 mass NEXT gathering*:ti,ab,kw
 #46 public NEXT gathering*:ti,ab,kw
 #47 ("ban" or "bans" or banned or banning):ti,ab,kw
 #48 (outbreak* NEAR/3 control*):ti,ab,kw
 #49 distancing:ti,ab,kw
 #50 MeSH descriptor Quarantine explode all trees
 #51 quarantine*:ti,ab,kw
 #52 (#27 OR #28 OR #29 OR #30 OR #31 OR #32 OR #33 OR #34 OR #35 OR #36 OR #37 OR #38 OR #39 OR #40 OR #41 OR #42 OR #43 OR #44 OR #45 OR #46 OR #47 OR #48 OR #49 OR #50 OR #51)
 #53 (#26 AND #52)

Ovid EMBASE search strategy

1 exp Influenza/
 2 influenza.tw.
 3 flu.tw.
 4 exp Common Cold/
 5 common cold.tw.
 6 exp Human Rhinovirus/
 7 rhinovirus*.tw.
 8 exp Adenovirus/
 9 adenovirus*.tw.
 10 exp Coronavirus/
 11 coronavirus*.tw.
 12 exp Respiratory Syncytial Pneumovirus/
 13 respiratory syncytial virus*.tw.
 14 respiratory syncytial virus.tw.
 15 (parainfluenza or para-influenza or para influenza).tw.
 16 exp Severe Acute Respiratory Syndrome/
 17 (severe acute respiratory syndrome or SARS).tw.
 18 acute respiratory infection*.tw.
 19 acute respiratory tract infection*.tw.
 20 or/1-19
 21 exp Hand Washing/
 22 (handwashing or hand washing or hand-washing).tw.
 23 hand hygiene.tw.
 24 (sanitizer\$ or sanitiser\$).tw.
 25 (cleanser\$ or disinfectant\$).tw.
 26 exp Glove/
 27 exp Surgical Glove/
 28 glov*.tw.
 29 exp Mask/
 30 mask*1.tw.

31 patient isolat*.tw.
 32 (barrier* or curtain* or partition*).tw.
 33 negative pressure room*.tw.
 34 reverse barrier nursing.tw.
 35 Cross Infection/pc [Prevention]
 36 school closure*.tw.
 37 (clos* adj3 school*).tw.
 38 mass gathering*.tw.
 39 public gathering*.tw. (5)
 40 (ban or bans or banned or banning).tw.
 41 (outbreak* adj3 control*).tw.
 42 distancing.tw.
 43 quarantine*.tw.
 44 or/21-43
 45 20 and 44

EBSCO CINAHL search strategy

S26 S10 and S24
 S25 S10 and S24
 S24 S11 or S12 or S13 or S14 or S15 or S16 or S17 or S18 or S19 or S20 or S21 or S22 or 23 or S24
 S23 TI outbreak* N3 control* or AB outbreak* N3 control*
 S22 TI (school closure* or mass gathering* or public gathering* or ban or bans or banned or banning or distancing or quarantine*) or AB (school closure* or mass gathering* or public gathering* or ban or bans or banned or banning or distancing or quarantine*)
 S21 TI (patient isolat* or barrier* or curtain* or partition* or negative pressure room* or reverse barrier nursing) or AB (patient isolat* or barrier* or curtain* or partition* or negative pressure room* or reverse barrier nursing)
 S20 TI (glov* or mask*) or AB (glov* or mask*)
 S19 TI (handwashing or hand washing or hand-washing or hand hygiene) or AB (handwashing or hand washing or hand-washing or hand hygiene)
 S18 (MH "Quarantine")
 S17 (MM "Cross Infection")
 S16 (MH "Isolation, Reverse")
 S15 (MH "Patient Isolation+")
 S14 (MH "Respiratory Protective Devices")
 S13 (MH "Masks")
 S12 (MH "Gloves")
 S11 (MH "Handwashing+")
 S10 S1 or S2 or S3 or S4 or S5 or S6 or S7 or S8 or S9
 S9 TI (influenza or flu or rhinovirus* or adenovirus* or coronavirus* or respiratory syncytial virus* or respiratory syncythal virus* or parainfluenza or para-influenza or para influenza or severe acute respiratory syndrome or SARS or respiratory viral infection* or viral respiratory infection*) or AB (influenza or flu or rhinovirus* or adenovirus* or coronavirus* or respiratory syncytial virus* or respiratory syncythal virus* or parainfluenza or para-influenza or para influenza or severe acute respiratory syndrome or SARS or respiratory viral infection* or viral respiratory infection*)TI (influenza or flu or rhinovirus* or adenovirus* or coronavirus* or respiratory syncytial virus* or respiratory syncythal virus* or parainfluenza or para-influenza or para influenza or severe acute respiratory (syndrome or SARS or respiratory viral infection* or viral respiratory infection*) or AB (influenza or flu or rhinovirus* or adenovirus* or coronavirus* or respiratory syncytial virus* or respiratory syncythal virus* or parainfluenza or para-influenza or para influenza or severe acute respiratory syndrome or SARS or respiratory viral infection* or viral respiratory infection*)
 S8 (MH "SARS Virus")
 S7 (MH "Severe Acute Respiratory Syndrome")
 S6 (MH "Respiratory Syncytial Virus Infections")
 S5 (MH "Respiratory Syncytial Viruses")
 S4 (MH "Coronavirus+")
 S3 (MH "Coronavirus Infections+")
 S2 (MH "Common Cold")
 S1 (MH "Influenza+")

WHAT'S NEW

Date	Event	Description
1 April 2020	New citation required and conclusions have changed	There is now sufficient randomised controlled trial (RCT) evidence to show that hand hygiene is likely to provide a modest-benefit. Uncertainty remains for the other interventions. Further RCT evidence is needed.
1 April 2020	New search has been performed	<p>Searches updated. In this 2020 update we only searched for RCTs and cluster-RCTs. We included 44 new trials (Aelami 2015; Aiello 2012; Alzaher 2018; Arbogast 2016; Azor-Martinez 2016; Azor-Martinez 2018; Ban 2015; Barasheed 2014; Biswas 2019; Canini 2010; Chard 2019; Correa 2012; DiVita 2011; Feldman 2016; Goodall 2014; Hartinger 2016; Hubner 2010; Huda 2012; Ibfelt 2015; Ide 2014; Ide 2016; Little 2015; MacIntyre 2011; MacIntyre 2013; MacIntyre 2015; MacIntyre 2016; McConeghy 2017; Millar 2016; Miyaki 2011; Najnin 2019; Nicholson 2014; Pandejpong 2012; Priest 2014; Radonovich 2019; Ram 2015; Savolainen-Kopra 2012; Simmerman 2011; Stebbins 2011; Suess 2012; Talaat 2011; Temime 2018; Turner 2012; Yeung 2011; Zomer 2015).</p> <p>We excluded 12 new trials (Azor-Martinez 2014; Bowen 2007; Chami 2012; Denbak 2018; Lennell 2008; Nandrup-Bus 2009; Patel 2012; Rosen 2006; Slayton 2016; Stedman-Smith 2015; Uhari 1999; Vessey 2007).</p> <p>We identified 5 new ongoing trials (NCT03454009; NCT04267952; NCT04296643; NCT04337541; Wang 2015) one of which – NCT04337541 – published as this review was going to press.</p> <p>We focused on RCTs and cluster-RCTs only and removed observational studies from this update.</p>

HISTORY

Protocol first published: Issue 4, 2006

Review first published: Issue 4, 2007

Date	Event	Description
22 October 2010	New search has been performed	Searches conducted. We included 7 new trials: 4 randomised controlled trials and 3 non-randomised comparative studies. We excluded 36 new trials.
22 October 2010	New citation required but conclusions have not changed	We updated the review again at the behest of the World Health Organization (WHO). External sources of support amended. External support from the WHO. The WHO interim guidelines document on 'Infection Prevention and Control of Epidemic and Pandemic Prone Acute Respiratory Diseases in Health Care' was published in 2007 to provide infection control guidance to help prevent the transmission of acute respiratory diseases in health care. The update of these guidelines will be evidence-based, and an update of this review was requested to assist in informing the evidence base for the revision of the WHO guidelines. Dr John Conly, Dr Mark Jones, and Sarah Thorning joined the review team.
7 May 2009	New search has been performed	For the 2009 update, we included 3 cluster-randomised controlled trials, Cowling 2009 ; MacIntyre 2009 ; Sandora 2008 , and 1

Date	Event	Description
		<p>individual randomised controlled trial (Satomura 2005, with its linked publication Kitamura 2007). We also included 1 retrospective cohort study (Foo 2006), 1 case-control study (Yu 2007), and 2 prospective cohort studies (Wang 2007; Broderick 2008).</p> <p>The content and conclusions of the 2007 review changed little, but the additional 8 studies add more information and certainty. Our meta-analysis remains unchanged as there were no new studies for pooling.</p>
30 April 2009	New citation required but conclusions have not changed	New author joined the review team.
8 July 2008	Amended	Converted to new review format.
20 August 2007	Amended	Review first published Issue 4, 2007.

CONTRIBUTIONS OF AUTHORS

Tom Jefferson (TOJ), Chris Del Mar (CDM), and Liz Dooley (LD) were responsible for drafting the protocol. TOJ, Eliana Ferroni (EF), Bill Hewak (BH), and Adi Prabhala (AP) extracted study data, and Sree Nair (SN) performed the analyses in the original review.

TOJ, EF, Lubna A Al-Ansary (LA), Ghada A Bawazeer (GB), and CDM adjudicated in data extraction in the 2009 update. Mieke van Driel (MvD) assisted in writing the review, updating with the most recent studies, and additional tables (apart from TiDIER). MvD constructed the summary of results table which was removed in the 2020 review update.

TOJ and John Conly (JMC) extracted data, and CDM checked extractions and arbitrated in the 2010 update. All three review authors checked the search strategy terms. Sarah Thorning (ST) designed and carried out the searches. All 2010 review authors contributed to the final report.

For the 2020 update:

Updated searches: ST

Co-ordinated the update: LD

Updated Background section: CDM

Designed Excel forms for extracting study characteristics and tested their usefulness/applicability: MJ

Screened titles and abstracts, excluded irrelevant citations: MJ

Excluded irrelevant citations based on title and text in the trial registry entry: ST

Excluded irrelevant citations based on titles/abstracts and the full-text articles: GB

Selection of studies: MvD, MJ, GB, JC

Data extraction and management: MJ, EF, LA, GB, TOJ, TH, EB

Assessment of risk of bias in the included studies: TOJ, EB, LA, GB, MJ, EF

Adjudicated data extraction: MJ

Data analysis: MJ, EB

Wrote the update: MJ, TOJ, LD, LA, JMC, EB, MVD, GB, TH, CDM, PG

DECLARATIONS OF INTEREST

Tom Jefferson: Tom Jefferson's full disclosure is available at restoringtrials.org/competing-interests-tom-jefferson/.

Chris B Del Mar: a grant from WHO paid to Bond University and a grant from UK National Institute for Health Research (NIHR) to be paid to Bond University on publication of the review in the Cochrane Library. Funding from National Health and Medical Research Council (NHMRC) for research on antibiotic resistance.

Liz Dooley: a grant from the World Health Organization (WHO) paid to Bond University and a grant from UK NIHR to be paid to Bond University on publication of the review in the Cochrane Library.

Eliana Ferroni: none known.

Lubna A Al-Ansary: none known.

Ghada A Bawazeer: none known.

Mieke L van Driel: Dr van Driel has acted as a consultant for Therapeutic Guidelines Ltd and NPS Medicinewise, for which fees have been paid to her institution.

Mark A Jones: a grant from WHO paid to Bond University and a grant from UK NIHR to be paid to Bond University on publication of the review in the Cochrane Library.

Sarah Thorning: none known.

Elaine M Beller: this review was supported in part by research grants from the NHMRC, Australia, to the Institute for Evidence-Based Healthcare, Bond University.

Justin Clark: none known.

Tammy Hoffmann: a grant from WHO paid to Bond University and a grant from UK NIHR to be paid to Bond University on publication of the review in the Cochrane Library. Funding from NHMRC for research on antibiotic resistance.

Paul Glasziou: none known.

John M Conly: John Conly holds grants from the Canadian Institutes for Health Research on acute and primary care preparedness for COVID-19 in Alberta, Canada and was the primary local Investigator for a *Staphylococcus aureus* vaccine study funded by Pfizer for which all funding was provided only to the University of Calgary. He also received support from the Centers for Disease Control and Prevention (CDC) to attend an Infection Control Think Tank Meeting.

SOURCES OF SUPPORT

Internal sources

- The Cochrane Collaboration Steering Group, UK

External sources

- National Institute of Health Research (NIHR), UK

Competitive grant awarded through The Cochrane Collaboration, 2009

- National Health and Medical Research Council (NHMRC), Australia

Competitive grant to Chris Del Mar and Tom Jefferson, 2009

- Sabbatical year (2010 to 2011) for John Conly while at the World Health Organization in Geneva, Switzerland was supported by the University of Calgary, Calgary, Canada

2020/1011941

- World Health Organization, Geneva, Switzerland

Requested and provided support to The Cochrane Collaboration for the 2011 update

- National Institute of Health Research (NIHR), UK

This study/project is funded by the NIHR Incentive Scheme Award Reference 130721. The views expressed are those of the authors and not necessarily those of the NIHR or the Department of Health and Social Care.

- World Health Organization, Geneva, Switzerland

Provided financial support for the 2020 update of this review. Reference number 2020/1011941

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

We changed the title of the review in 2010 (see [Published notes](#) below).

For this 2020 update, we added one additional outcome: adverse events related to the intervention, and we split the outcomes into primary and secondary outcomes. We also focused only on RCTs and cluster-RCTs and removed observational studies.

NOTES

In Issue 1, 2010, the title of the review was changed from 'Interventions for the interruption or reduction of the spread of respiratory viruses' to 'Physical interventions to interrupt or reduce the spread of respiratory viruses'.

The original review was subsequently published as Jefferson T, Foxlee R, Del Mar C, Dooley L, Ferroni E, Hewak B, Prabhala A, Nair S, Rivetti A. Physical interventions to interrupt or reduce the spread of respiratory viruses: systematic review. *BMJ* 2008;336:77-80 and Jefferson T, Del Mar C, Dooley L, Ferroni E, Al-Ansary LA, Bawazeer GA, van Driel ML, Foxlee R, Rivetti A. [Physical interventions to interrupt or reduce the spread of respiratory viruses: systematic review](#). *BMJ* 2009;339:b3675. DOI: 10.1136/bmj.b3675.

INDEX TERMS**Medical Subject Headings (MeSH)**

Case-Control Studies; Influenza, Human [transmission] [virology]; Randomized Controlled Trials as Topic; Respiratory Tract Infections [*prevention & control] [transmission] [virology]; Virus Diseases [*prevention & control] [transmission]; *Virus Shedding

MeSH check words

Humans