

# Effectiveness of home-based pulmonary rehabilitation: systematic review and meta-analysis

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Home-based pulmonary rehabilitation is as effective as centre-based in improving exercise capacity and quality of life, and is an option for people with COPD whose access to pulmonary rehabilitation centres is difficult. https://bit.ly/39HkMm4

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#### Abstract

*Introduction* Despite proven effectiveness for people with chronic respiratory diseases, practical barriers to attending centre-based pulmonary rehabilitation (centre-PR) limit accessibility. We aimed to review the clinical effectiveness, components and completion rates of home-based pulmonary rehabilitation (home-PR) compared to centre-PR or usual care.

*Methods and analysis* Using Cochrane methodology, we searched (January 1990 to August 2021) six electronic databases using a PICOS (population, intervention, comparison, outcome, study type) search strategy, assessed Cochrane risk of bias, performed meta-analysis and narrative synthesis to answer our objectives and used the Grading of Recommendations, Assessment, Development and Evaluations framework to rate certainty of evidence.

**Results** We identified 16 studies (1800 COPD patients; 11 countries). The effects of home-PR on exercise capacity and/or health-related quality of life (HRQoL) were compared to either centre-PR (n=7) or usual care (n=8); one study used both comparators. Compared to usual care, home-PR significantly improved exercise capacity (standardised mean difference (SMD) 0.88, 95% CI 0.32–1.44; p=0.002) and HRQoL (SMD -0.62, 95% CI -0.88-0.36; p<0.001). Compared to centre-PR, home-PR showed no significant difference in exercise capacity (SMD -0.10, 95% CI -0.25-0.05; p=0.21) or HRQoL (SMD 0.01, 95% CI -0.15-0.17; p=0.87).

**Conclusion** Home-PR is as effective as centre-PR in improving functional exercise capacity and quality of life compared to usual care, and is an option to enable access to pulmonary rehabilitation.

# Introduction

An estimated 545 million people globally are affected by chronic respiratory diseases such as COPD, remodelled asthma, pulmonary impairment after tuberculosis, interstitial lung disease (ILD), bronchiectasis and cystic fibrosis [1]. Chronic respiratory diseases are associated with breathlessness, fatigue and muscle dysfunction, which contribute to reduced physical activity levels and functional exercise capacity [2], and impaired health-related quality of life (HRQoL) [3, 4].





Pulmonary rehabilitation is an individually tailored multifaceted intervention that improves the physical condition and psychological wellbeing of people with chronic respiratory diseases [5–7]. Despite proven effectiveness [8, 9] and guideline recommendations [10, 11], pulmonary rehabilitation is under-utilised.

The reasons for poor attendance and completion rates are multifactorial, but inconvenient timing of programmes and geographical distance to pulmonary rehabilitation centres are commonly identified barriers [12–16]. While pertinent even in high-income countries [17–19], poor transport infrastructure in low- and middle-income countries (LMICs) exacerbates these barriers [20]. Typically, pulmonary rehabilitation is provided in hospital centres (centre-PR) [21], but community-based centres [22], home-based pulmonary rehabilitation (home-PR) with telephone mentoring [23], or telerehabilitation programmes [24], are attracting increasing interest. The ongoing coronavirus disease 2019 pandemic has necessitated remote delivery of the treatment for reasons of infection control [25].

Evidence of the effectiveness of these options varies. A subgroup analysis in a Cochrane review favoured centre-PR [8], while three systematic reviews concluded that home/community-PR could be as effective as centre-PR for people with COPD [26-29]. However, combining home and community services overlooks the distinction between a community-based group supervised in-person by a healthcare professional and a programme delivered to an individual in their own home. In addition, these reviews are limited by disease (COPD only), although there is evidence that pulmonary rehabilitation is of benefit in bronchiectasis and ILD [30-32]. More recently, a Cochrane review concluded that telerehabilitation for people with chronic respiratory diseases, achieved similar effectiveness and safety outcomes to centre-PR [33]. "Telerehabilitation" defines the intervention by the means of communication and the review included pulmonary rehabilitation delivered to individuals or groups (either physical or virtual) in any location. including in the patient's home or at a healthcare centre. In contrast, we defined home-PR as sessions undertaken by individuals by themselves (although a family member may be involved) and typically at home. Apart from baseline and post-PR assessments [32], the patient does not attend a centre (either a hospital centre or a local "satellite" community centre) and is not supervised face-to-face by a healthcare professional (though there may be remote communication from a healthcare professional for some or all of the sessions), is not part of an "in-person" group. In addition, to distinguish from "exercise training programmes" included in some reviews [26, 27, 32, 33], our definition of pulmonary rehabilitation comprised both exercise and at least one nonexercise component.

We aimed to systematically review the literature to assess the effectiveness, completion rates and components used in effective home-PR for people with chronic respiratory diseases. Our hypotheses were that 1) home-PR is superior to usual care, and 2) home-PR is noninferior to centre-PR. In people with chronic respiratory diseases, our objectives were to 1) assess the clinical effectiveness of home-PR compared to centre-PR or usual care at improving health outcomes (*i.e.* exercise capacity (primary outcome), HRQoL (primary outcome), dyspnoea, muscle fatigue, exacerbations and hospitalisations for chronic respiratory disease); 2) describe the components of home-PR that are associated with successful interventions (*e.g.* intensity of exercise, duration of the programme, education and/or other nonexercise components, frequency of supervision, information/resources, involvement of family members); and 3) compare the completion rate (defined as participating in  $\geqslant$ 70% of pulmonary rehabilitation sessions) of home-PR with centre-PR.

# Methods

We followed Cochrane methodology [34], and used Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines [35] to report our review findings. The review is registered at www.crd.york.ac.uk/PROSPERO (CRD42020220137) and the protocol has been published [36].

# Search strategy

We developed a search strategy to identify randomised controlled trials (RCTs) and controlled clinical trials of "chronic respiratory disease" AND "pulmonary rehabilitation" AND "home-PR" from 1990 (when pulmonary rehabilitation was first recommended by global COPD guidelines [37]) to 12 October 2020, without any language restrictions. In addition, we checked reference lists and conducted forward citation on included studies and on Cochrane reviews of pulmonary rehabilitation [8]. We searched MEDLINE, the Cumulative Index to Nursing and Allied Health, Cochrane, Embase, PeDRO, and PsycInfo (see appendix 1 in the supplementary material). Table 1 describes the PICOS (population, intervention, comparison, outcome, study type) search strategy and our definition of home-PR and centre-PR. A pre-publication update was conducted in August 2021 using forward citation of the Cochrane review [8] and all the studies included in this review [38].

#### Selection process

Following the search, all identified records were loaded into EndNote X9 (Clarivate Analytics, Philadelphia, PA, USA) and duplicates were removed. Six trained reviewers (M.N. Uzzaman, T. Jackson, J.P. Engkasan, F.T. Mirza, D. Agarwal, P. Jebaraj) worked in pairs to independently screen titles and

TABLE 1 PIC	COS (population, intervention	, comparison, outcome, study type) table	e for the search strategy
	Description, inclusion	Exclusion criteria	Operational rules
Population	Adults with primary diagnosis of CRDs Age >18 years Comorbidity will not be an exclusion criterion	Pregnant women and paediatric population Rehabilitation provided to predominant condition is nonrespiratory conditions Recovery from acute infections or injury (e.g. immediately post-COVID-19) until the condition has been stable for 6 months Conference abstract Lung cancer Pulmonary hypertension	PR delivered to people with CRDs such as COPD, remodelled asthma, PIAT, bronchiectasis, ILD, CF, stable post-COVID-19 (but excluding post-ICU rehabilitation) will be studied. We will also include PR delivered to people with more than one CRD, or undifferentiated chronic respiratory conditions  Conference abstracts will be excluded, but will prompt a search for a subsequent published paper
Intervention	Home-PR that comprises both exercise and at least one nonexercise component for a duration not less than 4 weeks	Formal hospital or community medical centre-based programmes	Home-PR: the key criterion is that the sessions are undertaken by individuals by themselves (although a family member may be involved) and typically at home. Apart from baseline and post-PR assessments, the patient does not attend a centre (either a hospital centre or a local "satellite" centre) and is not supervised face-to-face by a healthcare professional (although there may be remote communication from a healthcare professional for some or all of the session), and is not part of an "in-person" group Exercise sessions typically include aerobic, endurance, resistance and reconditioning exercises, although local resources and preferences may include other exercise modalities. Nonexercise components commonly include patient education, energy conservation training, smoking cessation, psychological support, self-management skill development or other recognised PR interventions, along with optimisation of pharmacotherapy
Comparison	Either population receiving centre-PR or receiving usual care	No control group	Centre-PR: the key criterion is that the sessions are under direct healthcare professional's supervision. The "centre" can be in a hospital, community setting or remote facility. Centre-based sessions are normally group-based (although it is recognised that this may be modified in the context of a pandemic). Telehealth services where patients attend a supervised satellite centre would be considered as centre-PR Usual care: the standard care received by individuals with CRD in the relevant healthcare system, but excluding the exercise components of PR
Outcomes	Either one of the following outcome measures: HRQoL Functional exercise capacity  ± Additional outcome(s): Uptake of the service, completion rates Assessment of motivation/ other intermediate outcome Activities of daily living Physical activity level Symptom control Psychological status Healthcare burden, e.g. exacerbation rates, hospitalisation, etc. Adverse effect	Not including HRQoL or any measurement of exercise capacity as outcome	Any validated instruments will be considered:  HRQoL: e.g. SGRQ, CRQ, EQ-5D  Functional exercise capacity: e.g. 6MWT, ISWT, ESWT. We will also include step tests and sit-to-stand tests that are sometimes used in home-PR assessments  Symptom control: e.g. mMRC, CCQ, Borg scale  Psychological status: e.g. HADS, PHQ-9, STAI, Beck's inventory tests
Setting	Any settings		Low- or high-resource settings, irrespective of level of economies of the countries
Study designs	RCTs, CCTs	Cohort study, case series, case report	We will exclude studies that do not have a control group. We will consider RCTs to answer all of the three research questions (i.e. effectiveness, components and completion rate of home-PR), and consider CCTs to answer research questions 2 and 3
Language	No language restriction		4

CRD: chronic respiratory disease; COVID-19: coronavirus disease 2019; PR: pulmonary rehabilitation; PIAT: pulmonary impairment after tuberculosis; ILD: interstitial lung disease; CF: cystic fibrosis; ICU: intensive care unit; HRQoL: health-related quality of life; SGRQ: St George's Respiratory Questionnaire; CRQ: Chronic Respiratory Questionnaire; EQ-5D: EuroQol Five Dimension; 6MWT: 6-min walk test; ISWT: incremental shuttle walking test; ESWT: endurance shuttle walking test; mMRC: modified Medical Research Council dyspnoea scale; CCQ: Clinical COPD Questionnaire; HADS: Hospital Anxiety and Depression Scale; PHQ-9: Patient Health Questionnaire-9; STAI: State-Trait Anxiety Inventory; RCT: randomised controlled trial; CCT: clinical controlled trial. Reproduced from [36] with permission.

abstracts, followed by full-text papers using the inclusion and exclusion criteria, defined by our operational rules (table 1). Disagreements were resolved by discussion with the review team (H. Pinnock, R.A. Rabinovich, Su May Liew (University of Malaya, Kuala Lumpur, Malaysia), G.M.M. Habib, N.S. Hanafi and S.C. Chan) as necessary. The process is reported in a PRISMA flow diagram (figure 1).

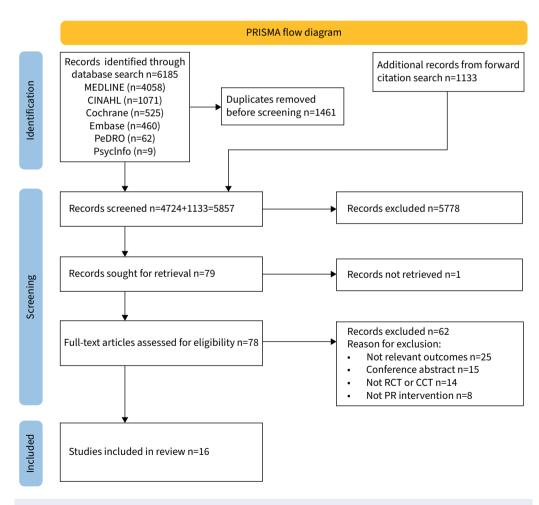
### Outcome measurement

Our primary outcomes were functional exercise capacity and HRQoL: 1) functional exercise capacity measured with any validated tools such as the 6-min walk test (6MWT) [39], Incremental Shuttle Walking Test (ISWT) [40] or Endurance Shuttle Walking Test (ESWT) [41]; 2) HRQoL measured with any validated tools such as the St George's Respiratory Questionnaire (SGRQ) [42], Chronic Respiratory Questionnaire (CRQ) [43], COPD Assessment Test (CAT) [42] or Short Form (SF-36 or SF-12).

We were interested in between-group differences at the post-pulmonary rehabilitation assessment (or first follow-up assessment if post-pulmonary rehabilitation assessment was not reported). Where multiple assessment tools for an outcome (exercise capacity or HRQoL) were reported, we used the most frequently reported measure (*e.g.* 6MWT, SGRQ) in the meta-analysis.

## Data extraction and risk-of-bias assessment

Data extraction was carried out by six reviewers (M.N. Uzzaman, T. Jackson, J.P. Engkasan, F.T. Mirza, D. Agarwal, P. Jebaraj) independently working in pairs, and checked by a third review author (H. Pinnock, R.A. Rabinovich). Data were extracted using a data extraction form in a Microsoft Excel spreadsheet and based on Cochrane Effective Practice and Organisation of Care guidance [44]. The following data were



**FIGURE 1** Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) flow diagram. CINAHL: Cumulative Index to Nursing and Allied Health; RCT: randomised controlled trial; CCT: clinical controlled trial; PR: pulmonary rehabilitation.

extracted from included studies: methods (study location, study design, duration of the intervention, duration of each pulmonary rehabilitation session, mode of supervision, follow-up period (if any)); participant characteristics (number, mean age, gender, diagnosis, severity of the condition); interventions (intervention, comparison); outcomes (primary and secondary outcomes specified and collected (at baseline and at the time of intervention completion) and follow-up measures at any other time point reported).

One review author (M.N. Uzzaman) transferred data into the Review Manager software (RevMan 2020, version 5.4.1) for conducting meta-analysis and another review author (R.A. Rabinovich) checked data accuracy. The six reviewers (M.N. Uzzaman, T. Jackson, J.P. Engkasan, F.T. Mirza, D. Agarwal, P. Jebaraj) also independently assessed methodological quality of all included studies using the Cochrane risk of bias tool for RCTs [45]. Discrepancies were resolved by discussion within the team. We assessed the risk of bias in the following domains: random sequence generation, allocation concealment, blinding of participants and personnel, blinding of outcome assessment, incomplete outcome data, selective outcome reporting, other sources of bias and overall risk of bias. We assessed each potential source of bias as high, low or unclear and summarised the "risk of bias" judgements across different studies for each of the domains in a risk of bias table. We contacted the author(s) of included studies to obtain any incomplete or missing data, but did not perform any statistical calculation for missing data to include in the meta-analysis.

# Heterogeneity and reporting bias

We assessed heterogeneity [46], and explored clinical and methodological reasons for substantial heterogeneity ( $I^2$  statistic >50%) in our primary outcome as defined in our *a priori* subgroups [34], and a sensitivity analyses for the effect of risk of bias. We were not able to pool >10 studies and therefore did not create a funnel plot to test for publication bias [47].

# Subgroups and sensitivity analyses

Our *a priori* subgroups were high-/low-income countries, chronic respiratory disease diagnosis, severity, intensity of intervention and arrangements for supervision of the home-PR programme [36]. We undertook a sensitivity analysis of our primary outcomes for the home-PR *versus* centre-PR comparison excluding studies at high risk of bias (there were too few studies for a sensitivity analysis of the home-PR *versus* usual care analysis).

# Data analysis to answer the three objectives

# Effectiveness of home-PR

We performed meta-analysis using Review Manager software for the primary and secondary outcomes, comparing home-PR with centre-PR or usual care. For continuous data, we calculated the mean difference (MD) (for same scale metric) or standardised mean difference (SMD) (for different scale metrics) with 95% confidence intervals. We used an inverse variance method, and chose a random-effects model to account for between-study heterogeneity in the meta-analysis. At least two studies were needed to perform a meta-analysis and measure the effect size. We used pooled mean differences if the same measurement tool was used in the included RCTs, or if the measurement tool varied among trials, we used SMDs for our primary analysis, but reported pooled MDs for the most commonly used outcome as a sensitivity analysis. A p-value <0.05 was considered statistically significant for the overall effect. For comparison of home-PR and centre-PR, if sufficient studies used the same measure for functional exercise capacity or HRQoL, we defined the noninferiority margin as the minimum clinically important difference (MCID) (e.g. 30 m for the 6MWT).

# Components of home-PR

We identified the components described in internationally recognised guidelines for pulmonary rehabilitation [5, 7, 11, 48] and constructed a matrix comparing components used in the included trials reporting effective interventions *versus* those reporting no effect.

# Uptake, adherence and completion

We used a narrative approach to synthesise reported uptake, engagement, completion and attrition in home-PR and centre-PR groups using the following definitions. Uptake: number of patients who attended the initial/baseline assessment and at least one pulmonary rehabilitation session; engagement: the proportion of pulmonary rehabilitation sessions attended. This reflects the "dose" of the intervention received and may be reported as the number of patients who attended a pre-defined proportion of pulmonary rehabilitation sessions (*e.g.* 70% of sessions); completion: the number of patients who attended the pulmonary rehabilitation discharge assessment and are regarded as having "completed" the pulmonary rehabilitation programme (regardless of the proportion of sessions attended); trial attrition: the number of

people who failed to attend for their post-pulmonary rehabilitation follow-up data collection in a trial. Trials of longer duration may have several follow-up assessments and thus several time points for recording attrition.

# Assessment of the certainty of evidence

To assess the quality of evidence of included studies, we used the five GRADE (Grading of Recommendations, Assessment, Development and Evaluations) considerations (study design, risk of bias, inconsistency, imprecision and indirectness) for the primary outcomes. Using GRADEpro GDT software (gradepro.org), we followed the techniques and guidelines outlined in the Cochrane Handbook for Systematic Reviews of Interventions [49]. We provide footnotes to explain any decisions to downgrade the quality of evidence.

#### Results

#### Study selection

We identified a total of 6185 records from six databases (figure 1) and found 1133 records from forward citation. After removing duplicates, a total of 5857 titles and abstracts were screened, and 78 full-text articles were considered for inclusion by the pairs of reviewers. All disagreements and decisions were discussed within the multidisciplinary team and 62 articles were excluded (supplementary table S1). Thus, we included 16 articles in our review [50–65]. No additional papers were added from the pre-publication update.

## Characteristics of included studies

Of the 16 included studies, 15 were individually randomised trials, and one was a cluster randomised implementation trial [59]. The latter, while relevant to our inclusion criteria, had a very different trial design informing the challenge of implementing home-PR within routine COPD care, rather than providing evidence of effectiveness, and we therefore did not include it in the meta-analysis. Eight studies compared home-PR *versus* usual care [50, 53, 58, 59, 61, 63–65] and seven studies compared home-PR *versus* centre-PR [51, 54–57, 60, 62]. One study compared home-PR against two different comparators (centre-PR and usual care) and is therefore included in both analyses [52] (supplementary table S2 presents key characteristics of included studies, main findings and interpretation).

The trials were conducted in Australia (n=3) [50, 56, 59], Brazil (n=2) [52, 63], Spain (n=2) [54, 65], the United Kingdom (n=2) [57, 58], Canada (n=1) [60], China (n=1) [51], Denmark (n=1) [55], Egypt (n=1) [53], India (n=1) [64], Iran (n=1) [61] and Turkey (n=1) [62]. Of these, nine were high-income countries [50, 54–60, 65], four were upper-middle-income countries [51, 52, 62, 63] and three were lower-middle-income countries [53, 61, 64].

All studies were in people with COPD. In total, 1800 people with a range of severities were recruited to the included trials (range 39–314 participants). Out of the 1733 participants with reported baseline demographic data, 1048 (62%) were male and the mean age ranged from 56 to 79 years.

All pulmonary rehabilitation programmes included either aerobic and/or resistance exercises (aerobic (n=15) [50, 52–65], resistance (n=13) [50–60, 62, 63], both (n=12) [50, 52–60, 62, 63]). Stretching exercises were included in two trials [52, 63] and inspiratory muscle training in one trial [53]. All studies except one [57] had 24 or more exercise sessions; five trials had more than 48 sessions of exercise [50, 53, 56, 58, 64]. All but two [55, 62] of the home-PR programmes included face-to-face training sessions either as inpatients [53, 61], outpatients [51, 52, 54, 57, 58, 60, 63–65] or home visits [50, 56, 59]. Most of the programmes described some form of supervision of the home-based sessions, most commonly telephone calls [52, 56–59, 61, 63, 65] although one used videoconferencing [55] and one study in housebound individuals provided repeated home visits. Other strategies included provision of a manual or written information [51, 57, 58, 61, 62] activity diaries [50, 52, 55, 56, 60, 62, 63, 65], pedometers [54, 56, 65] and heart rate monitors [52].

## Risk-of-bias assessment

Only three studies were at overall low risk of bias [55–57]. Two were at unclear/moderate risk of bias [58, 60] and 11 were at high risk of bias [50–54, 59, 61–65] (supplementary figure S1). Blinding of participants and personnel is impossible due to the nature of the intervention, but only six studies ensured outcome assessors were blind to allocation [54–58, 60]. Computer-generated randomisation sequence was used in 10 studies [51, 52, 55–61] and allocation concealment was described in seven [50, 54–58, 60]; the remaining studies did not provide sufficient information on randomisation [53, 62–65]. We were able to compare reported outcomes with published protocols or trial registrations for six studies [51, 52, 55–57, 59],

all of which were judged to be at low risk of selective reporting bias. Without a protocol for comparison, the remaining studies were designated as unclear risk of bias [50, 53, 54, 58, 60–65].

## Effectiveness of home-PR (objective 1)

Primary outcome: functional exercise capacity

## Home-PR versus usual care

Out of eight trials that compared home-PR with usual care, seven assessed at least one measure of functional exercise capacity [50, 52, 53, 58, 63–65]. Of these, five trials used the 6MWT [50, 52, 53, 63, 64], one trial used both ISWT and ESWT [58] and one trial used ESWT [65]. In one [52] of the seven studies, data were presented in a format that could not be retrieved for meta-analysis. Thus, we included six trials [50, 53, 58, 63–65] in the meta-analysis (figure 2). The pooled estimate showed a statistically significant increase in exercise capacity in home-PR compared with usual care (SMD 0.88, 95% CI 0.32–1.44; p=0.002). The only study not at high risk of bias showed no significant between-group differences [58].

In a subgroup meta-analysis of four RCTs [50, 53, 63, 64] with available data on 6MWT (supplementary figure S2), the pooled estimate showed a statistically significant increase in the mean difference in distance walked in home-PR compared with usual care (MD 61.58 m, 95% CI 45.88–77.29 m; p<0.01). Both the mean difference and the lower limit of the confidence interval exceeded the MCID for the 6-min walk distance (6MWD) of 30 m [66], indicating a clinically significant effect of home-PR.

#### Home-PR versus centre-PR

All the eight trials comparing home-PR with centre-PR assessed at least one measure of functional exercise capacity [51, 52, 54–57, 60, 62]. Of these, seven trials used the 6MWT [51, 52, 54–56, 60, 62], one trial used both ISWT and ESWT [57] and one trial used both cycle endurance test and 6MWT [60]. We included all eight trials [51, 52, 54–57, 60, 62] in the meta-analysis (figure 2). The pooled estimate showed no statistically significant difference in exercise capacity between home-PR and centre-PR (SMD -0.10, 95% CI -0.25-0.05; p=0.21). A sensitivity analysis including only the four studies at low/moderate risk of bias [55–57, 60] did not change the conclusion (SMD -0.02, 95% CI -0.18-0.15;  $I^2=28\%$ ; p=0.85) (supplementary figure S3).

In the meta-analysis of the seven RCTs [51, 52, 54–56, 60, 62] that used 6MWT (supplementary figure S4), the pooled estimates showed no statistically significant difference in the mean difference in distance walked in home-PR compared with centre-PR (MD -6.26m, 95% CI -18.55-6.02; p=0.32). This is within the noninferiority margin of 30 m for the 6MWT, indicating that the clinical effect of home-PR is not inferior to centre-PR for people with COPD.

# Primary outcome: health-related quality of life Home-PR versus usual care

All the eight trials comparing home-PR with usual care assessed at least one measure of HRQoL [50, 53, 58, 59, 61, 63–65]. Of these, four trials used the SGRQ [50, 59, 63, 65], two trials used CRQ [58, 64], one trial used both SGRQ and CAT score [59], one trial used SF-36 [53] and one trial used SF-12 [61]. We excluded the cluster RCT [59] from the meta-analysis because it informed implementation (as opposed to effectiveness) of home-PR in routine primary care management of COPD and was thus not comparable with the other trials. Thus, we included seven trials [50, 53, 58, 61, 63–65] in the meta-analysis (figure 2) and the pooled estimate (SGRQ-total, CRQ-mastery, SF-36-physical, SF-12) showed statistically significant improvement in HRQoL in the home-PR group compared with usual care (SMD -0.62, 95% CI -0.88-0.36; p<0.01). The only study not at high risk of bias showed no significant between group differences [58].

Meta-analysis of the three RCTs that used SGRQ [50, 63, 65] (supplementary figure S5) showed a statistically significant improvement that exceeded the MCID of 4.0 in all the domains except the "impact" domain. The effect on overall SGRQ in the home-PR group compared with usual care showed an MD -5.66 (95% CI -7.94-3.39; p<0.01) that exceeded the MCID.

Meta-analysis of the two RCTs [58, 64] that used the CRQ (supplementary figure S6) showed a statistically significant improvement (p=0.010) that exceeded the MCID of 0.5 in all the domains (dyspnoea, emotion, fatigue, mastery).

# Home-PR versus centre-PR

All seven trials comparing home-PR with centre-PR assessed at least one measure of HRQoL [51, 54–57, 60, 62]. Of these, four trials used the CRQ [54, 56, 57, 60] and three trials used the CAT score [51, 55, 62].

# Comparison: Home-PR versus usual care

## a) Functional exercise capacity

	Home-	PR	Usual c	are		SMD	SMD
Study or subgroup	Mean±sp	Total	Mean±sp	Total	Weight	IV, random (95% CI)	IV, random (95% CI)
Boxall, 2005 (6MWT)	39±91.85	30	4.2±114.36	30	17.9%	0.33 (-0.18-0.84)	+-
GHANEM, 2010 (6MWT)	52.71±30.01	25	-15.44±35.83	14	14.5%	2.07 (1.26-2.89)	<del></del>
JOHNSON-WARRINGTON, 2016 (ISWT)	45±51.85	39	30±70.37	39	18.6%	0.24 (-0.21-0.69)	<del> </del>
Pradella, 2015 (6MWT)	65.7±65.7	32	5.6±24.2	18	16.7%	1.08 (0.46-1.70)	
Singh, 2003 (6MWT)	54±62.19	20	6.3±222.74	20	16.7%	0.29 (-0.34-0.91)	<del> </del>
Varas, 2018 (ESWT)	6.1±5.7	21	-0.9±2.1	19	15.6%	1.57 (0.85-2.28)	<del>-</del>
Total (95% CI)		167		140	100.0%	0.08 (0.32-1.44)	•
Heterogeneity: Tau <sup>2</sup> =0.38; Chi <sup>2</sup> =25.7 Test for overall effect: Z=3.09 (p=0.00		001); I <sup>2</sup>	= 81%				-4 -2 0 2 4

b) Health-related quality of life

Favours usual care Favours home-PR

#### Home-PR Usual care SMD SMD IV, random (95% CI) Study or subgroup Weight IV, random (95% CI) Mean±sp Total Mean±sp Total BOXALL, 2005 (SGRQ) -1.4±17.07 -0.26 (-0.77-0.25) -5.8±16.13 30 30 17.5% GHANEM, 2010 (SF-36) -30±40.0494 25 2.18±21.459 14 11.2% -0.91(-1.60--0.22) JOHNSON-WARRINGTON, 2016 (CRQ) 20.7% -0.34 (-0.79-0.10) -1.41±1.48 39 -0.89±1.51 39 MOHAMMADI, 2013 (SF-12) -6.8±7.8309 20 -2.56±5.6748 20 12.7% -0.61 (-1.24-0.03) PRADELLA, 2015 (SGRQ) 14.1% -6.4±16.1 32 3.1±12.2 18 -0.63 (-1.22--0.04) SINGH, 2003 (CRQ) -0.89+1.3103 20 -0.05+1.1315 20 12.6% -0.67 (-1.31--0.03) VARAS, 2018 (SGRQ) -4.4±3.6 21 1±4.3 19 11.1% -1.34 (-2.03--0.65) Total (95% CI) 187 160 100.0% -0.62 (-0.88--0.36) Heterogeneity: Tau<sup>2</sup>=0.03; Chi<sup>2</sup>=8.18, df=6 (p=0.23); I<sup>2</sup>= 27% -2 0

Test for overall effect: Z=4.68 (p<0.00001)

Favours home-PR Favours usual care

Favours centre-PR Favours home-PR

# Comparison: home-PR versus centre-PR

#### c) Functional exercise capacity

, ,	Home-F	PR	Centre-	-PR		SMD			SMD		
Study or subgroup	Mean±sp	Total	Mean±sp	Total	Weight	IV, random (95% CI)		IV, rai	ndom (95	% CI)	
CHEN, 2018 (6MWT)	38.29±78.66	29	43.01±86.62	26	6.8%	-0.06 (-0.59-0.47)			-		
Güell, 2008 (6MWT)	54±103.13	28	65±126.67	29	7.0%	-0.09 (-0.61-0.43)		-			
Hansen, 2020 (6MWT)	17.2±46.74	67	23.5±46.74	67	13.3%	-0.13 (-0.47-0.21)					
Holland, 2017 (6MWT)	29.39±70.15	80	10.82±71.55	86	15.2%	0.26 (-0.04-0.57)			+	-	
Horton, 2018 (ISWT)	21±209.307	145	41±216.4133	142	20.7%	-0.09 (-0.33-0.14)		_			
Maltais, 2008 (6MWT)	8±51.0452	126	11±51.0452	126	19.4%	-0.06 (-0.31-0.19)		_	-		
Mendes de Oliveira, 2010 (6MWT)	73.2±50.2	42	93.6±70.6	46	9.7%	-0.33 (-0.75-0.09)					
Pehlivan, 2020 (6MWT)	18±61.18	39	66±102.75	32	8.0%	-0.58 (-1.050.10)		•	_		
Total (95% CI)		556		554	100.0%	-0.10 (-0.25-0.05)					
Heterogeneity: Tau <sup>2</sup> =0.01; Chi <sup>2</sup> =10.3	, ,,	'); I <sup>2</sup> = 3	3%				-1	-0.5	0	0.5	1
Test for overall effect: Z=1.26 (p=0.2	(1)						Eavo	urc contro l	DD Fav	ours hom	o DD

# d) Health-related quality of life

., 1										
	Home-F	PR	Centre-	-PR		SMD	SMD			
Study or subgroup	Mean±sp	Total	Mean±sp	Total	Weight	IV, random (95% CI)	IV, random (95	% CI)		
CHEN, 2018 (CAT)	-3.2±5.06	29	-1.27±4.95	26	7.5%	-0.38 (-0.91-0.15)		+		
GÜELL, 2008 (CRQ)	-0.1±1.09	28	-0.65±1.27	29	7.7%	0.46 (-0.07-0.98)		<del>                                     </del>		
Hansen, 2020 (CAT)	-1.7±6.1496	67	-0.3±6.1495	67	14.7%	-0.23 (-0.57-0.11)		+		
HOLLAND, 2017 (CRQ)	-2.43±4.6733	80	-2.02±4.6175	86	16.9%	-0.09 (-0.39-0.22)	_	+		
Horton, 2018 (CRQ)	-0.28±1.9146	145	-0.61±1.7755	142	22.6%	0.18 (-0.05-0.41)				
Maltais, 2008 (CRQ)	-0.49±0.9642	126	-0.51±0.9075	126	21.3%	0.02 (-0.23-0.27)			_	
Pehlivan, 2020 (CAT)	-5±8.76	39	-6±9.34	32	9.3%	0.11 (-0.36-0.58)		Ι'	_	
Total (95% CI)		514		508	100.0%	0.01 (-0.15-0.17)	•	lack		
Heterogeneity: Tau <sup>2</sup> =0.02; Chi <sup>2</sup> =9.	23. df=6 (p=0.16)	: I <sup>2</sup> = 35	5%			<b>—</b>	+	+		$\overline{}$
Test for overall effect: Z=0.17 (p=0		,				-2	-1	0	1	2
Ψ.	,						Favours home-PI	R Favo	ours centre	-PR

FIGURE 2 Comparison of primary outcomes. a, b) Comparing home-pulmonary rehabilitation (PR) with usual care for a) functional exercise capacity and b) health-related quality of life; c, d) comparing home-PR with centre-PR for c) functional exercise capacity and d) health-related quality of life. 6MWT: 6-min walk test; ISWT: incremental shuttle walking test; ESWT: endurance shuttle walking test; IV: inverse variance; SMD: standardised mean difference; SGRQ: St George's Respiratory Questionnaire; CRQ: Chronic Respiratory Questionnaire; CAT: COPD Assessment Test.

We included all seven trials [51, 54–57, 60, 62] in the meta-analysis (figure 2) and the pooled estimate (CRQ-mastery, CAT score) showed no statistically significant difference in the HRQoL in home-PR compared with centre-PR (SMD 0.01, 95% CI -0.15–0.17; p=0.87). A sensitivity analysis including only the four studies at low/moderate risk of bias [55–57, 60] did not change the conclusion (SMD -0.00, 95% CI -0.16–0.17;  $I^2$ =30%; p=0.98) (supplementary figure S7).

Meta-analysis of the four RCTs [54, 56, 57, 60] that used CRQ (supplementary figure S8) showed no statistically significant between-group differences (p=0.21) in any of the domains of CRQ (dyspnoea, emotion, fatigue, mastery).

Meta-analysis of the three RCTs [51, 55, 62] that used the CAT score (supplementary figure S9) favoured home-PR compared with centre-PR (MD -1.53, 95% CI -2.81-0.24; p=0.02).

# Secondary outcome: dyspnoea Home-PR versus usual care

Two trials [59, 65] assessed dyspnoea using the modified Medical Research Council (mMRC) scale and compared home-PR with usual care. The implementation cluster RCT [59] concluded that mMRC grades were not significantly different between groups. The other RCT also showed no statistically significant changes (p=0.22) in dyspnoea level associated with home-PR compared to usual care [65].

#### Home-PR versus centre-PR

Two trials [56, 62] assessed dyspnoea using mMRC and compared home-PR *versus* centre-PR. Meta-analysis showed no statistically significant changes between the groups in dyspnoea level (supplementary figure S10) between home-PR and centre-PR (MD -0.12, 95% CI -0.44-0.21; p=0.48).

# Secondary outcome: anxiety and depression

## Home-PR versus usual care

One trial [59] measured anxiety and depression using the Hospital Anxiety and Depression Scale (HADS) and compared the effect between home-PR and usual care. There was no statistically significant between-group difference in either anxiety (p=1.00) or depression (p=0.09).

# Home-PR versus centre-PR

Two trials [55, 57] measured anxiety and depression using HADS and compared the effects between home-PR and centre-PR. Meta-analysis showed no statistically significant between-group difference in anxiety or depression (supplementary figures S11 and S12) (anxiety: MD -0.33, 95% CI -1.81-1.15; p=0.66; depression: MD -0.03, 95% CI -1.28-1.22; p=0.97).

# Association of components of home-PR with effective interventions (objective 2)

Table 2 presents a matrix of components of home-PR mapped to effectiveness.

There were no obvious differences in the components of the home-PR between effective and ineffective studies or in the number of components included, supervision provided or duration of the course.

#### Uptake, engagement, completion and trial attrition (objective 3)

Table 3 shows details of recruitment, uptake, engagement, completion of pulmonary rehabilitation sessions and trial attrition.

# Screening and eligibility for the trials

Nine studies [51, 52, 55–61] provided details of the eligibility screening process, reporting recruitment rates between 12% and 56%. Five trials cited the presence of comorbidity as a reason for excluding between 3% and 14% of screened participants [51, 55–58]. Three studies reported that approximately one in five (22.8%, 18.3% and 12.0% [55–57]) potentially eligible patients declined to participate because of a strong preference for centre-PR. In contrast, one trial comparing home-PR *versus* centre-PR excluded 55% because they definitely wanted home-PR [55]. Distance/travel was cited as a reason for nonparticipation in two trials [51, 60].

# Uptake of pulmonary rehabilitation

The implementation cluster RCT reported an uptake of 66% among the 107 patients referred by their general practitioner [59]. Two trials reported that two patients did not attend any pulmonary rehabilitation sessions [50, 65].

Author, year		Exer	cise typ	oe .	Edu	PR frequency;	To	tal sessi	ons	Training, supervision and	FEC	HRQoL
[reference]	AE		Flex			duration	<24	24-48	≥48	monitoring in the home-PR group	0	و
Home-PR <i>versus</i> usual (	care	(n=9)										
GHANEM, 2010 [53]	✓	(II 3)	Х	1	1	2× per week; 8 weeks	Χ	✓	Χ	1 inpatient training session, then unsupervised home exercise	S <sup>#</sup>	S <sup>#</sup>
Pradella, 2015 [63]	✓	✓	1	Х	✓	3× per week; 8 weeks	Х	✓	Х	Outpatient training (first week), then diary + weekly TC to encourage home exercises	S <sup>#</sup>	S <sup>#</sup>
Varas, 2018 [65]	1	Χ	Χ	Χ	1	5× per week; 8 weeks	Χ	1	Χ	5 outpatient training sessions, then diary + pedometer + weekly TC	S <sup>#</sup>	S <sup>#</sup>
Singh, 2003 [64]	✓	Х	Х	Х	Х	2× per day; 4 weeks	Х	✓	Х	Outpatient training sessions, then weekly supervision (mode not described)	NS <sup>#</sup>	S <sup>#</sup>
BOXALL, 2005 [50]	1	✓	Χ	Х	Х	Daily; 12 weeks	Χ	Χ	✓	Weekly home visits for 6 weeks + diary, then fortnightly home visits	NS <sup>#</sup>	NS <sup>#</sup>
Johnson-Warrington, 2016 [58]	✓	✓	Х	Х	1	3× per week; 12 weeks	Χ	Х	✓	1 face-to-face introductory session and given manual, then fortnightly TC	NS	NS <sup>#</sup>
LIANG, 2019 [59]	1	1	Χ	Χ	1	NR; 8 weeks	1	Χ	Χ	1 home visit, then weekly TC	NS	NS
Mendes de Oliveira, 2010 [52]	✓	1	1	Х	1	3× per week; 12 weeks	Х	✓	Х	1 outpatient education + exercise training session, then diary + heart rate monitor + TC	NR	NS
Монаммарі, 2013 [61]	1	Χ	Χ	Χ	1	3× per week; 8 weeks	Χ	✓	Χ	3 inpatient training sessions, then manual + TC (alternate days)	NR	NS
Home-PR versus centre	-PR (	n=8)										
PEHLIVAN, 2020 [62]	1	✓	Х	Х	<b>√</b>	2–7× per week; 8 weeks	Х	1	X	Exercise sessions + daily walking + diary + manual Supervision NR	S	NS <sup>#</sup>
Снем, 2018 [51]	Х	1	Х	Х	✓	3× per week; 12 weeks	Х	1	X	1 outpatient education and training session, then home exercise + manual Supervision NR	NS <sup>#</sup>	NS <sup>#</sup>
GÜELL, 2008 [54]	1	✓	Х	1	✓	3× per week; 9 weeks	Х	✓	X	4 outpatient education and training sessions, then home exercise/ walking sessions + pedometer Supervision NR	NS <sup>#</sup>	NS
Hansen, 2020 [55]	/	✓	Х	Х	1	3× per week; 10 weeks	Х	1	Χ	3× per week exercise and education sessions, supervised by video-conference + diary	NS	NS
Holland, 2017 [56]	✓	✓	Х	Х	✓	Most days; 8 weeks	Х	Χ	1	1 home visit, then home exercise sessions supervised by weekly TC + diary + pedometer	NS	NS <sup>#</sup>
Horton, 2018 [57]	1	1	Х	Х	✓	3× per week; 7 weeks	Χ	✓	Χ	1 outpatient training session, then home exercise supervised by TC (×2) + manual	NS	NS
Maltais, 2008 [60]	1	1	Х	Х	1	3× per week; 8 weeks	Χ	1	X	Outpatient education and training sessions, then home-based exercise + diary	NS	NS
MENDES DE OLIVEIRA, 2010 [52]	1	1	1	Х	1	3× per week; 12 weeks	Х	1	Χ	1 outpatient education + exercise training session, then diary + heart rate monitor + TC	NS <sup>#</sup>	NR

AE: aerobic training; RT: resistance training; flex: flexibility training; RMT: respiratory muscle training; Edu: education; FEC: functional exercise capacity; HRQoL: health-related quality of life; S: significant between-group difference; TC: telephone call; NS: nonsignificant between-group difference; NR: not reported. #: improved above minimal clinically important difference.

# Engagement with the programme

Only four studies defined "engagement" as a pre-determined proportion of pulmonary rehabilitation sessions attended [55, 56, 59, 60]. Using the widely cited 70% threshold [67], Holland *et al.* [56] showed that engagement with home-PR was nearly twice that of centre-PR (91% *versus* 49%; relative risk of

uthor, year [reference]	Screened (reasons for ineligibility) Recruited: randomised	Uptake of PR/ usual care (reasons for nonstart)	Dose of PR and definition of engagement	Engaged with defined number of sessions (reasons for non-engagement)	Completed PR programme (reasons for dropout)	Trial attrition rate (reason for attrition)
ome-PR <i>versus</i> usua	l care					
BOXALL, 2005 [50] RoB: high	Eligibility screening not reported 60 recruited: home-PR n=30, usual care n=30	Uptake of PR: home-PR 28/30 (93%) (2 ill-health)	Home-PR: 11 sessions (12 weeks: 11 visits + daily unsupervised exercise) Engagement not defined	Engagement not reported	Home-PR: 23/30 (77%) (3 withdrew, 1 died, 1 ill-health)	Post-PR Home-PR: 7/30 (23%) (3 withdrew, 1 died, 3 ill-health) Control: 7/30 (23%) (2 withdrew, 2 died, 1 ill-health, 2 moved)
CHEN, 2018 [51] RoB: high	265 screened for eligibility (77 lost to contact, 44 distances >44 km, 53 declined, 36 comorbidity, 38 other) 55 (21%) recruited: home-PR n=29, usual care n=26	Uptake not reported	Home-PR: 36 sessions (12 weeks: 3× per week unsupervised exercise) Engagement not defined	Engagement not reported	Home-PR: 25/29 (86%) (3 ill-health, 1 moved) Usual care: 22/26 (85%) (1 ill-health, 3 not serious enough)	Post-PR Home-PR: 4/29 (14%) (1 moved, 3 ill-health) Usual care: 4/26 (15%) (1 not serious, 1 ill-health)
GHANEM, 2010 [53] RoB: high	Eligibility screening not reported 39 recruited: home-PR n=25, usual care n=14	Uptake not reported	Home-PR: 48 sessions (8 weeks: alternate days unsupervised exercise) Engagement not defined	Engagement not reported	Completion not reported	Post-PR Home-PR: 0/25 (0%) Usual care: 0/14 (0%)
JOHNSON-WARRINGTON, 2016 [58] RoB: moderate	464 screened for eligibility 175 declined, 76 not eligible, 49 comorbidity, 90 lost to contact 78 (17%) recruited: home-PR n=39, usual care n=39	Uptake not reported	Home-PR: 42 sessions (12 weeks 6 TCs + 3× per week unsupervised exercise) Engagement not defined	Engagement not reported	Home-PR: 35/39 (90%) (2 ill-health, 1 preferred centre-PR, 1 not COPD) Usual care: 36/39 (92%) (3 died)	Post-PR Home-PR: 4/39 (10%) (1 wanted centre-PR; 2 ill-health; 1 not COPD) Usual care: 3/39 (8%) (3 died)
Монаммаді, 2013 [61] RoB: high	106 assessed for eligibility 40 (38%) recruited: home-PR n=20, usual care n=20	Uptake not reported	Home-PR: 24 sessions (8 weeks: 3 sessions then daily TCs + unsupervised sessions 3× per week) Engagement not defined		Completion not reported	Not reported No attrition reported
PRADELLA, 2015 [63] RoB: high	Eligibility screening not reported 50 recruited: home-PR n=32, usual care n=18	Uptake not reported	Home-PR: 24 sessions (8 weeks: weekly TCs + 3 unsupervised sessions per week) Engagement not defined	Engagement not reported	Home-PR: 29/32 (91%) (1 withdrew; 1 died, 1 AECOPD)	Post-PR Home-PR: 3/32 (9%) (1 died, 1 withdrew, 1 AECOPD) Usual care: 3/18 (17%) (2 withdrew, 1 AECOPD)

Author, year [reference]	Screened (reasons for ineligibility) Recruited: randomised	Uptake of PR/ usual care (reasons for nonstart)	Dose of PR and definition of engagement	Engaged with defined number of sessions (reasons for non-engagement)	Completed PR programme (reasons for dropout)	Trial attrition rate (reason for attrition)
Singh, 2003 [64] RoB: high	Eligibility screening not reported 40 recruited: home-PR n=20, usual care n=20	Uptake not reported	Home-PR: 4 sessions (4 weeks: weekly visits + daily unsupervised exercise) Engagement not defined	Engagement not reported	Engagement not reported	Post-PR Home-PR: 0/20 (0%) Control: 0/20 (0%)
Varas, 2018 [65] RoB: high	Eligibility screening not reported 40 recruited: home-PR n=21, usual care n=19	Uptake of PR: home-PR 19/21 (90%) (2 withdrew)	Home-PR: 8 sessions (1× per week + unsupervised exercise ×8 weeks) Engagement not defined	Engagement not reported	Home-PR: 17/21 (81%) (2 did not complete)	Post-PR Home-PR: 4/21 (19%) (4 withdrew) Usual care: 3/19 (16%) (3 withdrew) 3 months and 12 months Home-PR: 4/21 (19%) Usual care: 3/19 (16%)
luster randomised	implementation trial: home-PR v	versus usual care				
LIANG, 2019 [59] RoB: high	Cluster RCT: 21 practices/ group 1050 screened for eligibility 272 (26%) recruited: home-PR n=157, control n=115	GP referred for PR: home-PR 107/157 (68%) Uptake of PR: home-PR 71/107 (66%)	Home-PR: 8 sessions (8 weeks: 1 session + weekly TC unsupervised exercise) Engagement defined as ≥70% sessions attended	Engaged ≥70% Home-PR: 49/107 (46%)	Completion not reported	6 months Home-PR: 39/157 (25%) Usual care: 21/115 (18%) 12 months Home-PR: 44/157 (28%) (27 lost to follow-up, 15 withdrew, 2 died) Usual care: 38/115 (33%) (29 lost to follow-up, 7 withdrew, 1 moved, 1 died)
lome-PR versus centre-PR						
GÜELL, 2008 [54] RoB: high	Eligibility screening not reported 57 recruited: home-PR n=28, centre-PR n=29	Uptake not reported	Home-PR: 27 sessions (9 weeks: 4 sessions + 3× per week unsupervised Centre-PR: 27 sessions (9 weeks: 3× per week) Engagement not defined	Engagement not reported	Home-PR: 23/28 (82%) (4 dropped out, 1 chest pain) Centre-PR: 28/29 (96%) (1 dropped out)	Post-PR Home-PR: 5/28 (18%) Centre-PR: 1/29 (4%) 6 months Home-PR: 8/28 (29%) (4 withdrew, 1 ill-health, 3 los to follow-up) Centre-PR: 6/29 (21%) (1 dropped out, 5 lost to follow-up)

TABLE 3 Continued						
Author, year [reference]	Screened (reasons for ineligibility) Recruited: randomised	Uptake of PR/ usual care (reasons for nonstart)	Dose of PR and definition of engagement	Engaged with defined number of sessions (reasons for non-engagement)	Completed PR programme (reasons for dropout)	Trial attrition rate (reason for attrition)
Hansen, 2020 [55] RoB: low	1099 assessed for eligibility: (608 declined centre-PR, 251 declined home-PR, 40 comorbidity, 66 other) 134 (12%) recruited: home-PR n=67, centre-PR n=67	Uptake not reported	Home-PR: 30 sessions (10 weeks: 3 sessions per week) Centre-PR: 20 sessions (10 weeks: 2 sessions per week) Engagement defined as ≥70% sessions attended Attendance defined as participating in the whole session	Engaged ≥70% Home-PR: 49/67 (73%) Centre-PR: 42/67 (63%) Median (IQR) number sessions attended Home-PR: 25/30 (20–28) Centre-PR 16/20 (8/19) Home-PR >70% engagement: OR 1.68, 95% CI 0.78–3.37; p<0.27	Home-PR: 57/67 (85%) (6 dropped out, 2 ill-health, 1 died, 1 AECOPD) Centre-PR: 43/67 (64%) (10 dropped out, 8 ill-health, 2 died, 4 AECOPD) Home-PR completing: OR 3.18, 95% CI 1.37–7.35; p<0.01	Post-PR Home-PR: 20/67 (30%) Centre-PR: 26/67 (39%) 3-month follow-up Home-PR: 29/67 (43%) Centre-PR: 26/67 (39%)
Holland, 2017 [56] RoB: low	295 assessed for eligibility (27 recent PR, 10 comorbidities, 5 recent AECOPD, 67 declined (54 wanted centre-PR), 120 other) 166 (56%) recruited: home-PR n=80, centre-PR n=86	Uptake not reported	Home-PR: 8 sessions (8 weeks: visit then weekly TCs + unsupervised sessions) Centre-PR: 16 sessions (8 weeks: twice weekly) Engagement defined as ≥70% sessions attended	Mean/total sessions attended (range) Home-PR: 7.4/8 (0–8) Centre-PR 8.3/16 (0–16) Engaged ≥70% Home-PR: 73/80 (91%) Centre-PR: 42/86 (49%) Relative risk of noncompletion in centre-PR 1.91 (95% CI 1.52–2.41)	Home-PR: 73/80 (91%) (1 died, 1 lost to follow-up, 5 declined) Centre-PR: 77/86 (89%) (1 died, 1 lost to follow-up, 7 declined)	Post-PR Home-PR: 7/80 (9%) Centre-PR: 9/86 (11%) 12-month follow-up Home-PR: 18/80 (24%) (4 lost to follow-up, 9 declined follow-up, 5 died) Centre-PR: 24/86 (28%) (10 lost to follow-up, 10 declined follow-up, 4 died)
Ноктом, 2018 [57] RoB: low	1162 assessed for eligibility (185 DNA, 32 comorbidities, 606 not eligible, 140 wanted centre-PR, 100 declined, 199 other) 287 (25%) recruited: home-PR n=145, centre-PR n=142	Uptake not reported	Home-PR: 21 sessions (7 weeks: 3 unsupervised sessions a week) Centre-PR: 14 sessions (7 weeks: twice weekly) Engagement not defined	Engagement not reported	Home-PR: 94/145 (85%) (16 lost to follow-up, 16 comorbidities, 2 died, 2 wanted centre-PR, 17 other) Centre-PR: 84/142 (59%) (30 lost to follow-up, 12 comorbidities, 1 died, 3 wanted home-PR, 12 others)	Post-PR Home-PR: 51/145 (35%) Centre-PR: 58/142 (41%) 6 months Home-PR: 70/145 (48%) (7 lost to follow-up, 3 DNA, 3 declined, 13 comorbidities, 3 other) Centre-PR: 72/142 (51%) (8 lost to follow-up, 1 comorbidity, 3 died, 2 others)

Continued

Author, year [reference]	Screened (reasons for ineligibility) Recruited: randomised	Uptake of PR/ usual care (reasons for nonstart)	Dose of PR and definition of engagement	Engaged with defined number of sessions (reasons for non-engagement)	Completed PR programme (reasons for dropout)	Trial attrition rate (reaso for attrition)
Maltais, 2008 [60] RoB: moderate	631 assessed for eligibility (214 declined, 27 transport problems, 1 died, 29 others) 252 (40%) recruited: home-PR n=126, centre-PR n=126	Uptake not reported	Home-PR: 24 sessions (8 weeks: 3 unsupervised sessions per week) Centre-PR: 24 sessions (4× 8 weeks: 3 sessions per week) Engagement defined as ≥60% sessions attended	Engaged ≥60% Home-PR: 123/126 (98%) Centre-PR:117/126 (93%)	Completion not reported	Post-PR Home-PR: 7/126 (6%) Centre-PR: 12/126 (10%) 12 months Home-PR: 19/126 (15%) (2 lost to follow-up, 16 withdrew, 1 died) Centre-PR: 17/126 (13%) (2 lost to follow-up, 14 withdrew, 1 died)
PEHLIVAN, 2020 [62] RoB: high	71 assessed for eligibility 71 recruited: home-PR n=39, centre-PR n=32	Uptake not reported	Home-PR: 32 sessions (8 weeks: 4 unsupervised sessions per week) Centre-PR: 16 sessions (8 weeks: 2 sessions per week) Engagement not defined	Engagement not reported 4 home-PR patients were excluded for "noncompliance"	Home-PR: 35/39 (4 discontinued) Centre-PR: 32/32	Post-PR Home-PR: 4/39 (10%) (4 withdrew) Centre-PR: 0/32 (0%)
hree-arm trial (home	e-PR versus centre PR versus us	ual care	0-0			
Mendes de Oliveira, 2010 [52] RoB: high	216 assessed for eligibility (65 declined, 32 ineligible, 2 died) 117 (54%) recruited: home-PR n=42, centre-PR n=46, usual care n=29	Uptake not reported	Home-PR: 36 sessions (12 weeks: TCs + 3 unsupervised sessions per week) Centre-PR: 36 sessions (12 weeks: 3 sessions per week) Engagement not defined	Engagement not reported	Home-PR: 35/42 (83%) (7 "abandoned" the programme) Centre-PR: 27/46 (59%) (7 "abandoned" the programme)	Post-PR Home-PR: 9/42 (21%) (2 los to follow-up) Centre-PR: 22/45 (50%) (4 lost to follow-up) Usual care: 0/29 (0%)

Home-PR: home-based pulmonary rehabilitation; RoB: risk of bias; centre-PR: centre-based pulmonary rehabilitation; TC: telephone contact; AECOPD: acute exacerbation of COPD RCT: randomised controlled trial; GP: general practitioner; IQR: interquartile range; DNA: did not attend.

nonengagement in centre-PR: 1.91, 95% CI 1.52–2.41). In contrast, two studies [55, 60] showed no between-group difference, although the latter used a lower threshold ( $\geq$ 60%) and reported that >90% of the participants in both groups achieved this threshold. The implementation cluster RCT reported 46% engaged with  $\geq$ 70% of the pulmonary rehabilitation programme.

# Completion of post-PR assessment and trial attrition

In the trial context, completion of the post-PR assessment was generally reported as attrition (*i.e.* loss to trial follow-up). Rates of attrition at the post-PR follow-up assessment ranged from 0% to 51%, but with no consistent pattern to suggest that mode of delivery affected follow-up.

#### Quality of evidence

Using GRADE, we judged primary outcomes (functional exercise capacity and HRQoL) of the review to provide low-certainty evidence when home-PR was compared with centre-PR and very low-certainty evidence when home-PR was compared with usual care. Downgrading for risk of bias was influenced by performance bias and some concerns in some or most of the domains of included studies. We additionally downgraded for imprecision because of use of SMD to assess the effect and/or small sample size, and for inconsistency due to heterogeneity in home-PR when compared with usual care (supplementary table S3).

#### Discussion

# Summary of findings

Our systematic review identified 16 studies involving a total of 1800 COPD patients from 11 different countries. The effects of home-PR on exercise capacity and/or HRQoL in people with COPD were compared to either centre-PR (n=7) or usual care (n=8). One study had both comparators [52]. Overall, statistically significant improvement was found in functional exercise capacity and HRQoL in home-PR groups when compared with usual care, but no statistically significant differences were found in exercise capacity and HRQoL between home-PR and centre-PR groups. All studies that compared home-PR with usual care were at high risk of bias, except one which was at moderate risk of bias [58]. Conversely, among the studies that compared home-PR with centre-PR, three were at low risk of bias, one was at moderate risk of bias and four were at high risk of bias. No distinguishable patterns were found in exercise components, supervision and monitoring among the three trials [53, 63, 65] that had statistically significant between-group differences and exceeded MCIDs for both the primary outcomes when compared to other included studies. Rates of attrition at the post-PR follow-up assessment ranged from 0% to 51%, but with no consistent pattern to suggest that mode of delivery affected follow-up.

# Strength and limitations

A strength of this systematic review is its comprehensive literature search constructed with the help of an expert librarian. We were open to including non-English language papers. We employed a rigorous methodology following a written protocol that has been published [36]. Although we searched for a wide range of chronic respiratory diseases, the included trials only recruited people with COPD, so the findings are not generalisable to people with other chronic respiratory diseases. We used generic terms for chronic respiratory diseases and named some of the commonest diseases, but our search might have missed some studies as all disease names were not explicitly included in the search strategy. Although we had low (home-PR *versus* centre-PR) or very low (home-PR *versus* usual care) confidence in our GRADE assessment for primary outcomes, this was influenced by multiple outcomes measures which we presented as an SMD in our meta-analysis. This emphasises the importance of agreed standardised outcomes for trials [68].

Six reviewers worked independently in pairs (as in the traditional model) and ensured that all titles and abstracts were duplicate-screened, and disagreements resolved in discussion involving the whole team as necessary. Involvement of six reviewers allowed us to complete the review in a timely manner and without overburdening any individual. The main limitation is the potential for inconsistency, so before starting screening, 100 articles were selected randomly from the total records by the study librarian and given to each pair to screen as a training exercise. Decisions were discussed within the study team and operational rules clarified and agreed.

# Interpretation in the light of published literature Effectiveness of home-PR

Our findings show that home-PR can be a clinically effective alternative to centre-PR for people with COPD in different settings [8, 27, 69] with the findings that both the MD and the lower limit of the confidence interval exceeded the MCID for the 6MWD [66] indicating a clinically significant effect in improving exercise capacity. This extends the findings of the recently published Cochrane review that

assessed the effect of telerehabilitation (either delivered in local community centres or at home) in two ways [33]. Firstly, the home-based programme remained effective despite the lack of the face-to-face group support available in a traditional centre-based pulmonary rehabilitation. This is of particular value in the context of a pandemic when infection control is an important consideration and may preclude group settings. Secondly, most of the telerehabilitation interventions in the Cochrane review [33] used video-conferencing or web-based systems to create virtual groups whereas in our home-based studies over half relied on individual telephone calls, and only one study provided a group-based structured pulmonary rehabilitation programme *via* video-conferencing [55]. This extends the findings to LMIC countries (and indeed some rural areas of high-income countries) with limited access to reliable internet connections. In addition to improving functional exercise capacity and HRQoL, meta-analysis of secondary outcomes showed that home-PR improved dyspnoea, anxiety and depression. These findings hint that home-PR may reduce stress associated with accessing and participating in centre-PR [13], as well as helping to develop confidence in the ability to exercise unsupervised [70].

#### Components of home-PR

Less than 2% of all patients with COPD globally can be served by the existing centre-PR programmes [71], and increasing access to and benefit from remote pulmonary rehabilitation remains a significant clinical and research priority [72]. To do this with confidence, providers of pulmonary rehabilitation services will want to know which components they should include and how to adapt them to home-PR. Although our review did not provide consistent evidence of which components or models of care were associated with effective interventions, others have reported that the intensity of supervision and monitoring increase chances of success in comparison to unsupervised programmes [73, 74]. Most of the interventions in our included studies provided between 24 and 28 home-based sessions with a broad range of arrangements for supervision, but no one approach was associated with effective interventions.

# Uptake, engagement, completion and trial attrition

The terms uptake, engagement and completion are often used interchangeably without clear definition. Data are rarely reported in full; a recent systematic review only identified one trial with comprehensive uptake and completion data [75]. Uptake, defined as the number of patients who attended the initial/baseline assessment and at least one pulmonary rehabilitation session, may be referred to as "enrolled" or in a trial context "recruited" [67]. In our review, uptake was not reported in any of the studies that compared home-PR to centre-PR. Engagement is the proportion of pulmonary rehabilitation sessions attended. This is often assessed as the number of patients who have attended a pre-defined proportion of pulmonary rehabilitation sessions (e.g. 70% of sessions) and is sometimes referred to as "completion rate" [55, 56], or "adherence" [60] or "compliance" [50]. Of the included studies, only four trials defined engagement [55, 56, 59, 60] and only three trials reported this clearly [55, 56, 60]. Engagement with defined sessions in home-PR varied from 73% to 98%, whereas in centre-PR engagement ranged from 49% to 93%. Completion can also be defined as the number of patients who attended the post-PR discharge assessment and are regarded as having "completed" the pulmonary rehabilitation programme (even if they attended very few of the sessions). Some trials referred to participants who did not complete as having "dropped out" of the pulmonary rehabilitation programme [67].

From a trial design perspective, attrition is the number of people who do not attend follow-up assessments and may be described as having "withdrawn" from the trial. Trials of longer duration may have several follow-up assessments and thus several time points for recording attrition. Attrition rates at the post-PR follow-up evaluation ranged from 0% to 51% in our review, but there was no consistent pattern to suggest that mode of delivery influenced follow-up.

# Implications for clinical practice and research

This systematic review gives confidence that home-PR can be an effective option to the traditional models of centre-PR programmes which could extend access for people with COPD to this effective intervention, although the low certainty of the evidence warrants further high-quality evaluation. Specifically, there is evidence that pulmonary rehabilitation improves outcomes in bronchiectasis [30] and ILD [31], but our studies were COPD-specific, so further investigation is required to establish whether home-PR is suitable for chronic respiratory diseases other than COPD. This may be of particular importance in rural areas of LMICs where poor access to investigations mean that the diagnosis may not be clear and limited facilities and travel infrastructure make remote delivery an important option [21].

While we may not have been able to identify specific components that contributed to effectiveness, providers will note that almost all the interventions included aerobic training and resistance training along with a programme of education. Regular remote supervision varied, but did not have to be technologically

complex; many used telephone calls often supplemented by maintaining an exercise diary. We recommend that future trials address issues of uptake, engagement, completion and attrition, and adopt standard terminology in order to provide clarity.

#### Conclusion

Our review concludes with low confidence that home-PR is as effective as centre-PR in improving functional exercise capacity and quality of life in people with COPD compared to usual care. Thus, home-PR is an option that could enable people whose lifestyles or geographical locations make attending a pulmonary rehabilitation centre difficult or who wish to socially distance to benefit from pulmonary rehabilitation.

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