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A 'whole of United Nations approach' to tackle antimicrobial resistance? A mapping of the mandate and activities of international organisations

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ABSTRACT The 2015 Wo

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Correspondence to Dr Didier Wernli; didier.wernli@unige.ch The 2015 World Health Organization Global Action Plan and other international policy documents have stressed the need for a 'whole of United Nations approach' in addressing antimicrobial resistance (AMR). As several years have passed, the goal of this paper is to take stock of the current role, mandate, and activities of international organisations and other global stakeholders on AMR. Relevant information is identified through a web-based search and a review of policy documents from international organisations. Based on the assessment of 78 organisations, 21 have AMR-specific activities in the broader sense, although for many of these organisations, their involvement is limited in scope, and 36 have AMRsensitive activities reflecting the wide scope of AMR. An interdisciplinary framework based on six relevant challenges of global collective actions regarding AMR as well as the main functions of international organisations in global governance is used to organise the findings into several 'clusters'. AMR is not a priority for many international organisations, but some of them can leverage current efforts to tackle AMR while contributing to their core agenda. Overall, a 'whole of UN approach' to AMR within the framework of Sustainable Development Goals is critical to move the global governance of AMR forward.

INTRODUCTION

Antimicrobial resistance (AMR) represents a formidable challenge for global governance. The cross-border transmission of AMR and the transnational nature of several drivers of antimicrobial use imply that no country can solve the problem on its own. International collaboration including bilateral and multilateral efforts is thus critical to successfully address AMR. In addition, the determinants of AMR are both social and ecological, therefore extending beyond the health sector; they are also related to sustainable development.¹ In 2015, the World Health O adopted a Global Action Plan on AMR (WHO GAP AMR).² This was followed by similar documents by the World Organization for Animal

WHAT IS ALREADY KNOWN ON THIS TOPIC

- ⇒ Addressing antimicrobial resistance (AMR) requires an integrated approach across sectors as recognised by the One Health approach.
- ⇒ The 2015 WHO Global Action Plan recognises the need for interorganisational collaboration and specifically identify several international organisations that could contribute to address the topic.

WHAT THIS STUDY ADDS

- ⇒ This study found that, in addition to core international organisations such as the WHO, the World Organization for Animal Health, and the Food and Agriculture Organization of the United Nations, several international organisations have activities that specifically address AMR, and many have activities that relate to AMR.
- \Rightarrow The findings also identify several collaborations between international organisations.

HOW THIS STUDY MIGHT AFFECT RESEARCH, PRACTICE AND/OR POLICY

- ⇒ As the global governance of AMR is embedded within wider processes of global health governance, the new findings emphasise the relevance of leveraging existing resources, mechanisms and institutions rather than reshuffling the governance of AMR.
- ⇒ A 'whole of United Nations approach' in an essentially polycentric governance system provides a realistic approach to address AMR as part of wider efforts towards sustainable development.

Health (OIE) and the Food and Agriculture Organization of the United Nations (FAO).³⁴ In 2016, a political declaration was adopted by Heads of States at the United Nations General Assembly (UNGA).⁵

The adoption of these soft law instruments marked renewed efforts in terms of increasing the profile of AMR on the global political agenda and addressing AMR from an integrated perspective. Such vision was already present in the 2001 Global Strategy for the containment of AMR.⁶ However, the 2015 WHO GAP AMR expanded this thinking by framing AMR as a 'One Health' issue that cannot be addressed within one sector but requires an 'all-out effort' and 'whole-of-society engagement'.² The 2015 WHO GAP AMR offers recommendations not only for Member States and the WHO Secretariat but also for other partners/ stakeholders. At the global level, several recommendations focus on strengthening the Tripartite collaboration between WHO, OIE and FAO, but others cover the role of other international organisations (IOs) including the United Nations Children's Fund (UNICEF), the United Nations Educational, Scientific and Cultural Organization (UNESCO), the World Bank (WB), the United Nations (UN), Organisation for Economic Co-operation and Development (OECD) and the Codex Alimentarius Commission (CAC).

More than 5 years after the adoption of the 2015 WHO GAP AMR and the subsequent political declaration at the UNGA, it is time to take stock of the progress accomplished in implementing the vision of a 'whole of UN approach' on AMR. In this article, we assess the contribution of different IOs which are part of or related to the UN system to address AMR with a focus on activities undertaken since the 2010s and more specifically in the wake of the adoption of the 2015 WHO GAP AMR. This aligns well with the waves of attention found by Overton et al.⁷ This assessment is based on an online collection of relevant sources. We report about the level of involvement of IOs in the global governance of AMR and organise these findings in several broad clusters to address AMR. Based on our findings, we discuss the pros and cons about expanding participation of IOs in AMR. We evaluate the potential future involvement of IOs and other global health stakeholders regarding AMR.

METHODS

Patient and public involvement

This study is based on online search of publicly available webpages and documents. No patients or members of the public were involved in the design of the study.

Selection of IOs

The UN system, which comprises the UN and several related organisations, is the largest intergovernmental organisation in the world. To select organisations, we used the UN system chart (https://www.un.org/en/pdfs/18-00159e_un_system_chart_17x11_4c_en_web.pdf). From this chart, we extracted the name of 105 organisations and entities in an Excel sheet. We included 78 organisations in the analysis including the 5 principal organs of the UN, the 15 specialised agencies (ie, autonomous organisations whose work is coordinated through Economic and Social Council (ECOSOC) and Chief Executive Board (CEB)), 23 department and offices of secretariat, 11 funds and programmes, 8 related organisations linked to the UNGA or the Security Council, 6 research and training organisations related to ECOSOC

and/or the UNGA, 2 subsidiary organs, 6 'other entities/bodies' and 1 mechanism for interorganisational collaboration (CEB). Furthermore, we added manually OIE which is not part of the UN system but is a member of the Tripartite. Because they are already represented by the main organs mentioned above, the subsidiary organs of the Security Council and the UNGA, the functional commissions and other bodies of ECOSOC, and the seven Regional Commissions of ECOSOC/Security Council were not considered for this analysis (some of these bodies are covered in online supplemental material 1). In addition, some other non-UN organisations were assessed but were not integrated in the main analysis (see online supplemental material 2).

Identification of activities

To identify IOs' activities on AMR, we conducted an online search with AMR and derivatives (eg. antibiotic resistance, antimicrobial resistance, resistant infections) and the name of the IO as the main keywords. As the focus of this article is on the global level, we limited our search to websites associated with global headquarters of each organisation. In addition, we browsed the websites of the relevant organisations to search for activities on AMR. In addition to written information directly available from websites, we downloaded the documents directly linked to the pages we visited. We qualitatively analysed these documents which encompass a wide range of sources such as policy and technical reports, normative guidance (eg, resolutions, action plan, strategy, etc), procedural documents (eg. agenda and meeting reports) with the software ATLAS.ti V.9 (Windows version) for relevant content on AMR activities (cf. the paragraphs below for the categories used). During the analysis, further documents were identified and collected through a snowball approach. In total, we collected 566 documents published by IOs on AMR from 1949 to March 2022 (a list of all documents considered for this analysis is available as a online supplemental material 3). While the core information used in this study was directly extracted from websites and recent documents, the broader information collected from the database of documents provided relevant information about the context of organisations' engagement in addressing AMR.

We distinguished between activities that are specific to tackle AMR and those that may influence AMR.⁸ These 'AMR-sensitive' activities such as the use of vaccines are not designed to tackle AMR directly but can contribute to address some aspects of the challenge. Because AMR touches on many different sectors, virtually all organisations of the UN system can have AMR-sensitive activities. For example, poverty and security might be upstream drivers of AMR. For this article, we included only AMRsensitive activities that are related to AMR in terms of six challenges of global collective action. However, to identify organisations that might contribute to tackle AMR through specific and actionable entry points, we assessed the mandate and activities of these organisations in the

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Box 1 Challenges of global collective action related to AMR

As addressing AMR requires global collective action, evaluating the mandate and activities of IOs requires understanding of strategic approaches and main policy goals. There have been several international policy frameworks to tackle AMR including a comprehensive WHO global strategy for the containment of AMR published in 2001.⁶ Renewed efforts resulted in the adoption by WHO Member States of the 2015 WHO GAP AMR. The GAP has been legitimised by the UNGA as the main instrument for guiding approaches for tackling AMR. FAO and OIE have published their own action plan based on the 2015 WHO GAP AMR.³⁴ IACG-AMR has developed its own framework.⁵⁰ Based on the literature, one can identify six areas/challenges of global collective action regarding AMR.⁵¹ The first is infection prevention which aims to limit infectious diseases in the first place. Relevant interventions include immunisation, sanitation, hygiene as well as infection prevention in healthcare settings. The second corresponds to all surveillance activities related to AMR (human health, animal health and the environment) and antimicrobial use to inform actions based on evidence. The third is conservation of antimicrobial effectiveness that aims to lower selective pressure and in turn reduces the emergence of AMR. This challenge mainly relates to interventions that target overuse and misuse of antibiotics and also limit antibiotic pollution in the environment. As transmission of AMR is an important factor driving the problem,⁵² the fourth challenge is about containing the different pathways of dissemination of AMR once it has emerged. The fifth is strengthening access to health including but not limited to access to drugs as inequalities are one of the most central challenges regarding AMR policy and global health.⁵³ Strengthening access to health can help not only to reduce the burden of infectious diseases but also improve how we use antibiotics. Finally, a sixth challenge of global collective action is innovation which designates all interventions and policies to support the development of health technologies with the direct goal to address AMR.

AMR, antimicrobial resistance; FAO, Food and Agriculture Organization; GAP, Global Action Plan; IACG, Interagency Coordination Group; IOs, international organisations; OIE, World Organization for Animal Health; UNGA, United Nations General Assembly.

health, agriculture/food and environmental sectors, all of which are important dimensions of AMR.⁹ We looked for these activities by searching the name of the organisation and relevant keywords regarding the abovementioned areas (eg, infectious diseases, malaria, tuberculosis (TB), HIV-AIDS) and by understanding how this relates to the main policy priorities regarding AMR. A summary of activities was created for each organisation.

Analytical framework

To help assess an organisation's role in tackling AMR, we built a multidimensional framework. First, we drew on the literature to identify six challenges of global collective action regarding AMR.^{10 11} Accordingly, AMR is not only an issue of classic strategies to tackle infectious diseases, that is, surveillance, prevention, containment, innovation, but also of conservation of a natural resource like several environmental challenges and of access to health technologies (cf. box 1 for definition of these challenges

of global collective action). Second, governance covers several functions. Based on the literature in international relations, we distinguished between different types of functions of IOs and other stakeholders in global governance (cf. box 2 for definition of these functions). Third, we defined several sectors that are relevant to address AMR including human health, agriculture, food production and the environment. Fourth, as AMR is pervasive regarding infectious diseases, but with highly variable scope and impact, we included activities to tackle AMR in specific diseases such as HIV-AIDS, malaria and TB in addition to common bacterial, viral and fungal infections. Fifth, we considered the population that can be targeted by different organisations (eg, children, migrants). Taken together, these five dimensions define a framework for evaluation of stakeholders' action on AMR.

To classify the involvement of IOs in the global governance of AMR, we defined five categories from core to distant organisations. Core organisations have received a mandate on AMR from their Member States. This mandate is associated with both AMR-specific and AMRsensitive activities. Core organisations have a large scope of activities regarding several of the five parameters above. Involved organisations do not necessarily have a mandate on AMR. However, they have some AMR-specific activities and may also have AMR-sensitive activities. Related organisations have only AMR-sensitive activities which can be related to the six challenges of global collective actions defined above. Relevant organisations have no mandate on AMR but have usually more distant AMR-sensitive activities in areas such as sustainable development, training and coordination. Finally, some organisations are distant as their mandate, priorities and activities are currently far removed from AMR.

RESULTS

Overall results

From the analysis conducted between January and September 2018 and updated in autumn 2021 on 78 international organisations, 21 organisations have AMRspecific activities in the broader sense, although for many of these organisations, their involvement is limited in scope, and 36 have AMR-sensitive activities reflecting the wide scope of AMR (the results based on the identification of relevant AMR activities are summarised in online supplemental material 4). A limited number of organisations have received a direct mandate on AMR from their Member States (WHO, OIE, FAO, the Joint United Nations Programme on HIV/AIDS (UNAIDS)) and more recently the United Nations Environment Programme (UNEP). Among these organisations, we identified WHO, OIE, FAO and UNAIDS as core organisations because they are active on several challenges of global collective action and exert multiple functions in the global governance of AMR. WHO is the main organisation in human health but also develops activities on food safety. OIE is the main organisation in animal health

Box 2 Functions of IOs and other global health actors

The IACG-AMR framework identified five levers (eg, awareness and capability building, measurement/surveillance, funding and financial incentives, policy and regulations, championing and piloting)⁵⁰ while the 2015 WHO GAP AMR does not cover specifically these areas. Based on the academic literature on the functions of IOs, we propose six core functions that form a continuum from purely technical support to countries to more policy-oriented and normative activities.^{54 55}

Knowledge synthesis

IOs often act as a knowledge broker which means that they seek to harness scientific research not only to produce evidence-informed policies, but also to legitimise policy choices, depoliticise action and ensure their institutional survival.⁵⁶ While the role of IOs in knowledge brokering predominantly involves the synthesis of existing knowledge and the classification of information,⁵⁷ it also involves the collection and production of new scientific data.⁵⁸ IOs also create and/or endorse new concepts that are then disseminated in the international community.⁵⁹ A good example is the coining of the concept of sustainable development by the World Commission on Environment and Development.

Normative and political forum

One of the core functions of IOs is normative.⁶⁰ This function is facilitated by their unique power for convening people and for providing a forum for negotiations among states. IOs are involved in all steps of the norm cycle: emergence, crystallisation and diffusion of norms.⁶¹ The UN system plays a key role in the codification of norms.⁶² Normative activities in IOs range from the creation of shared technical standards to the making of international law. As IOs have an authoritative role, they are also involved in the formulation of solutions to tackle international challenges.⁶³

Advocacy, awareness raising and capacity building

IOs are often involved in raising awareness about some problems that states are not aware of or insufficiently addressing. The focus here is on the diffusion and dissemination of information rather than the production of new knowledge. When common rules and policies are agreed at the international level, IOs can also play a key role in advocating change to their constituencies either by highlighting best practices or by providing technical guidance to support countries' implementation. Their capacity to draw on a vast expertise is often call upon to build capacities in countries. This is especially important in LMICs where capacities are often lacking to implement agreed policies. Some IOs specialise in training and capacity building (eg, UNITAR) and many IOs propose training activities.

Funding, procurement and delivery

Some IOs have mechanisms for project funding and delivery. For example, 'on average, WHO buys approximately US\$700 million worth of goods and services every year to maintain operations in the field and to react to upcoming and recurring demands'.⁶⁴ Funding and procurement activities from IOs and related organisations can be used to advocate changes and orient policies. A relevant example is the structural adjustment programmes conducted by the International Monetary Fund and the World Bank.⁶⁵

Oversight, monitoring and evaluation

Another function of IOs is to monitor and oversee the implementation of any agreement or shared policies. Monitoring and evaluation are essential to diffuse best practices, to promote learning and to benchmark countries' performance while oversight is key to foster compliance. Several mechanisms are possible from peer review to self-reporting, to technical experts.⁶⁶ In some cases, IOs play a key role in devising the oversight instrument through survey/questionnaire

Continued

Box 2 Continued

and/or in conducting on-site evaluation. A classic example is evaluation conducted by the International Atomic Energy Agency.

Coordination/interplay management

An important function of IOs is to coordinate work across different areas. Traditionally, IOs' coordination mandate has revolved around specific issues. A good example is the UN Office for the Coordination of Humanitarian Affairs which was created to improve the coordination between several humanitarian actors. In addition, some organs of the UN (eg, ECOSOC, UNGA) have a broader coordination mandate. With the growing interdependence between sectors, a growing coordination role is about the management of interplay between different sectors to improve policy coherence within the context of 'whole of UN system' response expressed by the Sustainable Development Goals.⁶⁷ Organisations that are representative of diverse groups are important.⁶⁸

AMR, antimicrobial resistance; ECOSOC, Economic and Social Council; GAP, Global Action Plan; IACG, Interagency Coordination Group; IOs, international organisations; LMICs, low/middle-income countries; UN, United Nations; UNGA, United Nations General Assembly; UNITAR, United Nations Institute for Training and Research.

and FAO has an important role regarding AMR in foodproducing animals and food systems. UNAIDS's activities on AMR relate to planning, advocacy, communication and capacity building in HIV-AIDS.

Involved organisations on AMR encompass 17 organisations that have had AMR-specific activities and, for some of them, AMR-sensitive activities. The United Nations Population Fund (UNFPA), the only one which has a primary health mandate, works on infection prevention, access and conservation of antimicrobials mainly in maternal health and HIV-AIDS. UNEP has recently become involved in tackling AMR in the environment. Beyond One Health, finance, trade and intellectual property organisations have been involved to some extent in tackling AMR. These include the WB, the United Nations Conference on Trade and Development (UNCTAD), the World Trade Organization (WTO) and the World Intellectual Property Organization (WIPO). Another relevant area is migration and refugees where the International Organization for Migration (IOM), the United Nations High Commissioner for Refugees (UNHCR), and the United Nations Relief and Works Agency for Palestine Refugees in the Near East (UNRWA) have had AMRspecific activities.

Most organisations in the category 'involved' are active in specific challenges of global collective action and/or functions of global governance and/or specific countries/populations. For example, the WB is not only interested in AMR in low/middle-income countries (LMICs) but also covers the economic aspect of AMR. The United Nations Development Programme and UNICEF are involved in access to antimicrobials. This suggests that involved organisations can address important drivers of AMR, but they do so in a way that are generally restricted to one or a few dimensions of the issue (eg, focus on a

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specific disease or population, functions or limited to one challenge of global collective action). In most cases, we found that IOs work at the project or programme levels without a mandate from their Member States. In some cases, they collaborate directly with an organisation that has a core mandate on AMR. This is for example the case of the International Telecommunication Union with WHO.

Among the 78 organisations assessed, 16 are currently 'related'. While they have a mandate in different areas including human rights such as the Human Rights Council and the Office of the United Nations High Commissioner for Human Rights, development such as the United Nations Industrial Development Organization (UNIDO) and the United Nations Research Institute for Social Development (UNRISD) or crime such as the United Nations Interregional Crime and Justice Research Institute (UNICRI) and the United Nations Office on Drugs and Crime, they undertake AMR-sensitive activities related to the six challenges of global collective action. Most of them have activities in infection prevention such as immunisation, sanitation, and hygiene and in access to medicines. We identified 15 organisations in the category 'relevant'. Their mandate includes economic and social development such as the United Nations Department of Economic and Social Affairs and the ECOSOC, education and training, such as UNESCO and the United Nations System Staff College, security such as the United Nations Security Council, or coordination such as the CEB. Lastly, 26 of the 78 organisations assessed are most likely distant from AMR. A quarter of them have a mandate in security, while others have their mandate in different areas such as marine affairs and internal management.

Finally, our analysis of the result led us to identify several mechanisms for intersectoral collaboration between (1) human health, animal health, agriculture and food; (2) human/animal health, food and trade; and (3) public health, intellectual property and trade. The roles of these mechanisms in addressing AMR are summarised in table 1. Some of these mechanisms such as the CAC are old. For several mechanisms, AMR is only part of their mandate. Other mechanisms such as WHO Advisory Group on Integrated Surveillance of Antimicrobial Resistance (AGISAR) were created specifically to address AMR. WHO is engaged in most of these mechanisms for intersectoral collaboration. Overall, the main areas of collaboration within these mechanisms are surveillance, conservation and infection prevention. These mechanisms are mainly dedicated to the management of interplay regarding knowledge synthesis and norms, and, to a lesser extent, policies including support to the development and implementation of national action plans and monitoring and evaluation.

Results in the six challenges of global collective action

While the findings above provide an overview regarding the work of each organisation or mechanisms for intersectoral organisation, using them in isolation offers limited insights into the global governance of AMR. A relevant approach is thus to understand involvement from these organisations in terms of the six challenges of collective action.

The global responsibility for surveillance of AMR and antimicrobial use is shared between several IOs. In human health, WHO activities on surveillance were already important in the 1960s and again in the 1980s.¹² In recent times, WHO set up the Global Antimicrobial Surveillance System in 2015, based on voluntary participation of countries. Several mechanisms are dedicated to pathogens/diseases such as Salmonella spp ('WHO Global Salm-surv'), Neisseria gonorrhoeae ('WHO Gonococcal Antimicrobial Surveillance Programme'), malaria (Worldwide Antimalarial Resistance Network) and TB (the Global Project on anti-Tuberculosis drug resistance surveillance). The responsibility for addressing global surveillance of drug resistance in HIV is shared between WHO ('WHO HIVresnet') and UNAIDS. Regarding antimicrobial use, WHO has issued technical guidance on surveillance for a long time, but there is currently no global mechanism to monitor antimicrobial use. In animal health and agriculture, OIE and FAO are responsible for global surveillance with activities since the 1960s. OIE has set up an ad hoc group on AMR to develop standards on surveillance and support the collection of data in Member States. FAO created an Assessment Tool for Laboratory and AMR Surveillance Systems, the goal of which is to evaluate surveillance capacities regarding AMR. For a long time, WHO has also been involved in surveillance of AMR in animal health and set up AGISAR in 2008. AGISAR has supported the development of standards regarding the surveillance of AMR but also maintained a list of critically important antimicrobials in human health. In animal health, there is currently no established global surveillance mechanism. In terms of antimicrobial use, activities have been initiated by OIE which collects data on a yearly basis. Finally, there is currently a gap in terms of the surveillance of AMR in the environment.¹³ Scientific studies are currently the main source of data for environmental surveillance.

Covered by the third objective of the 2015 WHO GAP AMR and relevant for both human and animal health, infection prevention is primarily about AMR-sensitive activities. A variety of activities and collaborations are relevant to AMR including on water sanitation, immunisation, patient safety and biosecurity. In human health, WHO is the main organisation responsible for infection prevention with a dedicated unit and the support of several WHO collaborating centres. In the healthcare setting, the single most effective and inexpensive intervention is hand hygiene, which has been supported by the WHO hand hygiene campaign and other initiatives. There are also targeted activities regarding specific diseases in the community as infection prevention is the basis of public health.¹⁴ For example, UNFPA is mainly active on AMR with regard to HIV infection and maternal health. Some organisations are active in infection prevention in specific

Acronym of the organisation			
organisation			
CAC (FAO and WHO)	Human health, animal health, agriculture and food The CAC adopts guidelines and standards regarding the conservation of antimicrobials and the containment of AMR in food products and animal husbandry. ²⁰ The main standards on AMR are the 1993 'International Code of Practice for Control of the Use of Veterinary Drugs', ⁶⁹ the 2005 'Code of Practice to Minimize and Contain Antimicrobial Resistance', ⁷⁰ and the 2011 'Guidelines for Risk Analysis of Foodborne AMR'. ⁷¹ These standards are recognised by the Sanitary and Phytosanitary (SPS) Agreement at the WTO. While collaboration was mainly between WHO and FAO through the CAC, OIE was also included in the discussion. The work has been conducted by the 'Ad hoc Codex Intergovernmental Task Force on AMR' from 2007 to 2011 and since 2017.		
Tripartite collaboration 2010 (FAO, OIE and WHO)	The Tripartite FAO/OIE/WHO Collaboration was created in 2010 and entails a common recognition of the role and functions of each organisation. For example, the standards published for terrestrial and aquatic animals are recognised by WHO. ⁴³ Efforts have been done towards better integration between the IHR and PVS instruments, regarding the assessment of national capacities to detect and respond to global health threats. ⁷² In recent years, AMR has become one of the main areas for action of the Tripartite. In 2017, the FAO/OIE/WHO Tripartite Collaboration on AMR renewed their commitment to work together. ⁷³ In 2019, the Tripartite established a standing Tripartite Joint Secretariat with special governance arrangements.		
AGISAR (WHO)	The WHO AGISAR is an expert-based intersectoral AMR-specific mechanism that mainly addresses AMR in food-producing animals. The mission of AGISAR is to support WHO's effort 'to minimise the public health impact of AMR associated with the use of antimicrobial agents in all food-producing animals'. ⁷⁴ AGISAR covers surveillance but is also responsible for some aspect of conservation, for example, by establishing the list of critically important antimicrobials.		
INFOSAN (WHO and FAO)	Created in 2005 by FAO and WHO, the INFOSAN shares food safety information between human and animal health sectors. INFOSAN has reported on AMR.		
	Human/animal health, food and trade		
SPS Agreement (WTO)	The need to manage the interface between trade and animal health resulted in the adoption of an agreement in 1998 'to act in collaboration and to consult each other on questions of mutual interest, in particular those concerning the sanitary aspect of international trade in animals and products of animal origin and zoonoses'. ⁷⁵ The standards adopted by OIE, FAO and the CAC are non-binding but 'are specifically identified in the WTO Agreement on the Application of Sanitary and Phytosanitary Measures as the international benchmark texts for food'. ⁷⁶ There is no specific mechanism for AMR, but FAO and WTO prepared a joint publication on 'Trade and food standards' in which AMR is briefly mentioned. ⁷⁷		
	Public health, intellectual property and trade		
Trilateral Cooperation on Public Health, Intellectual Property and Trade (WHO, WIPO and WTO)	WHO has a health mandate regarding both access and innovation, which are situated at the interface between the human rights and intellectual property regimes. Following the adoption of the 'Global Strategy and Plan of Action on Public Health, Innovation and Intellectual Property', WHO, WTO and the WIPO created the 'Trilateral Cooperation on Public Health, Intellectual Property and Trade'. ²⁵ The topic of AMR has been covered during a technical symposium held in 2016. ⁷⁸ The scope of formal participation remains limited as the WTO Secretariat participates 'as observers in the WHO Intergovernmental Working Group on Public Health Innovation and Intellectual Property and in the WHO's International Medical Products Anti-Counterfeiting Task Force (IMPACT), as well as in the field of technical cooperation'. ⁷⁹		
	Overall coordination mechanism		
IACG-AMR (Convened by the UN Secretariat)	The adoption of the 2016 Political Declaration by the UNGA led to the creation of the ad hoc IACG-AMR to provide advice on how to make the global AMR agenda move forward. ⁵ IACG-AMR, which is co-chaired by WHO and the UN Secretariat, has a broad membership of IOs and other global health actors. Four objectives have been laid out in its working plan for 2017–2019 and the work has been divided in six subgroups. ⁸⁰ In 2017 and 2018, IACG-AMR commissioned works in several areas and produced its own framework of action which put emphasis on innovation. The final recommendation of IACG-AMR stressed the need for countries' progress, collaboration, sustainable response and innovation, and mechanisms for global governance (eg, the creation of a Global Leaders Group on Antimicrobial Resistance and an Independent Panel on Evidence for Action Against Antimicrobial Resistance). ⁸¹		
Global framework (WHO, FAO and OIE)	Following the adoption of WHO Resolution WHA 68.7, and further supported by the 2016 UNGA Political Declaration, an ongoing effort by the FAO/OIE/WHO Tripartite Collaboration on AMR to design an overarching intersectoral mechanism to tackle AMR is the 'global development and stewardship framework'. ⁷⁶ As the framework aims to balance innovation, access and conservation, '[it] is envisaged to form an 'umbrella' uniting different instruments', ⁷⁶ but its legal form still needs to be defined. Several rounds of consultations have been organised with Member States, but it is yet unclear whether this initiative has so far garnered enough support to move to the WHA.		
AGISAR, Advisory Gro	oup on Integrated Surveillance of Antimicrobial Resistance; AMR, antimicrobial resistance; CAC, Codex Alimentarius Commission; FAO,		

AGISAR, Advisory Group on Integrated Surveillance of Antimicrobial Resistance; AMR, antimicrobial resistance; CAC, Codex Alimentarius Commission; FAO, Food and Agriculture Organization; IACG, Interagency Coordination Group; IHR, International Health Regulations; INFOSAN, International Food Safety Authorities Network; IOs, international organisations; OIE, World Organization for Animal Health; PVS, Performance of Veterinary Services; UN, United Nations; UNGA, United Nations General Assembly; WHA, World Health Assembly; WIPO, World Intellectual Property Organization; WTO, World Trade Organization.

contexts such as humanitarian crises (United Nations Office for the Coordination of Humanitarian Affairs), maternal health (UN-Women), refugees (UNHCR) or in relation to work activities (International Labour Organization). Other relevant organisations include those active in increasing immunisation such as UNICEF with longstanding collaborations between WHO and UNICEF to improve immunisation coverage in LMICs. UNICEF and UN-Water are the main organisations dedicated to improving sanitation with dedicated programmes in LMICs. UN-Habitat also focuses on infection prevention by working on sanitation, water and urban health. Other organisations are active in infection prevention issues through research and policy analysis (UNRISD) and training (United Nations University). In animal health, core institutions include those with a direct agriculture/ animal health mandate (OIE, FAO). However, WHO is also involved in food safety. The CAC, an interorganisational collaboration between FAO/WHO, focuses on internationally recognised food standards. Finally, the WTO rules that govern the standards that are compatible with international trade of animal and food products also impact infection prevention (eg, the agreement on the Application of Sanitary and Phytosanitary Measures (SPS Agreement)).

Conservation corresponds to the fourth objective of the 2015 WHO GAP AMR and combines educational and regulatory strategies to optimise the use of antimicrobials. In human health, WHO provides guidance about the best practices for the rational use of drugs including antimicrobial stewardship. UNAIDS has similar functions for addressing drug resistance in HIV. WHO also raises awareness about AMR including through the provision of education material and the World Antibiotic Awareness Week. AGISAR performed repeated assessments of critically important antimicrobials for human medicine. In 2019, WHO further released a categorisation of antibiotics as access, watch and reserve to maximise their good use.¹⁵ In animal health and agriculture, the mandate on conservation is shared between FAO and OIE. OIE set standards through the terrestrial and aquatic animal health codes. With no global mechanism for conservation, OIE reported that 35 countries are allowing antibiotics for growth promotion in 2018.¹⁶ WHO has also been involved in non-human use of antimicrobials including regarding foodborne pathogens since the end of the 1990s. WHO collaborates with FAO through the CAC since the 1960s. Several organisations have been involved in some activities related to conservation of antimicrobial effectiveness including UNFPA on TB and HIV, UNICEF in children, UNHCR and UNRWA in refugees. Furthermore, some activities to reduce the issue of counterfeit medicines from UNICRI and United Nations Office for Project Services are indirectly related to the conservation of antimicrobial effectiveness.

Against the backdrop of globalisation, international dissemination of AMR is an externality of travel and trade.¹⁷ Issues related to the **containment of AMR** are not well developed in recent global policy documents and we found limited activities related to containment. In human health, WHO has issued guidelines for control of some highly drug-resistant bacteria such as carbapenemresistant Enterobacteriaceae,¹⁸ and specific diseases such as HIV-AIDS, malaria and TB.¹⁰¹¹ In addition, it has been argued that the broad scope of the International Health Regulations encompasses AMR, but guidance regarding their applicability has been limited.¹⁹ In animal health and food production, some guidelines have been issued by FAO and OIE. Furthermore, any restriction on international trade is governed by the SPS Agreement at WTO which regulates the rules and processes that countries need to respect when adopting standards that might

represent barriers to trade. Countries' policies to restrict imports of products contaminated with AMR appear to have been limited.²⁰ A few activities from the International Civil Aviation Organization and the International Maritime Organization are relevant to the containment of AMR.

The main actors classically dedicated to innovation including research and development of new drugs and other relevant technologies are multinational companies. Given the market failures regarding the development of novel antibiotics,^{21 22} innovation for technologies to tackle AMR has received growing attention in global governance since the mid-2000s. There has been more emphasis regarding human health than animal health as the latter area is seen as more profitable for pharmaceutical companies. Innovation is well represented in the 2015 WHO GAP AMR (objective 5). Several organisations contribute to this objective at the global level. In 2011, WHO created the 'Consultative Expert Working Group on Research and Development: Financing and Coordination' that evaluated innovative approaches to incentivise further the development of new health technologies. This led to the creation of the Global Observatory on Health R&D. In 2017, WHO also issued a list of priority pathogens for the development of new antibiotics.²³ Against the background of the development of product development partnerships and several initiatives on AMR, a Global Antibiotic Research & Development Partnership was created in 2016 to address the innovation gap in AMR.²⁴ As intellectual property has been the main mechanism to reward the research and development of new technologies, both WTO and WIPO have an upstream influence on research and development of technologies for AMR. The topic has been discussed within the Trilateral cooperation on public health, intellectual property and trade.²⁵

WHO has historically been the main IO dealing with access to health. Its programme on essential medicines started in 1977.²⁶ Both objectives 4 and 5 of the 2015 WHO GAP AMR cover aspects related to access to medicines. Several global stakeholders contribute to access to medicines with mainly AMR-sensitive activities. While WIPO is responsible for intellectual property, the most important international agreement regarding access is the Agreement on Trade-Related Aspects of Intellectual Property Rights (TRIPS) at WTO. The current regime allows for some flexibilities (eg, compulsory licensing) for drugs of public health importance. UNAIDS has advocated access to antiviral drugs against HIV-AIDS since its creation in 1996. Since 2010, interorganisational collaborations have taken place between WHO, WTO and WIPO under the 'Trilateral cooperation on public health and intellectual property' but have not led to the establishment of a specific international mechanism dedicated to improving access to antimicrobials. Other relevant AMRspecific activities to facilitate access have been undertaken at the programme/project level by UNCTAD, UNHCR and UNICEF for specific populations. Finally, the Human Rights Council and UNIDO have activities on access to medicines that indirectly relate to AMR. In 2015, the UN Secretary-General convened a 'High Level Panel on access to medicine'. In its final report published in 2016, the Panel reiterated the importance of the flexibilities within the TRIPS agreement,²⁷ as did the 2018 UNGA political declaration on TB.²⁸

DISCUSSION

Our assessment provides insights about past and current efforts of IOs to tackle AMR with a focus on activities conducted since the 2010s. First, it suggests that addressing a complex issue such as AMR is shaped by different international actors. While there is a core of IOs with a myriad of activities on AMR, several IOs without a formal mandate on AMR are also contributing to address the challenge, although the respective impact of different regimes on AMR is poorly understood (figure 1 summarises the current levels of involvement from IOs on AMR). Second, despite the call from WHO and the UNGA to expand participation in the global governance of AMR, our assessment reveals a mixed picture. On the one hand, several activities were undertaken over the past years and several IOs have become involved on AMR. The WB and UNEP are the best examples. On the other hand, the level of involvement of other IOs remains often limited to the project level. While the environment is a critical dimension to AMR, it remains under-represented in the global governance of AMR.7 A possible explanation for the growing but still limited involvement of UNEP is that it is faced with several other pressing global challenges. Overall, this suggests a certain amount of 'output effectiveness' in current effort to govern AMR. While the goal of this article is not primarily to measure the impact of the current governance system, it appears that the current governance system has managed to raise awareness about AMR and made progress to address several challenges of collective action (eg, surveillance and conservation) and several governance functions (eg, monitoring and evaluation).²⁹

Several factors may explain the current situation. From a formal perspective, an important obstacle is that most IOs have not received a formal mandate from their Member States in the respective organisation. So, the lack of engagement primarily reflects the weak will of States, and the current subeffective organisation of the international system when it comes to addressing complex challenges such as AMR. Moreover, the current situation stresses the discrepancy between engagement at the UNGA by Head of States and further engagement in different organisations by ministries. This problem of policy coherence is hampering collective action on AMR.³⁰ Insights on the international policy process suggests that several other factors might play a role in whether an issue get addressed or not by the international community. Getting stakeholders involved require making the issue tractable and actionable.^{31 32}

For example, there is ongoing discussion in the literature about the relevance of developing environmental surveillance of AMR.³³ Given the existence of multiple pressing global issues and the competition for attention and resources, one must demonstrate that addressing AMR will reinforce issues that are central to the mandate of organisations whose primary mandate is not AMR.³⁴ For example, organisations working on specific diseases are more likely to provide expertise and logistical support to support broader AMR activities in LMICs (eg, resistance testing for extensively drug-resistant (XDR) TB in remote areas), if these activities do not lead to a reduction of support to their core mandate. More attention is thus needed on the issue of co-benefits to show how AMR can strengthen the core mandate of other organisations.³⁵

Addressing several challenges regarding AMR depends to various extent on the capacity of organisations with different and sometimes conflicting mandate to collaborate. While this stresses the importance of interorganisational collaboration to address AMR,^{36 37} current efforts take place within 'a history of heterogeneous, often uncoordinated, and at times conflicting reform efforts, whose legacies remain apparent today'.³⁸ The WHO Secretariat frequently reports to the World Health Assembly about its collaboration within the UN system and with other intergovernmental organisations.³⁹ In addition, specific mechanisms ranging from soft to hard law (eg, an international treaty) are needed in specific areas as it has been discussed in the literature.^{40–42} While collaboration between human and animal health is not new as illustrated by the CAC, the framing of AMR as a One Health issue in the 2015 WHO GAP AMR has supported collaboration between human, animal and environmental health.^{43 44} Collaboration ranges from sharing information to avoid surprise, to raising awareness and finally to sharing of resources and decision-making power.⁴⁵ Moreover, the creation of a Tripartite Joint Secretariat on Antimicrobial Resistance in 2019 signals an impetus to move to joint management of priorities and decision-making. Finally, interorganisational collaboration extends beyond the UN system as demonstrated with Global Antibiotic Research and Development Partnership and other public-private partnerships which have a critical role in funding and delivery of essential medicines in LMICs (table 2 provides an assessment regarding several global health stakeholders). The engagement with global stakeholders is necessary,⁴⁶ but raises questions regarding transparency and accountability of governance processes.⁴⁷ Overall, a better understanding of the resources devoted to intersectoral organisation as well as the practical barrier to collaboration is needed. Past experiences in other issues such as HIV-AIDS, tobacco control or climate change suggest that there are potential benefits and costs (eg, increasing conflict) in using a collaborative approach.

Our findings contribute to the debate about how to govern AMR globally. Not surprisingly, the analysis of current arrangements for the global governance of AMR reveals that the main responsibilities for addressing AMR

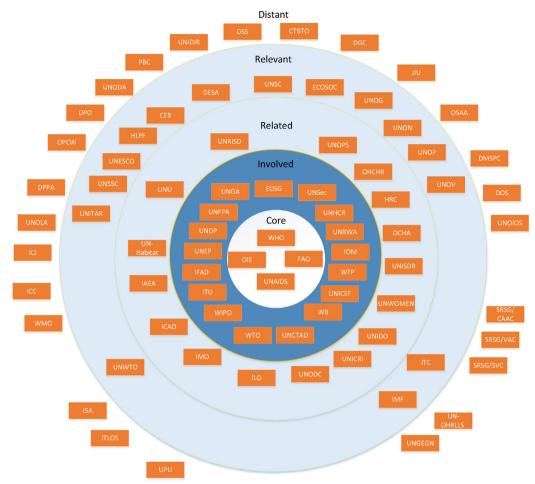


Figure 1 Current levels of IOs' involvement on tackling AMR. AMR, antimicrobial resistance; CAAC, Children and Armed Conflict; CEB, Chief Executive Board; CTBTO, Comprehensive Nuclear-Test-Ban Treaty Organization; DESA, Department of Economic and Social Affairs; DGC, United Nations Department of Global Communications; DPPA, United Nations Department of Political Affairs: EOSG, Executive Office of the Secretary-General: DMSPC, Department of Management Strategy, Policy and Compliance: DOS. Department of Operational Support: DPO. Department of Peacekeeping Operations: ECOSOC, Economic and Social Council; FAO, Food and Agriculture Organization; HLPF, High-Level Political Forum on sustainable development; HRC. Human Rights Council; ICAO, International Civil Aviation Organization; IAEA, International Atomic Energy Agency; ICC, International Criminal Court, ICJ, International Court of Justice; IFAD, International Fund for Agricultural Development; ILO, International Labour Organization; IMF, International Monetary Fund; IMO, International Maritime Organization; IOM, International Organization for Migration; ITC, International Trade Center; ITLOS, International Tribunal for the Law of the Sea; ITU, International Telecommunication Union; ISA, International Seabed Authority; JIU, Joint Inspection Unit; OCHA, Office for the Coordination of Humanitarian Affairs; OHCHR, Office of the United Nations High Commissioner for Human Rights; OHRLLS, United Nations Office of the High Representative for the Least Developed Countries, Landlocked Developing Countries and Small Island Developing States OIE, World Organization for Animal Health; OPCW, Organisation for the Prohibition of Chemical Weapons; OSAA, Office of the Special Adviser on Africa; PBC, Peacebuilding Commission; SRSG, Special Representative of the Secretary-General: SVC. Sexual Violence in Conflict: UNAIDS. Joint United Nations Programme on HIV/AIDS: UNCTAD. United Nations Conference on Trade and Development: UNDP. United Nations Development Programme: UNDSS, United Nations Department for Safety and Security; UNEP, United Nations Environment Programme; UNFPA, United Nations Population Fund; UNGA, United Nations General Assembly; UNGEGN, United Nations Group of Experts on Geographical Names; UNHCR, United Nations High Commissioner for Refugees; UNICRI, United Nations Interregional Crime and Justice Research Institute; UNIDIR, United Nations Institute for Disarmament Research; UNIDO, United Nations Industrial Development Organization; UNISDR, United Nations Office for Disaster Risk Reduction; UNITAR, United Nations Institute for Training and Research; UNODA, United Nations Office for Disarmament Affairs; UNODC, United Nations Office on Drugs and Crime; UNOG, United Nations Office at Geneva; UNOIOS, United Nations Office of Internal Oversight Services; UNOLA, United Nations Office of Legal Affairs; UNOP, United Nations Office for Partnerships; UNOPS, United Nations Office for Project Services; UNOV, United Nations Office at Vienna; UPU, Universal Postal Union; UNRISD, United Nations Research Institute for Social Development; UNRWA, United Nations Relief and Works Agency for Palestine Refugees in the Near East; UNSC, United Nations Security Council; UNSec, United Nations Secretariat; UNSSC, United Nations System Staff College: UNU. United Nations University: UNWTO, World Tourism Organization: VAC, Violence against Children WB, World Bank; WFP, World Food Programme; WIPO, World Intellectual Property Organization; WMO, World Meteorological Organization; WTO, World Trade Organization.

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Other IOs and mechanisme OECD, ISO, EU, ICRC, TATFAR The international system has seen a proliferation of used and the seen a proliferation of used in the support region. ECOL aggregates data there while other work at the regional level (e.g. EU while bell will write work at the regional level (e.g. EU while other IOS and PLAME, JPAMR) supports research on AMR. OECD has also published reports on AMR. Global health initiatives Global health initiatives Global health initiatives Global health initiatives Global health initiatives are hybrid organisations which mix elements of the private and public sector. They are targeted funding and health technologies in LMICs. ⁵⁰ Global Hundt in Fight ADS, Tuberculosis, and Mataris, GAW. Global health initiatives are hybrid organisations shore Global mealth initiatives are hybrid organisations which mix elements of the private and public sector. They are targeted funding and health technologies in LMICs. ⁵⁰ Tuberculosis, and Mataris, GAW. A central innovation in global health the sector organisations such as GAV have been reduction base that the sector organisations active in global funditin mix elements of the private and public sector. They are targeted funding and nealth technologies in LMICs. ⁵⁰ They take different forms such as a finance mechanism with organisations active in global funditin mixel (SARDP; MPP Whith Methods active) mixel sector organisations active in global funditin mixel sectors the prove the finant introl (SARDP mixel sectors) Finant Mixel sectors mixel sectors the prove the sector of sectors mixel sectors that pose the most significant threat to health infectors that pose the most significant threat to health infectors that pose the most significant mixel sectors to develop action of the proving in sectors to develop fundition (GARDP privath infectore mital, sectors mans and have developed form infectors	Table 2 Actors of the global health system and their role on AMR			
Bilateral development cooperation agencies. Ministris of rorigin Atlans; Ministris of Neath are the traditional actors in global health. National, and comment time. While the second seco	Types of actor and examples	Role in the global health system	Role in the global governance of AMR	
OECD, ISO, EU, ICRC, TATFAR Among these organisations, some are active at the global AMR. For example, the EU adopted plans on AMR. ³ evel while others work at the regional level (e.g., EU which adopted a strategy on global health). With the European region, EOCD agregates data from EU countries on AMR. JPIAMR supports research on AMR, OECD has also publicated reports on AMR. ³ Global health initiatives Global Fund to Fight ADS, Tuberciolesia, and Mataria, GAV. A central innovation in global health governance has been the development of new mechanisms to divers trageted funding and health technologies in LMICs. ⁴¹ Global health initiatives are hypola specific diseases (Global Fund to Fight ADS, Tuberciolesia, and Mataria; GAV. Unitad; GARDP; MPP A central innovation in global health technologies in LMICs. ⁴¹ Global health initiatives are hypola specific diseases tradecine start from HUCs to subisidia access to endicine start from HUCs. The were created to address major communicable diseases the editivery of health technologies, medications and partnership which aims to research and develop and mans and have developed strong expertise in the delivery of health technologies, medications and partnership which aims to research and develop action to the increase of funding in global health in the larget products such as drugs for neglected tropical diseases. These organisations have been endowed with large mans and have developed strong expertise in the delivery of health technologies, medications and partnership which aims to research and divelop new tradication. Rocketeller Foundation, Nacketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller foundation, Rocketeller	Bilateral development cooperation agencies; Ministries of Foreign Affairs; Ministries of	are the traditional actors in global health. National, and to some extent, international public health depends on Ministries of Health. Donor countries are involved in development assistance for health through their bilateral development cooperation agencies. Recently, as health has gained political prominence as 'soft' power, Ministries of Foreign Affairs have been increasingly	policies and implement them. ⁶ The responsibility to address AMR usually spans several ministries including health, trade and economy. Since the adoption of the WHO GAP AMR in 2015, national governments have developed national action plans to combat AMR. There is however a gap in implementation. In this regard, national and subnational actors are ultimately the ones that need to be involved and convinced about the need and urgency of implementing AMR policies through a coordinated multisectoral and multilevel approach where	
Drugs for Neglected Diseases; Global Fund to Fight ADS. Tuberculosis, and Materia; GAV; Unitadi; GARDP; MPPDens the development of new mechanisms to deliver targeted funding and health inticheologies in LMICs. The routed in the othologies in LMICs and the carbonic of the private and public sector. They are there created to address major communicable diseases in the routed to address major communicable diseases in the creation of GARDP with new about 100 organisations active in global health in the routed to address major communicable diseases. They take different forms such as a finance mechanism implementation programme or product develop mem products such as drugs for neglected tropical diseases. These organisations have contributed significant threat to health to the increase of funding in global health in the last proundation, Meckefeller Foundation, Mellocome Trust, Aga Roundation, Mellocome Trust,		Among these organisations, some are active at the global level while others work at the regional level (eg, EU which	AMR. For example, the EU adopted plans on AMR. ⁸² Within the European region, ECDC aggregates data from EU countries on AMR. JPIAMR supports research on AMR. OECD has also published reports on AMR. The International Committee of the Red Cross has been involved in the fight against AMR mainly in conflict areas. TATFAR was created in 2009 to enhance expert	
Bill and Melinda Gates Foundation, Rockefeller Foundation, Nellcome Trust, Aga Khan Foundationto the increase of funding in global health in the last 20 years. Founded in 2000, the Bill and Melinda Gates Foundation has become a prominent player with total payments of more than US\$58.8 billion since inception. With their huge financial means, the largest health agenda.collaboration between different actors to develop action and funding programmes against AMR. Through its forums, the World Economic Forum, which included AMR as part of its global risks report, plays a leading role in calling for cooperation between the public and private sectors to develop innovative solutions to combat AMR as part of its global risks report, plays a leading role in calling for cooperation between the public and private sectors to develop innovative solutions to combat AMR as part of a wider organisations for defective scores borders. International NGOs have been or access to effective drugs for access to effective drugs for all.Global civil society organisations Frontières); OXFAM International Committee of the Red CrossGlobal civil society organisations are non-governmentation non-business organisations or movements that are active across borders. International NGOs have been critical in promoting access to medicines as illustrated by the access to medicine campaign which drugs in LMICs). Global civil society and plays a critical role in advocacy and scrutiny of health policy.Some NGOs have a mandet specifically on AMR while many other NGOs address AMR as part of a wider portfolio of activities on health or humanitarian issues. Nut a global reach such as Doctors without Borders have been critical in promoting access to medicines as illustrated by the access to effective drugs for all.Global civil society orga	Drugs for Neglected Diseases; Global Fund to Fight AIDS, Tuberculosis, and Malaria; GAVI;	been the development of new mechanisms to deliver targeted funding and health technologies in LMICs. ⁸³ Global health initiatives are hybrid organisations which mix elements of the private and public sector. They are typically programmes targeted at specific diseases that were created to address major communicable diseases in LMICs. Their number has risen sharply since 2000 with now about 100 organisations active in global health. They take different forms such as a finance mechanism, implementation programme or product development partnership which aims to research and develop new products such as drugs for neglected tropical diseases. These organisations have been endowed with large means and have developed strong expertise in the delivery of health technologies, medications and	regarding innovation, access to medicines and infection prevention. Organisations such as GAVI have been critical to step immunisation. Regarding access, the main solution has been for HICs to subsidise access to antimicrobials in LMICs through the establishment of funding and procurement organisations (eg, Global Fund or the Medicine Patent Pool, UNITAID). On innovation, several organisations are conducting R&D activities. Recently, efforts have resulted in the creation of GARDP	
IOs) G7 and G20that were primarily created to address financial and macroeconomic issues but have come to cover many pressing issues in global governance.organisations have played a key role in putting AMR on the global political agenda.Global civil society organisations Care International; Doctors without Borders (Médecins Sans Frontières); OXFAM International; Save the Children; People Health Movement; International Committee of the Red CrossGlobal civil society organisations or movements that are active across borders. International NGOs have been an important player in global health in implementation programmes as well as in advocacy (eg, the campaign for access to essential medicines by Doctors without Borders, which aims at increasing the availability of drugs in LMICs). Global civil society also encompasses networks such as the People Health Movement, which focuses on health equity and plays a critical role in advocacy and scrutiny of health policy.Some NGOs have a mandate specifically on AMR while many other NGOs address AMR as part of a wider portfolio of activities on health or humanitarian issues. With a global reach such as Doctors without Borders, which aims at increasing the availability of drugs in LMICs). Global civil society also encompasses networks such as the People Health Movement, which focuses on health equity and plays a critical role in advocacy and scrutiny of health policy.To address market failure in the research and development of new drugs, some governments have step up incentives. ²⁴ Most research on R&D on new antimicrobials is led by small firms. In addition, big pharmaceutical companies are contributing to global	Bill and Melinda Gates Foundation, Rockefeller Foundation, Wellcome Trust, Aga	to the increase of funding in global health in the last 20 years. Founded in 2000, the Bill and Melinda Gates Foundation has become a prominent player with total payments of more than US\$58.8 billion since inception. With their huge financial means, the largest philanthropies can set priorities and influence the global	collaboration between different actors to develop actions and funding programmes against AMR. Through its forums, the World Economic Forum, which included AMR as part of its global risks report, plays a leading role in	
organisations Care International; Doctors without Borders (Médecins Sans Frontières); OXFAM International; Save the Children; People Health Movement; International Committee of the Red Crossnon-business organisations or movements that are active across borders. International NGOs have been an important player in global health in implementation programmes as well as in advocacy (eg, the campaign for access to essential medicines by Doctors without Borders, which aims at increasing the availability of drugs in LMICs). Global civil society also encompasses networks such as the People Health Movement, which focuses on health equity and plays a critical role in advocacy and scrutiny of health policy.many other NGOs address AMR as part of a wider portfolio of activities on health or humanitarian issues. With a global reach such as Doctors without Borders have been critical in promoting access to medicines as illustrated by the access to medicine campaign which advocates for access to effective drugs for all.Private industry Pharmaceutical companies; biotechnology companies information and communication companiesThe private industry researches, develops and disseminates products such as drugs, diagnostics and technologies through global markets. Some private companies have set up their own foundations and/or participated in public-private partnerships.To address market failure in the research and development of new drugs, some governments have step up incentives. ²² Most research on R&D on new antimicrobials is led by small firms. In addition, big pharmaceutical companies are contributing to global	IOs)	that were primarily created to address financial and macroeconomic issues but have come to cover many	organisations have played a key role in putting AMR on	
companies; biotechnology companies; information and communication companies communication companies communication companies companies have set up their own foundations and/or participated in public–private partnerships.	organisations Care International; Doctors without Borders (Médecins Sans Frontières); OXFAM International; Save the Children; People Health Movement; International	non-business organisations or movements that are active across borders. International NGOs have been an important player in global health in implementation programmes as well as in advocacy (eg, the campaign for access to essential medicines by Doctors without Borders, which aims at increasing the availability of drugs in LMICs). Global civil society also encompasses networks such as the People Health Movement, which focuses on health equity and plays a critical role in	many other NGOs address AMR as part of a wider portfolio of activities on health or humanitarian issues. With a global reach such as Doctors without Borders have been critical in promoting access to medicines as illustrated by the access to medicine campaign which	
	companies; biotechnology companies; information and	disseminates products such as drugs, diagnostics and technologies through global markets. Some private companies have set up their own foundations and/or	development of new drugs, some governments have step up incentives. ²² Most research on R&D on new antimicrobials is led by small firms. In addition, big	

Continued

Types of actor and examples	Role in the global health system	Role in the global governance of AMR
Academic institutions and professional associations ReAct	Academic institutions and networks play an essential role in global health education and research. Professional societies gather members of the same profession such as physicians or nurses and usually set professional standards. In global health, professional societies can enact recommendations for best practices and play a role in global health education.	Academic institutions conduct both fundamental and applied research on AMR including drug discovery, clinical research and wider public health and governance aspects of AMR. Additionally, academic experts contribute to technical meetings from IOs.

AMR, antimicrobial resistance; ECDC, European Centre for Disease Prevention and Control; EU, European Union; GAP, Global Action Plan; GARDP, Global Antibiotic Research and Development Partnership; GAVI, The Vaccine Alliance; HICs, high-income countries; ICRC, International Committee of the Red Cross; IOs, international organisations; ISO, International Organization for Standardization; JPIAMR, Joint Programming Initiative on Antimicrobial Resistance; LMICs, low/ middle-income countries; MPP, Medicine Patent Pool; NGOs, non-governmental organisations; OECD, Organization for Economic Co-operation and Development; R&D, research and development; TATFAR, Transatlantic Taskforce on Antimicrobial Resistance; UN, United Nations.

fall mainly on a few IOs, whereas several other IOs are involved or simply relate to the governance of AMR. Given their strong mandate on AMR, one can expect that the current core organisations will remain so in the future. However, our assessment suggests several opportunities for further involvement in terms of specific AMR policy goals, functions of IOs, diseases, populations, sectors and regions (these activities are summarised in table 3). We believe that the thematic and functional clusters can further guide institutional involvement in the global governance of AMR. First, organisations within the same cluster do not necessarily all need to be involved or work together. For example, although several UN organisations are specialised in training activities (eg, United Nations Institute for Training and Research, United Nations University, etc), a partnership between the Tripartite and one of those organisations might be sufficient to reach the goal set. Second, organisations can be part of several clusters. This is especially the case of organisations that have a broad mandate on AMR such as WHO. Third, a few organisations such as UNEP might become a core organisation to tackle AMR (Tripartite+). However, the role of most organisations would be limited to some specific aspects of the challenge, functions or populations regarding AMR. Even when limited, involvement can have multiplying effects within a strongly interlinked system.

Finally, some limitations can be identified. As data collection was web-based, some activities might have been undertaken without being reported on the internet. In addition, search was more likely to find recent activities than older ones. This is particularly the case before the 1990–2000s, where relevant literature is often accessible through the grey literature. Interviews with key informants could be useful to complement the findings of the web and document searches. Second, while we discussed the overall 'output effectiveness', the reported activities are not assessed in terms of 'outcome/impact effectiveness' and the capabilities or organisation in terms of know-how, resources and knowledge. Although our search of documents allowed us to provide relevant background about the broader context of engagement of IOs to address AMR, we were not able to evaluate the level

of engagement of IOs for each activity. For example, it may be that an organisation published once on AMR but did not sustain any activity through time. Third, the distinction between relevant and distant depends to some extent on what is considered as relevant. Here we considered activities that focus on several challenges of global collective action but there might be other ways to organise results that the ones identified by our interdisciplinary framework. Given the challenge of collecting all relevant documents on AMR-sensitive activities, our assessment of AMR-sensitive activities might be incomplete. Fourth, our assessment focuses on the UN while the global governance of AMR extends to other organisations. In addition, while several entities within a specific IO can be involved in tackling AMR, the analysis did not have such granularity. Internal dynamics of competition may have substantial implications, for example, when specific units within a specific IO seek to secure funding for their own purposes. Another limitation is the focus of the analysis at the global level whereas several IOs work at the regional level (eg, European Union) or at both the international and regional levels (eg, WHO regional offices) where priorities between global and regional activities can differ.⁴⁸ Finally, our study focuses on the international level but national and subnational actors are ultimately the ones that need to be involved and convinced about the need and urgency of implementing AMR policies through a coordinated multisectoral and multilevel approach where authority is increasingly dispersed.649

CONCLUSION

In conclusion, our assessment demonstrates the complexities of the global governance of AMR within and beyond the UN system. While WHO has been the traditional actor tackling AMR, there is a growing realisation that it is unable to address the problem on its own given the strong intersectoral nature of AMR. This takes place in a broader context of global governance marked by the evolution from the responsibility of one organisation to a network of organisations that seek to achieve common goals.

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Cluster of relevant	Addressing gaps in global	
organisations	governance	Potential for further involvement of global stakeholders
Infection prevention DESA, FAO, GAVI, ILO, ISO, OIE, UNISDR, UN- Water, UNICEF, WHO, WTO	AMR is another strong argument for the implementation of water, sanitation and hygiene (WASH) interventions in LMICs (SDG6) and for better immunisation (SDG3). Synergies between AMR and interventions to tackle infectious diseases and pandemic preparedness should be better assessed. There is currently limited coordination to design intervention based on co-benefits.	 The ILO could raise awareness of the standards in occupational health and hygiene and develop the capacity of national actors to use them. GAVI activities on improving immunisation coverage in LMICs can be better linked to the goal of reducing AMR. In food-producing animals, standards of hygiene could be set up on the ISO model as there are already standards for food safety. UNISDR may incorporate AMR concerns into disaster reduction strategies. DESA could strengthen its AMR objectives by encouraging Member States to use the SDGs review processes to highlight the links between AMR and SDG under review. UN-Water could take action to reduce water contamination by advocating for improvements in the sewage system.
Surveillance UNEP, WHO, OIE, FAO, IMF	The Tripartite has strengthened surveillance of AMR but there are gaps in capacities between countries. In addition, there is limited effort for surveillance of antimicrobial use in animal health and a lack of environmental surveillance of AMR. Another need is a better integration/harmonisation of surveillance standards across sectors.	 The WHO-OIE-FAO Tripartite could improve integration of AMR surveillance with the IHR and PVS pathways to foster an integrated approach to zoonosis. UNEP should become more involved in environmental surveillance of antibiotic residues and AMR. IMF could integrate vulnerabilities of countries to infectious diseases including AMR in its evaluation about countries' economic growth and stability.⁸⁵ What has been created for AGISAR could be scaled up by combining its strength with the work of GLASS for the collection of epidemiological data.
Conservation WHO, IFAD, UN- Habitat, ISO, FAO, OIE	Current global mechanisms for the conservation of antimicrobials rely mainly on the diffusion of information and adoption of standards. Promotion of antimicrobial use is poorly controlled. The responsibility for conservation is shared between different entities. More efforts are needed to translate this into actionable mechanisms, which are currently more developed in human health than in other sectors. An important challenge is to define common standards for the different areas (ban of antibiotics as growth promoters, critically important antibiotics).	 The Tripartite could work together to ban the use of antibiotics as growth promoters. A code of conduct, like the international code on breastmilk substitute, could be used to restrict promotion and advertising of antimicrobials by pharmaceutical companies. The Tripartite could set up a mechanism like the consultative Expert Working Group on research and development to assess the potential effectiveness and feasibility of conservation strategies. IFAD can build global capacity in the agricultural sector to address AMR. UN-Habitat could provide a relevant entry point for the important topic of AMR dissemination through wastewater and the treatment plan. ISO could be involved in setting standards regarding antimicrobial use and further embed actions against AMR in the global economy.
Containment ICAO, IMO, UNEP, UNWTO, WHO, OIE, FAO, Wellcome Trust, BMGF	Expansion of containment strategies should be based on the identification of new ways to prevent the dissemination of AMR in different settings, including in the environment.	 Organisations related to transport (IMO or ICAO) or tourism (UNWTO) could help address international containment of AMR through their respective mandate. Funding agencies might support more research on the ecology and transmission of AMR including the flow of antimicrobial residues and genetic elements.
Innovation GARDP, ITU, UNCTAD, Wellcome Trust, WIPO, WHO	Addressing market failures against AMR requires market incentives and innovative funding mechanisms. Current efforts should build on existing structures that have proven to be effective.	 ITU could help facilitate the discussion regarding innovation. UNCTAD could play a supporting role given its wide range of activities in the development of new therapies and diagnostics, and its efforts to improve access to these therapies and diagnostics.

 Table 3
 Opportunities for further development of the global governance of AMR based on clusters of competencies and functions

Continued

Table 3 Continued

Addressing gaps in global governance	Potential for further involvement of global stakeholders
Access to antimicrobials has been mainly supported in specific diseases. A current challenge is that access to medicines can conflict with other challenges of global collective action such as conservation. However, enhancing access can also reduce the use of substandard medicines. The report of the UN Secretary-General's high- level panel on access to medicines stated that an important issue for improved coherence is the diversity of accountability mechanisms between access and innovation. ²⁷	 Extending the mandate of relevant organisations, with and without a primary health mandate, such as the Global Fund, UNAIDS and UNDP, to increase access and delivery to essential drugs in LMICs. HRC, OHCHR and other human rights bodies can raise awareness and build common international norms about access to medicines. IOs working to combat fake medicines (eg, UNICRI, UNODC) can contribute to limit the use of substandard antimicrobials (in partnership with the WHO, which created the Global Surveillance System for substandard and fake medical products in 2003). UNCTAD could play a supporting role given its wide range of activities in the development of new therapies and diagnostics, and its efforts to improve access to these therapies and diagnostics. UNOPS, in partnership with other organisations, could provide demand-driven operational support for the procurement and distribution of quality medicines.
The Tripartite produces communication material (eg, global campaign) but other IOs with specific commitments and targets are needed to raise awareness and build evidence on AMR, particularly in areas where AMR is not commonly discussed. An important contribution would be to better measure the (cost-) effectiveness of interventions about AMR and understand how to tailor intervention to the context.	 IOs involved in knowledge creation and transmission (UNESCO, UNITAR, UNU, UNSSC) could leverage their expertise in training and education to raise awareness on AMR. IOs with specific constituencies that are important to tackle AMR (eg, chronic diseases (UNIATF), women (UN-Women), tourism (UNWTO) could also raise awareness about AMR for different constituencies. Several business-oriented organisations could play a role in advocacy for AMR. These organisations include UNCTAD, ITC or UNIDO but also the WEF which has already put AMR on its global health risks report. Several private foundations such as the Wellcome Trust, the BMGF and the Flemming fund have critical funding capacities for research and education on AMR.
AMR has been a growing problem in several specific diseases including TB, malaria and HIV-AIDS. AMR is another strong argument to build the case for tackling these diseases through an intersectoral approach. AMR is an important case for maximising positive synergies in strengthening health system in LMICs. ⁸⁶	 support of GFATM and other organisations on specific diseases to build broader AMR capacity. Given its large funding power, the Global Fund could become a key player to address and coordinate AMR challenges regarding TB, HIV and malaria in LMICs among the different actors.
Fragile and discriminated people may be more at risk of AMR. ⁸⁹ There is still a lack of empirical evidence on the link between inequalities and AMR but there is a growing sense that AMR can be an obstacle to poverty reduction. ^{8 90} Several IOs may contribute to raise awareness and, in some cases, mitigate the effect of AMR in these populations.	 (Humanitarian) agencies working in fragile context such as OCHA, WFP, UNHCR, UNICEF, MSF and ICRC can improve access, infection prevention, surveillance, containment and conservation. Some IOs can also raise awareness about AMR in specific populations, particularly women and poor communities. This can be done in collaboration with the Tripartite and UNDP, which already play a leading role in assessing the impact of AMR on the SDGs. Some mechanisms such as the UNIATF as well as treaty-based organisations (ie, Convention on the Rights of Persons with Disabilities) can also raise awareness about AMR in specific populations.
	governanceAccess to antimicrobials has been mainly supported in specific diseases. A current challenge is that access to medicines can conflict with other challenges of global collective action such as conservation. However, enhancing access can also reduce the use of substandard medicines. The report of the UN Secretary-General's high- level panel on access to medicines stated that an important issue for improved coherence is the diversity of accountability mechanisms between access and innovation.The Tripartite produces communication material (eg, global campaign) but other IOs with specific commitments and targets are needed to raise awareness and build evidence on AMR, particularly in areas where AMR is not commonly discussed. An important contribution would be to better measure the (cost-) effectiveness of interventions about AMR and understand how to tailor intervention to the context.AMR has been a growing problem in several specific diseases including TB, malaria and HIV-AIDS. AMR is another strong argument to build the case for tackling these diseases through an intersectoral approach. AMR is an important case for maximising positive synergies in strengthening health system in LMICS. ⁶⁶ Fragile and discriminated people may be more at risk of AMR. ⁸⁹ There is still a lack of empirical evidence on the link between inequalities and AMR but there is a growing sense that AMR can be an obstacle to poverty reduction. ^{8,90} Several IOS may contribute to raise awareness and, in some cases, mitigate the

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Table 3 Continued

Cluster of relevant organisations	Addressing gaps in global governance	Potential for further involvement of global stakeholders
Sustainable funding Asian Infrastructure Investment Bank, IMF, World Bank	Sustainable funding is a key issue for many AMR efforts. A first challenge is funding dedicated to strengthening global governance. Some countries have helped through earmarked funding for AMR. The other is development aid to strengthen capacities in LMICs. While savings regarding One Health approach have been estimated for important components in tackling infectious diseases, more precise assessment and pilot studies are needed regarding AMR. ⁹¹	 IMF could include assessments of countries' systemic risk caused by infectious diseases in financial evaluation to strengthen nations' government incentives to act on AMR as access to financial market is capital for them.⁹² The World Bank can further identify and prioritise interventions that are relevant to LMICs. The World Bank, IMF and/or the Asian Infrastructure Investment Bank along with other financial players can better link the financial sector to health goals.⁹³ Financial organisations could provide expertise in quantifying AMR risks and efforts, which will be important in securing funding for such projects amidst fierce competition for attention and resources. Engaging insurance (or, more importantly, reinsurance) companies that could play an important role in this group. These evaluations should not only focus on costs but also on potential savings.
Coordination CEB, ECOSOC, JIU, UNGA, UNSG, HLPF	AMR requires coordination and collaboration across many different areas. In addition to mechanisms specific to AMR, several IOs and mechanisms share a coordination mandate which could provide relevant fora for discussion on AMR. This should facilitate the definition of a common agenda on infectious disease and issue linkages with other pressing global challenges.	 Due to near universal membership, the UNGA is an appropriate forum to encourage continuous high-level support on AMR. ECOSOC might be an important forum for discussion related to the socioeconomic impact of AMR. EOSG could advocate for AMR and encourage alignment across the One Health agenda. Other high-level relevant fora for attention grabbing include the G7 and G20 which have already included AMR in their agenda. With its broad mandate on sustainable development, the HLPF provides a platform for follow-up and review of AMR-sensitive activities.

AGISAR, Advisory Group on Integrated Surveillance of Antimicrobial Resistance; AMR, antimicrobial resistance; BMGF, Bill and Melinda Gates Foundation; CEB, Chief Executive Board; CRPD, Convention on the Rights of Persons with Disabilities; DESA, Department of Economic and Social Affairs; ECOSOC, Economic and Social Council; EOSG, Executive Office of the Secretary-General; FAO, Food and Agriculture Organization; GARDP, Global Antibiotic Research and Development Partnership; GAVI, The Vaccine Alliance; GFATM, Global Fund to Fight AIDS, Tuberculosis and Malaria; GLASS, Global Antimicrobial Surveillance System; HLPF, High-Level Political Forum on sustainable development; HRC, Human Rights Council; ICAO, International Civil Aviation Organization; ICRC, International Committee of the Red Cross; IFAD, International Fund for Agricultural Development; IHR, International Health Regulations; ILO, International Labour Organization; IMF, International Monetary Fund; IMO, International Maritime Organization; IOM, International Organization for Migration; IOs, international organisations; ISO, International Organization for Standardization; ITC, International Trade Center; ITU, International Telecommunication Union; JIU, Joint Inspection Unit; LMICs, low/middle-income countries; MSF, Médecins Sans Frontières; OCHA, Office for the Coordination of Humanitarian Affairs; OHCHR, Office of the United Nations High Commissioner for Human Rights; OIE, World Organization for Animal Health; OSAA, Office of the Special Adviser on Africa; PVS, Performance of Veterinary Services; SDGs, Sustainable Development Goals; TB, tuberculosis; UN, United Nations; UNAIDS, United Nations Programme on HIV/AIDS; UNCTAD, United Nations Conferences on Trade and Development; UNDP, United Nations Development Programme; UNEP, United Nations Environment Programme; UNFPA, United Nations Population Fund; UNGA, United Nations General Assembly; UNHCR, United Nations High Commissioner for Refugees; UNIATF, United Nations Interagency Task Force on Non-Communicable Diseases; UNICRI, United Nations Interregional Crime and Justice Research Institute; UNIDO, United Nations Industrial Development Organization; UNISDR, United Nations Office for Disaster Risk Reduction; UNITAR, United Nations Institute for Training and Research; UNODC, United Nations Office on Drugs and Crime; UNOPS, United Nations Office for Project Services; UNRISD, United Nations Research Institute for Social Development; UNRWA, United Nations Relief and Works Agency; UNSG, United Nations Secretary-General; UNSSC, United Nations System Staff College; UNU, United Nations University; UNWTO, World Tourism Organization; WEF, World Economic Forum; WFP, World Food Programme; WIPO, World Intellectual Property Organization; WTO, World Trade Organization.

Our assessment demonstrates that the 'whole of UN approach' adopted in the wake of the 2015 WHO GAP AMR has supported increased collaboration in certain areas but that the outcome/impact effectiveness of such approach deserves further attention. In addition, further research is needed to better account for the varying degrees of engagement across time. Within this deliberate attempt from the World Health Assembly and the UNGA to expand participation in addressing AMR, the key governing principle should be to ensure that the governance system remains fitted to the evolving challenge of AMR. A key strategy for maximising institutional fit is to reflect on the nature of the six challenges of collective action.

6

While some organisations might be more relevant, more willing, and above all, more likely to receive support from their Member States than others to address AMR, our approach by cluster of organisations offers a range of options to consider in seeking to expand and improve AMR governance in relevant areas.

As the global governance of AMR is embedded within wider processes of global (health) governance, these findings emphasise the relevance of leveraging existing resources, mechanisms and institutions rather than reshuffling the governance of AMR. A 'whole of UN approach' in an essentially polycentric governance system currently provides a realistic yet underevaluated approach to address AMR. This 'whole of UN approach' should be part of wider efforts towards building more sustainable and resilient societies.

The Tripartite+ constituted of the core organisations (WHO, OIE and FAO + UNEP) should continue to lead the way for developing the AMR agenda. Harnessing the power of the UN system to move the AMR agenda forward can not only help tackle AMR, but it can also contribute to transform the UN system to make it ripe to address the challenges of the 21st century.

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