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ANALYSIS

ACCELERATING ACTION ON SUSTAINABLE DEVELOPMENT GOALS

# Trade is central to achieving the sustainable development goals: a case study of antimicrobial resistance

Johanna Hanefeld and colleagues highlight the links between trade and health and argue for greater consideration of trade agreements in actions to meet the sustainable development goals

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The 17 sustainable development goals adopted by 198 nations in September 2015 provide a transformative framework for global action across a wide range of areas. Health is an essential component of many of the goals, such as those relating to hunger, the environment, and education. The third goal (SDG3) focuses explicitly on health: "Ensure healthy lives and promote well-being for all at all ages."<sup>1</sup>

Like health, trade is explicitly targeted in several of the goals. However, it is also a critical determinant of other goals where it has not been mentioned. We discuss the often overlooked centrality of trade as a determinant of health through a case study of antimicrobial resistance.<sup>2</sup> Although not explicitly mentioned in the sustainable development goals, antimicrobial resistance threatens health, prosperity, and sustainable development.<sup>3</sup> It is responsible for an estimated 700 000 deaths annually, forecast to be 10 million by 2050.<sup>3</sup>

#### Trade and health

The link between trade (agreements) and health (systems) has been well documented over the past decade.<sup>4-8</sup> The interaction can be considered in four parts:

*Commodities traded*—This may be commodities harmful to health, such as tobacco and alcohol,<sup>9 10</sup> or those which may be beneficial, such as fresh fruit and vegetables.<sup>11-13</sup>

*Trade agreements and the provisions*—The legal framework provided for which goods can cross borders. Here the initial focus was on access to medicines and intellectual property (and this connection is explicitly acknowledged in SDG3 (target 3B), mainly through the Agreement on Trade Related Aspects of Intellectual Property Rights (TRIPS).<sup>14-16</sup> Over the past five years the focus has been increasingly on regional and bilateral trade agreements, often related to non-communicable diseases and their risk factors.<sup>17-20</sup>

*Trade in services*<sup>21</sup>—This includes the movement of health workers, telemedicine, medical tourism, and foreign direct investment in health systems.<sup>22-24</sup>

*Wider determinants of trade on health*—These are both positive and negative,<sup>25</sup> including patterns of employment, urbanisation, income, and housing, with considerable evidence to suggest trade as determinant of health inequities.<sup>8-27</sup>

## Trade and antimicrobial resistance

Antimicrobial resistance has rapidly ascended the political agenda and is now recognised as a major threat to health, prosperity, and global health security.<sup>28</sup> Substantial funding—with estimates of \$40bn (£31bn; €35bn) over the next 10 years—is being sought to tackle the problem, and the World Health Assembly endorsed a global action plan on antimicrobial resistance in 2015.<sup>28</sup> But meeting these challenges will not be straightforward as it requires the collaboration of a large number of stakeholders across animal, human, and environmental spheres, often with conflicting interests and networks.<sup>29</sup>

Here we outline core areas where trade is critical to tackling antimicrobial resistance, highlighting key areas for action to achieve the SDGs and the 2030 agenda (box 1). We will focus

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mainly on commodities traded and the possible effect of trade related regulation and agreements.

Trade and antimicrobial resistance intersect in several ways. On a fundamental level trade (mobility) of people, animals, and goods is connected to the spread of microbes, and in human transmission there is a clear association with travel routes and the emergence of antimicrobial resistance.<sup>5 30</sup>

## Commodities

# Trade in food products that contain resistant bacteria

The most important sector in terms of trade relating to antimicrobial resistance is in livestock, food animals, and their feed, particularly in animals that host resistant bacteria. For example, the global trade in meat is large: the United States Department of Agriculture predicted that global production of beef and veal would rise to 62 million tonnes in 2017, with global exports predicted to be 9.6 million tonnes. Export of broiler chicken meat was expected to be a record 11.2 million tonnes.<sup>37</sup>

Drug resistant *Escherichia coli* can live on beef carcasses even after 24 hours in a chiller and in minced beef that has been stored for up to eight days.<sup>38</sup> A study of raw meat samples provided by retail traders in the Netherlands found meticillin resistant *Staphylococcus aureus* (MRSA) in many of the samples, ranging from 35.3% of turkey samples to 2.2% of game samples.<sup>39</sup> While many low and middle income countries export food products and animals, their capacity to monitor antimicrobial resistance adequately may be limited by overall budget constraints.<sup>35 40</sup>

Trade is a possible route for spreading resistance. One response, where animals are tested for resistant strains as part of efforts to control the spread of antimicrobial resistance, is likely to affect trade immediately as animals with positive results would be banned. A challenge here is that although many farmers are advised to vaccinate their animals, it is hard to enforce and to differentiate between livestock vaccinated and those infected. Vaccination is also not currently routine in all countries. Thus, national strategies to contain antimicrobial resistance through vaccination may clash with global antimicrobial resistance control and prevent farmers from trading their animals across national borders.

#### Availability of antimicrobials

Trade in antimicrobials also affects their availability for human and animal consumption. Part of the complexity of addressing antimicrobial resistance is balancing access to medicines against overconsumption; at the global level the consensus is that we are probably consuming too many antimicrobials, but many of the worlds' poorest people still lack access to the essential medicines they need. More than a million children die each year from sepsis and untreated infection,<sup>31</sup> which is inconsistent with target 8 of SDG3: "Achieving universal health coverage including access to safe, effective, quality, and affordable essential medicines for all."

Clearly, efforts to tackle antimicrobial resistance have to acknowledge that appropriate access is part of a wider global trading system, which links incentives and profits from medicines to volume of sales. Access to good quality antimicrobials is determined by intellectual property provisions set out as part of trade agreements, which set availability and price within a country, as well as regulations concerning safety and substandard medicines, relying on local action and enforcement. These intellectual property provisions are either insufficient or insufficiently implemented to ensure access to all those who require it. Here trade and the existing trade agreements, including TRIPS, have a central role. Indeed, this is reflected in one of the targets linked to SDG3, which calls for better use of flexibilities granted under TRIPS to increase access to medicines:

"Provide access to affordable essential medicines and vaccines, in accordance with the Doha Declaration on the TRIPS Agreement and Public Health, which affirms the right of developing countries to use to the full the provisions in the Agreement on Trade Related Aspects of Intellectual Property Rights regarding flexibilities to protect public health, and, in particular, provide access to medicines for all."

Addressing antimicrobial resistance and maintaining drug effectiveness while increasing access will be a key challenge and require nuanced, equity sensitive policy implementation. WHO's recent reclassification of antimicrobials into "access", "watch", and "reserve" marks a step in this direction.<sup>32</sup>

Trade also affects the availability of substandard "counterfeit" antimicrobial drugs. Trade in counterfeit drugs is estimated to run into billions.<sup>34</sup> Availability of substandard antimicrobials is likely to undermine attempts at public regulation of antimicrobials, such as oversubscribing or the provision of antimicrobials through informal providers. Little is known about the effect of counterfeit trade in animal drugs.<sup>35</sup> However, trade agreements and greater regulation through bodies such as the World Trade Organisation (WTO) offer solutions to this challenge.

#### **Trade agreements**

The importance of trade and intellectual property provisions to health and drug innovation is explicitly set out in target 3b of SDG3. It is acknowledged as a core aspect of the Global Strategy to Fight Antimicrobial Resistance and affirmed by the political declaration resulting from the UN General Assembly special session on antimicrobial resistance in 2016. The political declaration calls on governments to: "recognise the importance of delinking the cost of investment in research and development on antimicrobial resistance from the price and volume of sales so as to facilitate equitable and affordable access to new medicines, diagnostic tools, vaccines."

This is important as some of the largest unmet needs in the research and development of new antimicrobials are found in diseases such as tuberculosis, which contribute substantially to the global antimicrobial resistance burden.<sup>33</sup> New drugs that would shorten the six month TB treatment course would alleviate the huge burden on patients and health systems and help to reduce the emergence of roughly 480 000 new multidrug resistant cases each year; these are at least partly related to poor adherence to the long treatment course.<sup>33</sup>

However, research into TB treatment is not profitable enough to warrant sufficient investment from drug companies. This is reflected in the huge funding gap for TB; less than one third of the \$3.7bn for new drugs called for in the Stop TB Partnership's Global Plan to Stop TB 2011-15 was raised. Concerns extend to diagnostic tools, vaccines for disease prevention, and wider antimicrobial resistance related innovation. For example, healthcare facilities may require new types of furniture and tools with antibacterial surfaces. Such products may be an area of future innovation but need to be affordable by low and middle income countries.

#### Box 1: Trade and health framework for antimicrobial resistance

Commodities traded

- · Availability of antimicrobials
- · Availability of food, including livestock, that may contain resistant bacteria

Trade agreements

- · Regulation of counterfeit medicines
- · Availability of new antimicrobials
- · Interpretive labelling of food

Trade in services, including movement of people

Spread of infection through human mobility

Effect on wider determinants of health

- · Pattern of urbanisation
- · Livestock farming practices

Food, animal, and plant safety in trade is regulated globally through the WTO's Sanitary and Phytosanitary Agreement. It sets out basic standards of food safety based on science, while providing some room for countries to set and retain their own regulation. A recent analysis of interpretive nutrition labelling and trade discussions under WTO regulation found that these were often at odds and there was a risk that trade agreements may constrain public health regulation.<sup>36</sup>

#### Trade in services

Other areas of trade and health also affect antimicrobial resistance, even though these have had less attention. Trade in services, including medical tourism, has been documented as contributing to the spread of resistant bacteria.<sup>41</sup>

#### Wider determinants of health

Increases in global trade and the resulting changing patterns of employment have resulted in intensive urbanisation and changes in farming.<sup>5</sup> These in turn affect resistance in animals and its transmission. This also links closely to SDG11, which is focused on urbanisation.

#### Discussion

Our case study highlights the need to consider trade when seeking to tackle antimicrobial resistance and achieving the sustainable development goals. Given the close connection between food products, including food animals, and the spread of antimicrobial resistance, this seems likely to become the subject of future trade discussions and dispute—for example, if individual countries wish to introduce public health regulation that requires labelling of antibiotic use for food, or restrict imports from specific countries.

Similar public health regulations around interpretive nutrition labelling for food have led to trade discussions in the past, such as regulation surrounding fatty cuts of meat in Ghana,<sup>42</sup> or the EU-US dispute on the use of growth hormone in cattle.<sup>43</sup> It has yet to be established whether antimicrobial resistance could be used as a rationale for a "public health emergency" which allows for greater flexibility around intellectual property regulation under TRIPS.<sup>44,43</sup>

#### Regulation

It is therefore essential that trade agreements and regulating bodies, such as the WTO and the World Intellectual Property Organisation (WIPO), consider antimicrobial resistance in the discussions on, and adjudication of disputes on, intellectual property protection. Standard setting bodies such as the Codex Alimentarius Commission and the OIE (World Organisation for Animal Health) also have an important role since WTO members are obliged to base regulations on relevant international standards. By setting stronger regulations these bodies can help tackle antimicrobial resistance and contribute to achieving the sustainable development goals.

Free trade agreements are especially important because they typically liberalise trade in agriculture to a greater extent than WTO commitments and can include more specific obligations. This relates closely to SDG16 (transparent institutions and participatory policy processes).

## Role of civil society, think tanks, and academic institutions

It is important to develop international regulation and work with national regulators to prevent pitching trade against public health advocates in relation to antimicrobial resistance.<sup>45 46</sup> Civil society has been active in highlighting the intersection between trade and health, ranging from the HIV access movement to the recent opposition to the TTIP agreement between the EU and the US.<sup>47</sup>

Involvement of civil society, think tanks, and academic institutions is crucial for effective action on antimicrobial resistance. In Sweden, which has responded to antimicrobial resistance faster and earlier than other countries, public mobilisation and non-governmental organisations such as ReAct have been critical to placing the issue high on public and policy agenda.<sup>48</sup> Equally, recent protest by doctors and drug sellers in Punjab, Pakistan, in response to stricter laws on substandard drugs highlight the importance of involving civil society from the outset and the need for greater public debate to improve understanding of the challenges of antimicrobial resistance.

Advances are also needed in governance mechanisms to better facilitate collaboration between health and trade, both in terms of where health convenes, as well as where health needs to engage for a better governance for health.<sup>25</sup>

These challenges throw down the gauntlet to think tanks and academic institutions to go beyond knowledge generation and dissemination. There is need to develop new ways of working on health and trade to tackle the big issues in sustainable development—such as the threat of antimicrobial resistance. The myriad ways in which trade will affect sustainable development also apply to sectors other than health. The lessons for all sectors include the need to identify interlinkages between sectors and goals, the importance of governance, and the need for strong and just institutions as well as inclusive processes.

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#### ANALYSIS

#### Key messages

Antimicrobial resistance shows how trade can affect health and the sustainable development goals

Trade and its intersection with health is critical to achieving sustainable development, yet it is little understood

Civil society, including think tanks, and academic institutions have a key role in exploring and acting on the intersection between trade and health