

Follow-up of the report of the Consultative Expert Working Group on Research and Development: Financing and Coordination

Report by the Director-General

1. In resolution WHA69.23 (2016), the Health Assembly requested the Director-General to expedite the full implementation of the strategic workplan endorsed in resolution WHA66.22 (2013). Resolution WHA69.23 includes the following elements:

- development of WHO's Global Observatory on Health Research and Development;¹
- full implementation of the strategic workplan;
- establishment of an Expert Committee on Health Research and Development to provide technical advice on the prioritization of health research and development;
- exploration of the feasibility of a voluntary pooled fund to support research and development for Type III and Type II diseases and specific research and development needs of developing countries in relation to Type I diseases.

2. More specifically, the Health Assembly in resolution WHA69.23 requested the Director-General to submit to the Seventieth World Health Assembly, through the Executive Board at its 140th session, two texts: terms of reference and a costed workplan for the Global Observatory on Health Research and Development; and a proposal with goals and an operational plan for a voluntary pooled fund to support research and development for Type III and Type II diseases and specific research and development needs of developing countries in relation to Type I diseases.

3. The terms of reference and a costed workplan of the Global Observatory on Health Research and Development are submitted in Annex 1.

4. The proposal for a voluntary pooled fund is contained in Annex 2. Further details on the operationalization of the fund are currently being developed by the UNICEF/UNDP/World Bank/WHO Special Programme for Research and Training in Tropical Diseases. The Secretariat has

¹ Available at: <http://www.who.int/research-observatory/en/> (accessed 31 October 2016).

undertaken a study that presents some possible financing options that could be used to feed the voluntary pooled fund as requested in the resolution.¹

5. The Health Assembly further requested the Director-General in resolution WHA69.23 to submit to the Board for consideration at its 140th session the terms of reference for the Expert Committee on Health Research and Development whose establishment it also requested. The terms of reference are contained in the accompanying document EB140/22.

Global Observatory on Health Research and Development

6. The Global Observatory on Health Research and Development aims “to consolidate, monitor and analyse relevant information on health research and development needs of developing countries, with a view to contributing to the identification and the definition of gaps and opportunities for health research and development priorities”.²

7. In January 2016, a demonstration version of the Global Observatory was published online which incorporated data on: funding for health research and development (from Policy Cures Research’s Grant Finder survey³); health products that are under development (from four data sources); clinical trials (from the WHO International Clinical Trials Registry Platform); research publications (from PubMed); gross domestic expenditures on health research and development by country (from UNESCO, OECD, La Red de Indicadores de Ciencia y Tecnología – Iberoamericana e Interamericana and Eurostat) and other relevant country-level macroeconomic data such as total health expenditures (from WHO’s global health expenditures database) and burden of disease (from the Global Health Observatory).

8. Following consideration of feedback from users, the Global Observatory on Health Research and Development will be released in early 2017. It will include new elements, such as indicators for monitoring activities in and resources for health research and development, comprehensive analyses of data on health research and development by disease, comparative analyses of health research and development activities between countries and across diseases, and a section on classifications and standards, as a first step towards harmonization of future data collection efforts. The Observatory will continue to be expanded and updated as new resources and information become available.

HEALTH RESEARCH AND DEVELOPMENT DEMONSTRATION PROJECTS

9. Based on resolution WHA66.22, the following six demonstration projects were finally selected:

(a) the visceral leishmaniasis global research and development and access initiative (proponents: Drugs for Neglected Diseases initiative and United States Food and Drug Administration);

(b) exploiting the pathogen box: an international open-source collaboration to accelerate drug development in addressing diseases of poverty (proponent: Medicines for Malaria Venture);

¹ The results of the study will be made available at: <http://www.who.int/phi/implementation/research/en/>.

² Resolution WHA69.23, eighth preambular paragraph, available at: http://apps.who.int/gb/ebwha/pdf_files/WHA69/A69_R23-en.pdf (accessed 25 October 2016).

³ <http://www.policycuresresearch.org/g-finder/> (accessed 15 November 2016).

- (c) development of easy to use and affordable biomarkers as diagnostics for types II and III diseases (proponents: African Network for Drugs and Diagnostics Innovation, Chinese Network for Drugs and Diagnostics Innovation et al.);
- (d) development of a vaccine against schistosomiasis based on the recombinant Sm14, a member of the fatty acid-binding protein family: controlling transmission of a disease of poverty (proponent: Oswaldo Cruz Foundation, Brazil);
- (e) multiplexed point-of-care test for acute febrile illness (proponent: Translational Health Science and Technology Institute, India);
- (f) demonstration of the potential of a single-dose malaria cure of artemether-lumefantrine through reformulation in a nano-based drug delivery system (proponent: Council for Science and Industrial Research, South Africa).

10. In 2015, the Ad-hoc Committee for the Demonstration Projects/Global Health R&D Observatory¹ reviewed the workplans and budget proposals for five demonstration projects – (a) to (e) – as well as those of the Global Observatory on Health Research and Development, and recommended allocation of funding for the first year of implementation of those five demonstration projects and the Observatory. Letters of agreement were signed and funds were disbursed for demonstration projects (a) to (c) in 2015 and for demonstration project (d) in September 2016, using all available funds. The fifth demonstration project (e) and the Global Observatory on Health Research and Development have not received any funding so far. The sixth demonstration project (f) was only recognized as a demonstration project in April 2016 and will receive funding subject to approval by the Ad-hoc Committee and availability of funds.

LINKING MONITORING, COORDINATION AND FINANCING OF HEALTH RESEARCH AND DEVELOPMENT

11. In resolution WHA69.23, the Health Assembly requested a description of how the Global Observatory on Health Research and Development, the proposed Expert Committee on Health Research and Development and the Scientific Working Group of a voluntary pooled fund would work together in terms of specific disease examples.

12. The role of the Global Observatory on Health Research and Development is to provide the fundamental data needed to prioritize decisions on research and development. Such analyses would include data on the unaddressed public health needs for new products, ongoing research and development activities (such as products in the pipeline and clinical trials), investments, gaps, list of approved products, patents, and any established priorities for research and development for the diseases and conditions within its scope.

13. The analyses produced by Global Observatory on Health Research and Development will be used by the Expert Committee on Health Research and Development to recommend priority areas for research and development of specific health products and technologies; for example, the need for an innovative vaccine against pulmonary tuberculosis in adults, a condition that accounts for most cases

¹ See http://www.who.int/phi/news/adhoc_committee/en/ (accessed 31 October 2016).

of tuberculosis worldwide, or an accurate and specific, easy-to-use diagnostic tool that can be used in rural health settings.

14. The Scientific Working Group of the proposed pooled fund would then operationalize the priorities established by the Expert Committee by defining the detailed product characteristics. The interaction of this system is illustrated in the recent report on Health product research and development fund: a proposal for financing and operation.¹

PROMOTING AND ADVOCATING FOR SUSTAINABLE AND INNOVATIVE FINANCING

15. In resolution WHA69.23 the Health Assembly also requested the Director-General to promote and advocate sustainable and innovative financing for all aspects of the strategic workplan and to consider its inclusion in WHO financing dialogues.

16. The estimated total financial requirement over the period 2014–2017 for the implementation of the demonstration projects and establishment of the Global Observatory is US\$ 85 million.² As at 30 September 2016, a total of US\$ 2.26 million over the four years had been contributed or pledged by France, Germany, Switzerland, the United States of America and the European Commission to the Global Observatory, and a total of US\$ 7.17 million had been contributed by Brazil, India, Norway, South Africa and Switzerland to the voluntary fund designated for demonstration projects. Another US\$ 1.18 million has been contributed or pledged by Norway and Switzerland as matching grants for contributions from developing countries on the basis of half a dollar for each dollar contributed, and US\$ 1.40 million of Swiss matching funds are still available, pending developing country contributions.

17. In total, US\$ 72.99 million is required for all the selected demonstration projects to be completed as proposed and to finalize the development of the Observatory.

18. Given the importance of the subject matter, the specific nature of research and development funding in comparison to the funding of emergencies and other parts of WHO's Twelfth General Programme of Work, the Secretariat has decided to hold a specific high-level event between the 140th session of the Executive Board and the Seventieth World Health Assembly with the purpose of promoting increased investment into research and development funding in areas where the current investment levels are insufficient to meet global public health needs. The Secretariat is undertaking further fundraising for the demonstration projects and full development of the Global Observatory.

Promoting policy coherence

19. The Health Assembly in resolution WHA69.23 additionally requested the Director-General to promote policy coherence within the Organization. The Secretariat thus ensures that key principles agreed with respect to research and development as a follow up to the report of the Consultative

¹ Figure 4.1 in Health product research and development fund: a proposal for financing and operation. Geneva: World Health Organization on behalf of the Special Programme for Research and Training in Tropical Diseases; 2016, available at: http://www.who.int/tdr/publications/r_d_report/en/ (accessed 31 October 2016).

² See document A69/40, available at: http://apps.who.int/gb/ebwha/pdf_files/WHA69/A69_40-en.pdf (accessed 31 October 2016).

Expert Working Group are also applied in its other areas of engagement, namely in new initiatives such as the Research and Development Blueprint to foster research and development preparedness for infectious diseases with epidemic potential, or the Global Antibiotic Research and Development Partnership, a joint venture by WHO and the Drugs for Neglected Diseases initiative.

ACTION BY THE EXECUTIVE BOARD

20. The Board is invited to note the report and to provide guidance on future strategic directions.

ANNEX 1

TERMS OF REFERENCE OF THE GLOBAL OBSERVATORY ON HEALTH RESEARCH AND DEVELOPMENT

GOAL AND SCOPE

1. The establishment of a Global Observatory on Health Research and Development was mandated by the Sixty-sixth World Health Assembly in resolution WHA66.22 (2013).¹ In May 2016, the Sixty-ninth World Health Assembly in resolution WHA69.23 recognized the central role of the Global Observatory on Health Research and Development and re-emphasized its overall goal “to consolidate, monitor and analyse relevant information on health research and development needs of developing countries”.² The Global Observatory will do so by “building on national and regional observatories (or equivalent functions) and existing data collection mechanisms, with a view to contributing to the identification and the definition of gaps and opportunities for health research and development priorities, and supporting coordinated actions on health research and development.”²
2. The scope of data and analyses covered by the Global Observatory on Health Research and Development will prioritize information on “Type II and Type III diseases and on the specific research and development needs of developing countries in relation to Type I diseases, as well as needs for information on potential areas where market failures exist, and also on antimicrobial resistance and on emerging infectious diseases likely to cause major epidemics.”²
3. The Global Observatory on Health Research and Development functions as a centralized and comprehensive source of information and analyses on global health research and development activities, building on existing data and reports from a wide range of data sources and gathering new information, where needed and feasible, with the aim of enabling decisions on priorities in research and development.
4. WHO’s Global Observatory on Health Research and Development has the following terms of reference:
 - (a) to produce comprehensive analyses and syntheses of existing data and information on health research and development for specific health conditions (for example, tuberculosis, malaria and leishmaniasis) in order to identify gaps and opportunities for health research and development and inform the setting of priorities;
 - (b) to monitor and report on global trends related to health research and development, including investments in health research and development and capacity for health research at national level; registered clinical trials and health technologies that are under development (the research and development pipeline); as well as approved medicines;
 - (c) to benchmark and compare health research and development activities across countries and health conditions; for example, comparing investments in health research and development

¹ http://www.who.int/phi/resolution_WHA-66.22.pdf (accessed 31 October 2016).

² http://apps.who.int/gb/ebwha/pdf_files/WHA69/A69_R23-en.pdf (accessed 31 October 2016).

across countries and health conditions with indicators such as burden of disease, gross domestic product, gross domestic expenditures on health, and expenditures on other types of research;

(d) to contribute to improving data collection and sharing standards by making available classifications and terminologies it uses as a step towards consensus building and better harmonization of future data-collection efforts;

(e) to make all data and analyses available to the general public in a web portal;

(f) to conduct comprehensive analysis and syntheses of data based on the advice of the Expert Committee on Health Research and Development.

5. To ensure adequate expert input into the Global Observatory's technical areas of work and analyses, the Secretariat will work closely with its technical departments and their established expert groups and committees in order to develop and/or review analyses and syntheses produced by the Global Observatory. The comprehensive analyses will be produced in close collaboration with experts in the respective subject areas and supported by the Global Observatory's data.

6. In addition, the Secretariat will also seek regular user feedback on the Global Observatory's structure and outputs from national policy-makers, academia, WHO's technical experts and other international governmental organizations and global partnerships, WHO regional offices, civil society and industry stakeholders to continue improving the Observatory's functionality and usability in future versions.

7. The analyses and data provided by the Global Observatory will constitute a major resource for the work of the Expert Committee on Health Research and Development, which aims at providing technical advice to the Director-General on priorities for health research and development.

WORKPLAN AND BUDGET FOR THE GLOBAL OBSERVATORY ON HEALTH RESEARCH AND DEVELOPMENT

8. The estimated costs for continued development of the Global Observatory for the period 2016–2019 are shown in Table 1. Table 2 shows available funds for the two bienniums 2016–2017 and 2018–2019 and the funding gaps. The estimated total net cost for 2016–2019 is US\$ 6.3 million. The total net earmarked funds received or pledged (as of September 2016) are US\$ 1.5 million (net of programme support costs) for 2016–2017. Taking into account the allocated WHO programme budget funds from assessed contributions, the total gross funding gap for 2016–2017 is US\$ 1.7 million. Without additional financial contributions, the total funding gap is estimated at US\$ 2.6 million for 2018–2019.

Table 1. Summary of planned activities and costs for the period 1 January 2016 to 31 December 2019 (in US\$)

Budget item	Budget 2016–2017	Budget 2018–2019
Global Observatory on Health Research and Development portal	585 000	675 000
Research and development knowledge generation and dissemination	400 000	575 000
Total activity costs	1 085 000	1 250 000
Total staff costs	1 961 133	1 961 133
Total net* biennial costs	3 046 133	3 211 133

* Net of programme support costs at 13%.

Table 2. Available funding and funding gap for the years 2016–2019 (in US\$)

Source	Total 2016–2017	Total 2018–2019
Total earmarked funds* for the Observatory (net of programme support costs**)	1 502 842	
Programme budget: assessed contributions	1 000 000	1 000 000
Grand total net of programme support costs**	2 502 842	1 000 000
Funding gap net of programme support costs**	1 543 291	2 211 133
Total gross funding gap (including programme support costs**)	1 743 919	2 628 580
Total gross funding gap	1 700 000	2 600 000

* Funds received or pledged from France, Germany, Switzerland and the European Commission as of September 2016.

** Programme support costs at 13%.

ANNEX 2

A VOLUNTARY POOLED FUND: OPERATIONAL PLAN AND GOALS

1. As outlined in the study “Health Product Research and Development Fund: A Proposal for Financing and Operation,”¹ several conditions should be met in order to accelerate and fill the gaps in the research and development pipeline for diseases primarily affecting low- and middle-income countries:

- minimum size of a research and development fund with a disbursement of US\$ 100 million annually over a 10-year period;
- a diversified portfolio of 35–40 research and development projects, including shorter-term development projects (e.g. repurposing of existing medicines) as well as longer-term discovery efforts (e.g. development of new chemical entities);
- transparent operations, with clear objectives and non-political, evidence-based decision-making processes;
- a sustainable funding mechanism.

2. The pooled fund would focus on the priorities as defined by the Expert Committee on Health Research and Development through focused research and development projects and build up a mixed research and development project portfolio over time. Figures 5.7 and 5.8 of the report on health product research and development fund¹ illustrate the potential number of research and development projects and the potential associated costs for a US\$ 100 million annual fund. The model used predicts that the fund could support 39 projects from year 11 onwards. The report also illustrates the funding increase over time from a US\$ 15 million fund supporting seven selected projects in the first year to US\$ 100 million supporting 39 portfolio projects over an 11-year period.

3. Based on these assumptions, the mechanisms would fund several development projects (for example, funding phase III trials to bring a single intervention through to approval). If the fund becomes operational in 2017, the UNICEF/UNDP/World Bank/WHO Special Programme for Research and Training in Tropical Diseases estimates that three reformulated or repurposed medicines, one simple new chemical entity and one complex repurposed medicine may be launched by 2030 as a result of this investment.

OPERATION OF THE FUND

4. In its study, the Special Programme for Research and Training in Tropical Diseases formulated some basic principles:

- a simple, evidence-based process should be used for reviewing projects quickly and deciding which to incentivize;

¹ Health product research and development fund: a proposal for financing and operation. Geneva: World Health Organization on behalf of the Special Programme for Research and Training in Tropical Diseases; 2016, available at: http://www.who.int/tdr/publications/r_d_report/en/ (accessed 31 October 2016).

- projects that have the potential to deliver impact should be prioritized rather than those that only build research and development capacity;
- the operating cycle should be based on transparent, objective and non-political decision-making.

5. The Scientific Working Group would play a critical role in the operating model, being responsible for translating research and development priorities defined by WHO into a portfolio of projects. It would then be responsible for two main processes: further detailing the priorities to an actionable level; and managing the project portfolio and financing (including soliciting, selecting, monitoring and evaluating projects).¹ It would prepare calls for proposals on the basis of analyses of the existing pipeline and product profile characteristics. It would also prepare a recommendation on the most appropriate incentive/disbursement mechanism. Assisted by the Secretariat, the Scientific Working Group would monitor and review funded projects to measure progress and evaluate their impact potential.

AFFORDABILITY OF PRODUCTS COMING OUT OF THE MECHANISM

6. The pooled fund would operationalize some of the principles and recommendations formulated by the Consultative Expert Working Group, including the core principles of affordability, effectiveness, efficiency, equity and delinking the costs of the investment into research and development from the volume and price of the resulting health products. It would favour open collaboration and sharing of research and development results. Recipients of grants would have to adhere to these principles of transparency and knowledge sharing. Those who would bring products to the market that were developed with funding from the pooled fund would have to commit to an affordable pricing policy and to manage any intellectual property in a way which prioritizes access.

OPERATIONAL COSTS

7. An estimated additional operating cost of up to US\$ 7.6 million² including fund-hosting costs would be required to run the pooled fund (with a size envisaged to be US\$ 100 million) hosted by the Special Programme for Research and Training in Tropical Diseases.

OPTIONS FOR SUSTAINABLE FUNDING

8. The success of a voluntary pooled fund will depend on its ability to attract sufficient amounts of funding, with a minimum size of US\$ 100 million per year. As highlighted by the Expert Working Group on Research and Development Financing³ and the subsequent Consultative Expert Working Group on Research and Development: Financing and Coordination,⁴ there are very many possible ways available for reference by Member States to finance such a fund. To come up with some realistic options, the Secretariat is currently undertaking a study into financing options that could be used to

¹ This governance structure is exemplified in Figure 4.1 of the report on Health product research and development fund, available at: http://www.who.int/tdr/publications/r_d_report/en/ (accessed 31 October 2016).

² Representing an operational cost of between 6% and 8%.

³ The report can be found on: http://www.who.int/phi/publichearing_researchdev/en/ (accessed 2 November 2016).

⁴ The report can be found on: <http://www.who.int/phi/implementation/research/en/> (accessed 2 November 2016).

feed the voluntary pooled fund and the results will be made available on the WHO website before the 140th session of the Board.¹

9. Member States are invited to consider the options presented in the study, focusing on a mixed model that combines different instruments and involves different sources of funding. For example, Member States could commit themselves to contribute a certain amount through voluntary contributions. These extra voluntary contributions by Member States could be combined with matching funds contributed by the private sector. The remaining funds needed to reach an annual budget of US\$ 100 million could be financed through a financial instrument such as a social impact bond guaranteed by Member States or philanthropic research and development investors. Where there are buyers for future products, advance market commitments could be another option to generate funds. Where eligible products are being developed, revenues from selling priority review vouchers could be another source of income of the fund. The last two options can, however, only be used with respect to specific products and projects and would still require a loan to finance the initial research and development cost and to bear the risk of failure.

10. Member States could also consider setting up a fund that would invest its revenues in health research and development, following the example of the Australian Medical Research Future Fund. A fund amounting to US\$ 2000 million could guarantee an annual 5% investment, namely US\$ 100 million, into research and development.

11. In addition, Member States could opt to use a replenishment model, in which each donor independently fixes its contribution in the form of a pledge. This model is less sustainable than creating a mechanism that creates revenue. An example of the latter is the introduction of a fee or levy on marketing activities of pharmaceutical companies, following the Italian model. The Italian Government introduced a 5% levy on the promotional expenditures of pharmaceutical companies. The resulting revenue has been used to fund an independent research and development programme on pharmaceuticals; this represented about €78 million between 2005 and 2007. These options will be further described in the study that is currently undertaken by the Secretariat.

12. In any case, the pooled fund should also be able to accept voluntary, preferably unspecified, funding from non-State actors, such as philanthropic foundations, following WHO's rules on acceptance of donations. The Special Programme for Research and Training in Tropical Diseases would ensure appropriate management of conflicts of interest in subsequent decisions on the allocation of funds, in other words ensuring that such contributions have no impact on the process of selection of projects.

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¹ Information will be made available on the WHO website, at: <http://www.who.int/phi/implementation/research/en/>.