

### Research during outbreak response:

from Ebola to the COVID-19 pandemic

### Ana Maria Henao Restrepo MD, MSc

Unit Lead, R&D Blueprint
WHO Health Emergencies Preparedness & Response
World Health Organization

February, 2021



### A consolidated approach to combating epidemics

Preparing for the inevitable

Improved regulatory pathways

Equitable data and sample sharing including of benefits

Develop research infrastructure

Conducting research during epidemics

More innovative approaches to funding essential work

Sustainability of funding and research capacity during inter-epidemic periods

Established surveillance systems

Resources at critical stages of the early development process

A more productive, integrated approach to research

Improved regulatory pathways

**Expand clinical trials infrastructure** 

Strengthening local research capacity

Understanding of the funding landscape: Timely tracking for course correction

Ethics oversight to facilitate research

Resilient health systems

**Build true global partnerships** 

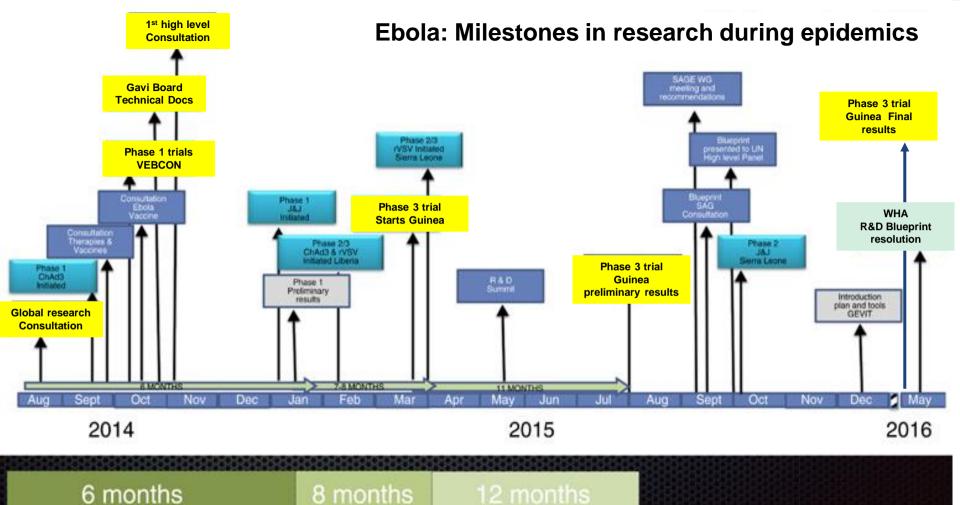


# A central and historic responsibility for WHO has been the management of the global regime for the control of the international spread of disease

The Constitution of WHO confers upon the WHA the authority to adopt regulations "designed to prevent the international spread of disease" which, after adoption by the WHA, enter into force for all WHO Member States

- 1951 International Sanitary Regulations
- 1969 The International Health Regulations
- **2003** WHA requested their revision following the emergence of severe acute respiratory syndrome (the first global public health emergency of the 21st century).
- 2005 revised IHR were adopted by the WHA
- 2007 revised IHR entered into force (15 June 2007)

Adequate preparedness for pandemics is a national obligation under the International Health Regulations (2005)





# **R&D**Blueprint

Powering research to prevent epidemics

At the request of its 194 Member States in 2015, WHO has convened a broad network of experts to develop an R&D Blueprint for Action to Prevent Epidemics

In the event of an outbreak, R&D Blueprint activities shift from R&D preparedness to an emergency response plan





#### DISEASES TO BE URGENTLY ADDRESSED UNDER THE R&D BLUEPRINT, AS OF MAY 2016







Filovirus diseases (i.e. EVD & Marburg)

Highly pathogenic emerging coronaviruses relevant to humans (MERS Co-V & SARS)

Lassa fever virus







a new severe infectious disease

### A consolidated approach to combating epidemics requires research preparedness and coordinated and sustained investments

Yet there remains a shortfall in funds even for the current pandemic

Health systems support and investments in research during interepidemic periods are also insuficient

#### SERIOUS DISEASES NECESSITATING FURTHER ACTION AS SOON AS POSSIBLE. AS OF MAY 2016



Chikunguya virus



Severe fever with thrombocytopenia syndrome



Congenital abnormalities and other neurological complications associated with Zika



### Research during epidemics

The R&D Blueprint systematic approach to accelerate research and address gaps

DISEASE	Generic/ methodol.	CCHF	Ebola & Marburg	Lassa fever	MERS-Cov & SARS	Nipah & henipavirviru ses	Rift valley fever	Zika virus	Pathogen X (COVID-19)	Plague	Chikungunya
R&D roadmap	<b>√</b>	✓	<b>√</b>	✓	<b>√</b>	<b>√</b>	✓	<b>√</b>	<b>√</b>		✓
TPP Vx	✓		<b>√</b>	✓	✓	✓	<b>√</b>	✓	✓	✓	
ТРР Тх	✓								✓		
TPP Dx	✓	✓	<b>√</b>	✓		✓		✓	✓		
Regulatory standards	<b>√</b>	✓	✓	✓		✓		✓	✓		
Vx trials design	<b>√</b>	✓	✓	✓	✓	✓	<b>√</b>	✓	CORE	✓	<b>√</b>
Tx trials design	<b>√</b>	✓	✓	✓	✓	✓	<b>√</b>		CORE	✓	✓
Decision tree design	✓	$\checkmark$						✓			
Trial simulator	✓	✓					✓				
Innovative analysis	✓	Accumulating evidence from randomized clinical trials across outbreaks						<b>√</b>			





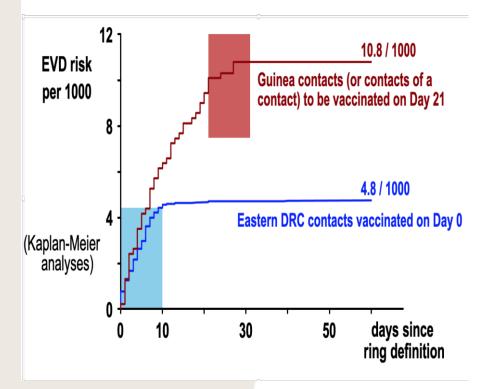
# Since 2016, rVSV ZEBOV has been deployed using Expanded Access/Compassionate Use as part of the response in 6 outbreaks

Country/year	Days start after outbreak declaration	Number vaccinated
Guinea/Sierra Leone (2016)	7	1510
Equator, DRC (2018)	7	5000 +
Eastern Congo, DRC (2018-2020)	7	300,000+
Equator, DRC (2020)	7	40,000
Eastern Congo DRC (2021)	8	325 (ongoing)
Guinea (2021)	8	Starts tomorrow



### EVD onset in rings defined in Guinea (2015) and Eastern Congo (2018-19)

Shading indicates the first 10 days after vaccination, when little, if any, efficacy is expected. This non-randomised comparison suggests vaccine efficacy about 90-95% in Eastern Congo against late EVD onset (in days 10-84)





A more productive, integrated approach to research during epidemics would encompass many disciplines and involve pursuit of innovative study designs and aligned and expedient regulatory pathways





### "It will take enlightened, brave leadership and concerted action to promote multilateral, cooperative solutions. [...] Doing the right thing, doing the thing that is appealing from a

humanitarian perspective, is also the efficient thing where ending the pandemic is concerned."

R. Hatchett, CEPI



The COVID-19 Research GloPID-R Synergies Meeting Working Group BMC Medicine **BMC Medicine** https://doi.org/10.1186/s12916-020-01807-3 Ending COVID-19: progress and gaps in research—highlights of the July 2020 GloPID-R COVID-19 Research Synergies

The COVID-19 Research GloPID-R Synergies Meeting Working Group 123,456,7 and Meeting Co-chairs 89,10,11,1213,14

Keywords: CDVID-19. SARS-CoV-2. GloPID-R. COVID-19 vaccines. COVID-19 therapeutics. SARS-CoV-2 transmission.

Hamilton Canada ISARC, Oxford University, Oxford, UK

Meetings

In mid-luly, GloPID-R convened its members, scientists 19 and building on a collaborative approach between rewho had received COVID-19 funding, and world leaders searchers and institutions to achieve health equity, uniin pandemic preparedness and response for a series of versal access and data sharing is critical to reduce the COVID-19 Research Synergies Meetings with the purpose of establishing collaboration and identifying knowledge gaps, in order to build a collective path forward to end COVID-19. GloPID-R is an international consortium of 29 funding organisations invested in research related to new or re-emerging infectious diseases [1]. By offering a coordination platform, GloPID-R aims to maximise the use of available resources and streamline research efforts amongst national and global funders. The COVID-19 pandemic has caused unprecedented devastation to the health and economies of countries across the world and has highlighted basic inequalities

The COVID-19 Research GioPID-R Synergies Meeting Working Group:

Geneviève Bolly-Larouche, Gall Carson, Josie Golding, Evelyn Depoortere, Info Rannel de Almeiria, Richard Vaux, Gusenne Panarella, Danielle Vitali.

Deborah Khursigara, Claire Madelaine, A. Morgan Lav. Rarbara Kensténs, Yazdan Yazdanganah, Charu Kauthir Meeting Co-chain: Anita Zaidi, Melanie Saville, Debra Yeskey, Glenda Gray,

Valdida Veloso, Marion Koopmans, David Risman, Kristy Crooks, Kenneth

Full list of author information is available at the end of the article

Canadian institutes of Health Research – Institute of Infection and Immunity.

impact of the COVID-19 pandemic. "Lockdowns, in my view, have been successful in helping to reduce transmission and new infections. but they haven't changed the fundamentals of this infection. The virus biology, its transmission, how infections it is the clinical syndrome it causes the

in society. Understanding the fundamentals of COVID

inequalities it drives through the world, and of course the tertiary consequences in economics and in geopolitics. So, I think we are only very much at the start and we need to be humble about the challenge we face. We do not yet have a clear exit strategy and we need to define one." J. Farrar, Wellcome

#### Ending COVID-19: vaccine

Balancing speed, scale and access.

Presenters highlighted the progress in COVID-19 vaccine development, regulatory considerations, as well as gaps in advancing the demonstration of clinical efficacy and effectiveness and challenges that must be overcome to meet the global demand for equitable access (Table 1).



Research funders recognise that there is a need to coordinate COVID-19 research funding at all levels to prevent duplication and improve impact. This coordination should not be limited to the COVID-19 pandemic, but instead needs to facilitate ongoing improved coordination for future infectious disease epidemics and pandemics.

**eglopid-R eukcor** 

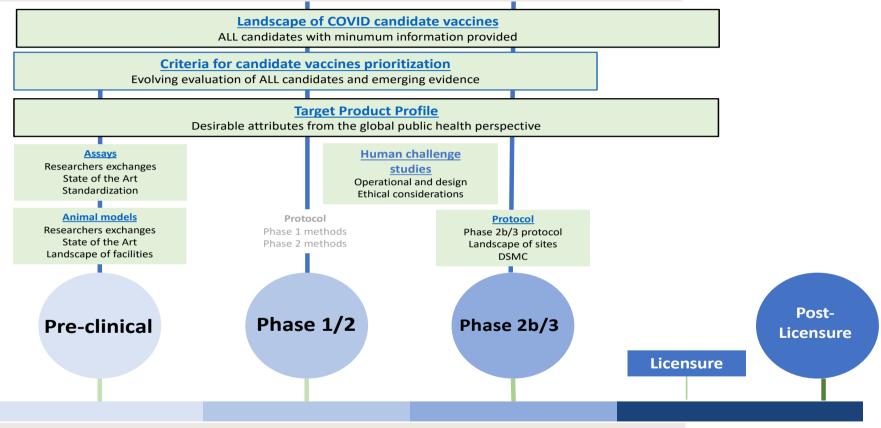
The UK Collaborative on Development Research (UKCDR) and the Global Research Collaboration for Infectious Disease Preparedness [GloPID-R] have agreed a set of principles to align research funders towards a coordinated effort for supporting high-quality research for the most pressing global needs in epidemics and pandemics.

FUNDER PRINCIPLES FOR SUPPORTING HIGH-QUALITY RESEARCH FOR THE MOST PRESSING GLOBAL NEEDS IN EPIDEMICS & PANDEMICS





### WHO's contribution to accelerate COVID-19 vaccines evaluation







# The world needs efficient, speedy, and reliable evaluation of more candidate vaccines against COVID-19.

70 candidates in clinical phase

181 candidates in pre- clinical phase

## Schedule and route of administration of candidates in clinical phase

Schedule       Candidate vaccines (no. and %)         1 dose       13       19%         Day 0       13       2         2 doses       42       60%         Day 0 + 14       6       6         Day 0 + 21       16       6         Day 0 + 28       20       3         3 doses       1       1%         Day 0 + 28 + 56       1       20%         TBD / No Data (ND)       14       20%         Route of administration       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%         TBD / No Data (ND)       10       14%						
Day 0       13         2 doses       42       60%         Day 0 + 14       6       6         Day 0 + 21       16       16         Day 0 + 28       20       3 doses       1       1%         Day 0 + 28 + 56       1       20%       70         Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	Schedule		Candidate vaccines (no. and %)			
2 doses	1 dose		13	19%		
Day 0 + 14       6         Day 0 + 21       16         Day 0 + 28       20         3 doses       1       1%         Day 0 + 28 + 56       1       20%         TBD / No Data (ND)       14       20%         Route of administration       70         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	Day 0		13			
Day 0 + 21       16         Day 0 + 28       20         3 doses       1       1%         Day 0 + 28 + 56       1       20%         TBD / No Data (ND)       14       20%         Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	2 doses		42	60%		
Day 0 + 28       20         3 doses       1       1%         Day 0 + 28 + 56       1       20%         TBD / No Data (ND)       14       20%         Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	Day 0 + 14		6			
3 doses       1       1%         Day 0 + 28 + 56       1       1         TBD / No Data (ND)       14       20%         Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	Day 0 + 21		16			
Day 0 + 28 + 56       1         TBD / No Data (ND)       14       20%         70         Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	Day 0 + 28		20			
TBD / No Data (ND)     14     20%       70       Route of administration       Oral     2     3%       Injectable     58     83%       SC     2     3%       ID     3     4%       IM     53     76%	3 doses		1	1%		
70   Route of administration   2   3%   Injectable   58   83%   SC   2   3%   ID   3   4%   IM   53   76%	Day 0 + 28 + 56		1			
Route of administration         Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%	TBD / No Data (ND)	)	14	20%		
Oral       2       3%         Injectable       58       83%         SC       2       3%         ID       3       4%         IM       53       76%			70			
Injectable         58         83%           SC         2         3%           ID         3         4%           IM         53         76%	Route of administra	ation				
SC     2     3%       ID     3     4%       IM     53     76%	Oral	Oral		3%		
ID 3 4% IM 53 76%	Injectable		58	83%		
IM 53 76%	SC	SC		3%		
	ID		3	4%		
TBD / No Data (ND) 10 14%	IM		53	76%		
	TBD / No Data (ND)		10	14%		



# A coordinated global approach is critical WHO must continue to facilitate coordination

An effective response to new variants will depend on continued collaboration.

WHO's is establishing a risk assessment framework for SARS-CoV-2 variants.

If a range of vaccine products are required, this could exacerbate supply issues.

 Therefore decisions on need for modified or new vaccines should be the outcome of global coordinated efforts.

To address the threat of new variants, new vaccines may be needed, or existing platforms could be modified.

 WHO and regulatory authorities are working to achieve regulatory alignment for vaccines targeting new variants and efforts to coordinate approaches globally.

