



COVID-19 : Lessons for Public Health and Clinical Practice

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Global Situation

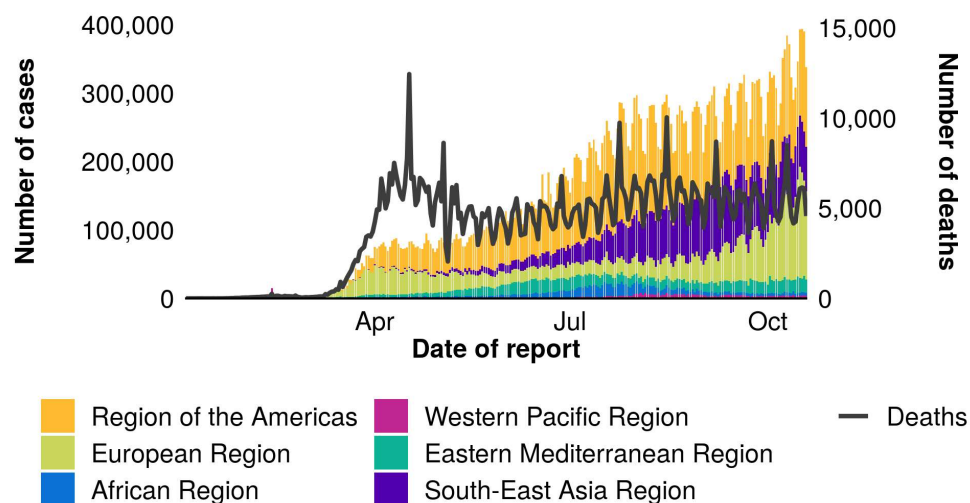
(as of 19 October 10H CEST)

Previous 24 hours:

- 338,096 new confirmed cases.
- 4,556 new deaths.

Cumulative:

- 39,944,882 confirmed cases.
- 1.1 million deaths.

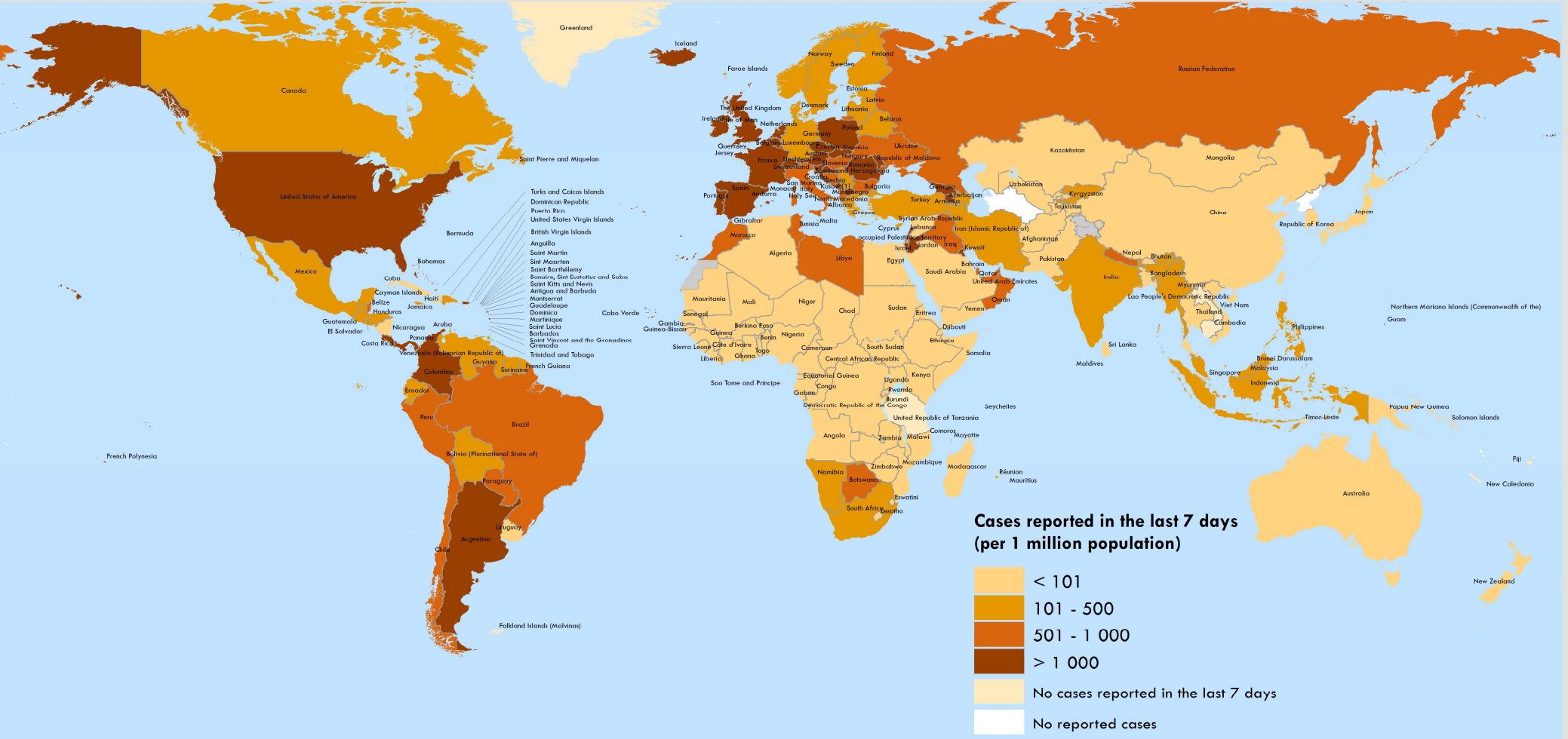


Countries with the highest number of new cases in previous 24 hours

Country		New Cases	Total Cases	New Deaths	Total Deaths
India		55,722	7,550,273	579	114,610
United States of America		52,508	8,019,237	588	217,659
France		29,833	867,978	85	33,204
Brazil		24,062	5,224,362	461	153,675
United Kingdom		16,981	722,413	67	43,646
Russian Federation		15,982	1,415,316	179	24,366
Argentina		13,510	979,119	384	26,107
Italy		11,705	414,241	69	36,543
Poland		8,536	175,766	49	3,573
Netherlands		8,439	228,234	23	6,751

COVID-19 cases reported in the last 7 days per 1 million population

(from 12 October 2020, 10:00AM to 18 October 2020, 10:00AM (CEST))



Data Source: World Health Organization,
United Nations Population Division (population prospect 2020)
Map Production: WHO Health Emergencies Programme

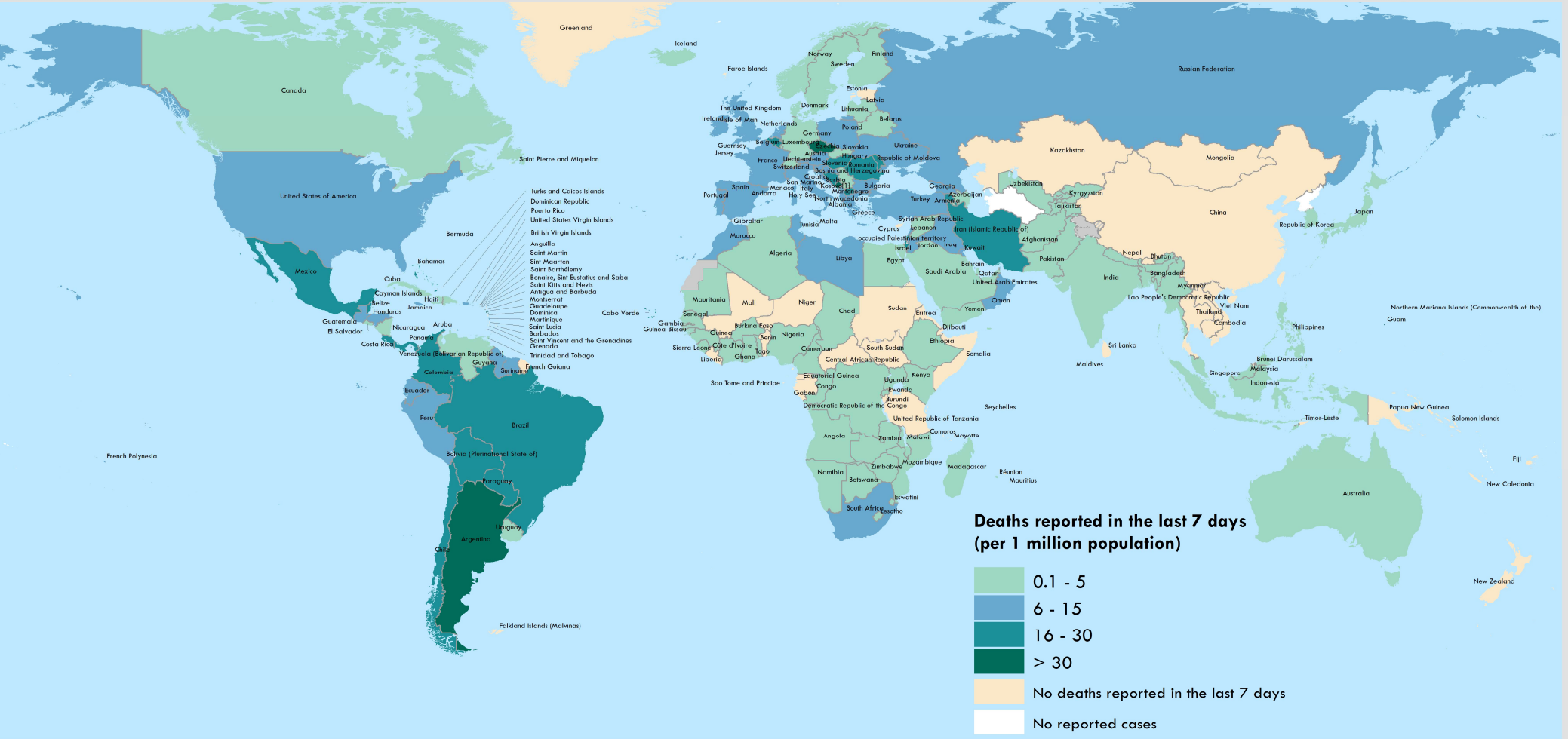
Not applicable

0 2,500 5,000 km
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COVID-19 deaths reported in the last 7 days per 1 million population

(From 12 October 2020, 10:00AM to 18 October 2020, 10:00AM (CEST))



Data Source: World Health Organization,
United Nations Population Division (population prospect 2020)
Map Production: WHO Health Emergencies Programme

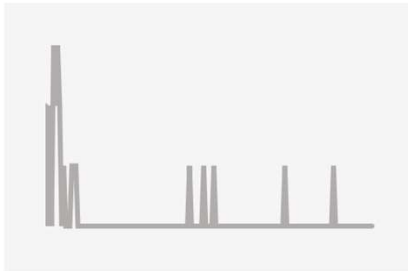
Not applicable

0 2,500 5,000 km
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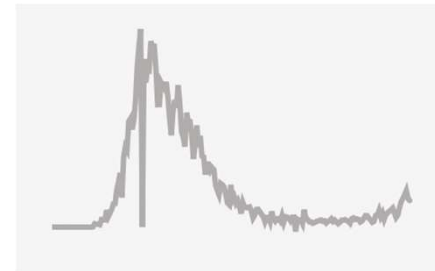
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COVID-19: currently facing different transmission situations

Avoided large outbreaks



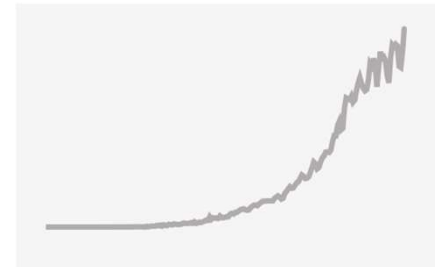
Major outbreak brought under control



Major outbreak brought under control, but having eased restrictions, are now seeing resurgence



Intense transmission



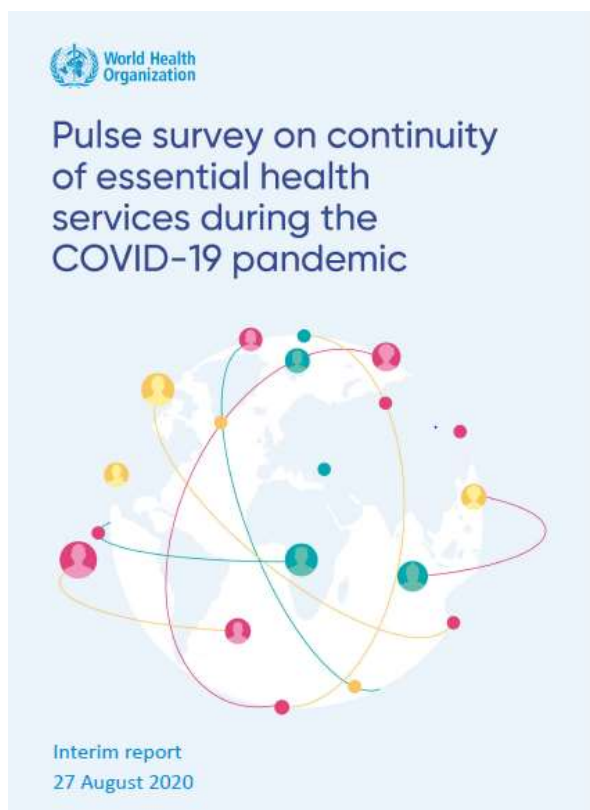
COVID-19 Challenges and Lessons Learned

Health systems facing COVID-19 outbreak

- Huge stress posed by increasing demand for care of people with COVID-19
- Lack of supplies and equipment
- Burden on health care workers
- Disruption of essential health services

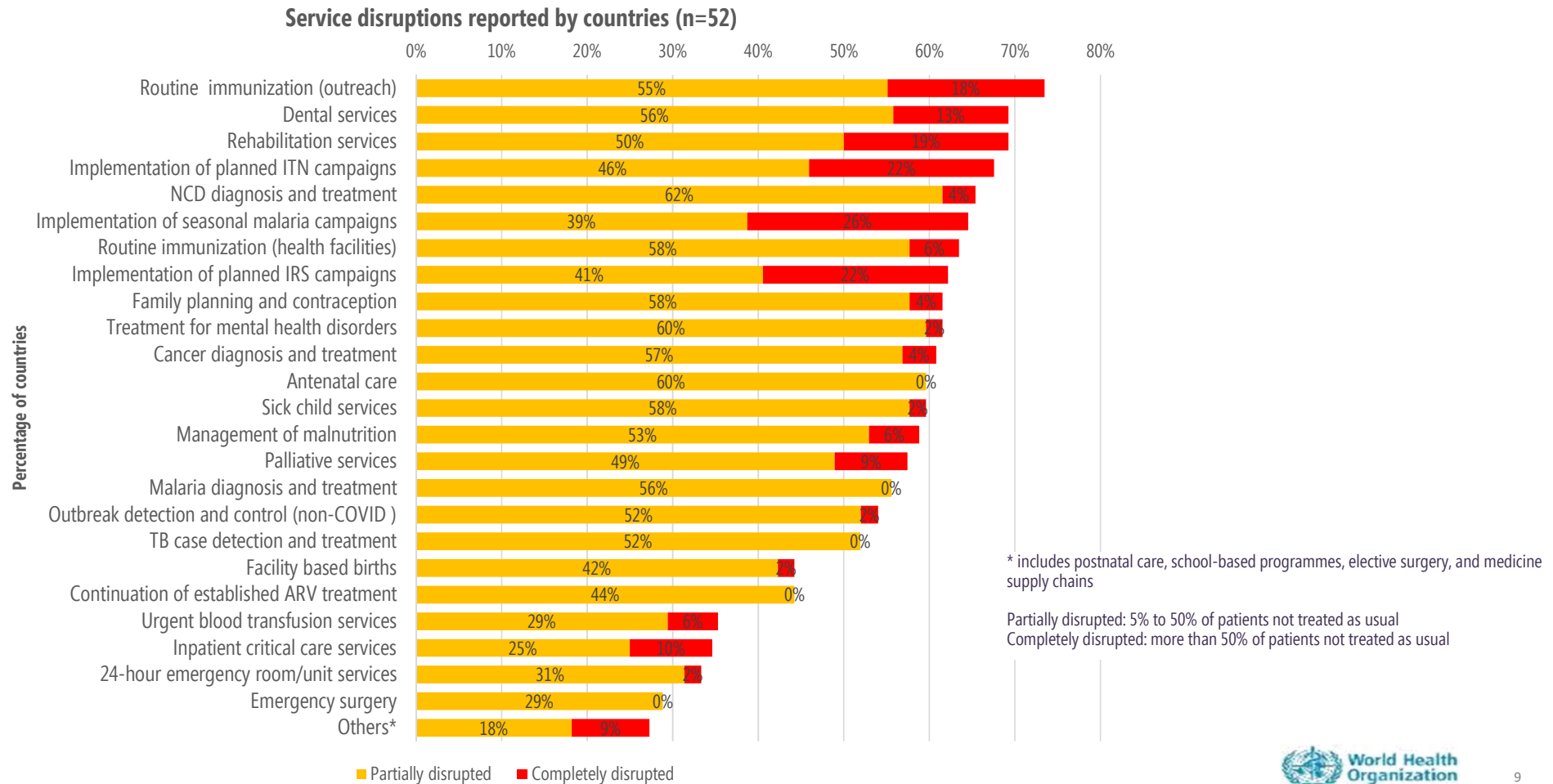
Dilemma: balance the demands of responding to COVID-19 with strategic planning and coordinated action to maintain quality essential health services

Pulse survey on continuity of essential health services during the COVID-19 pandemic

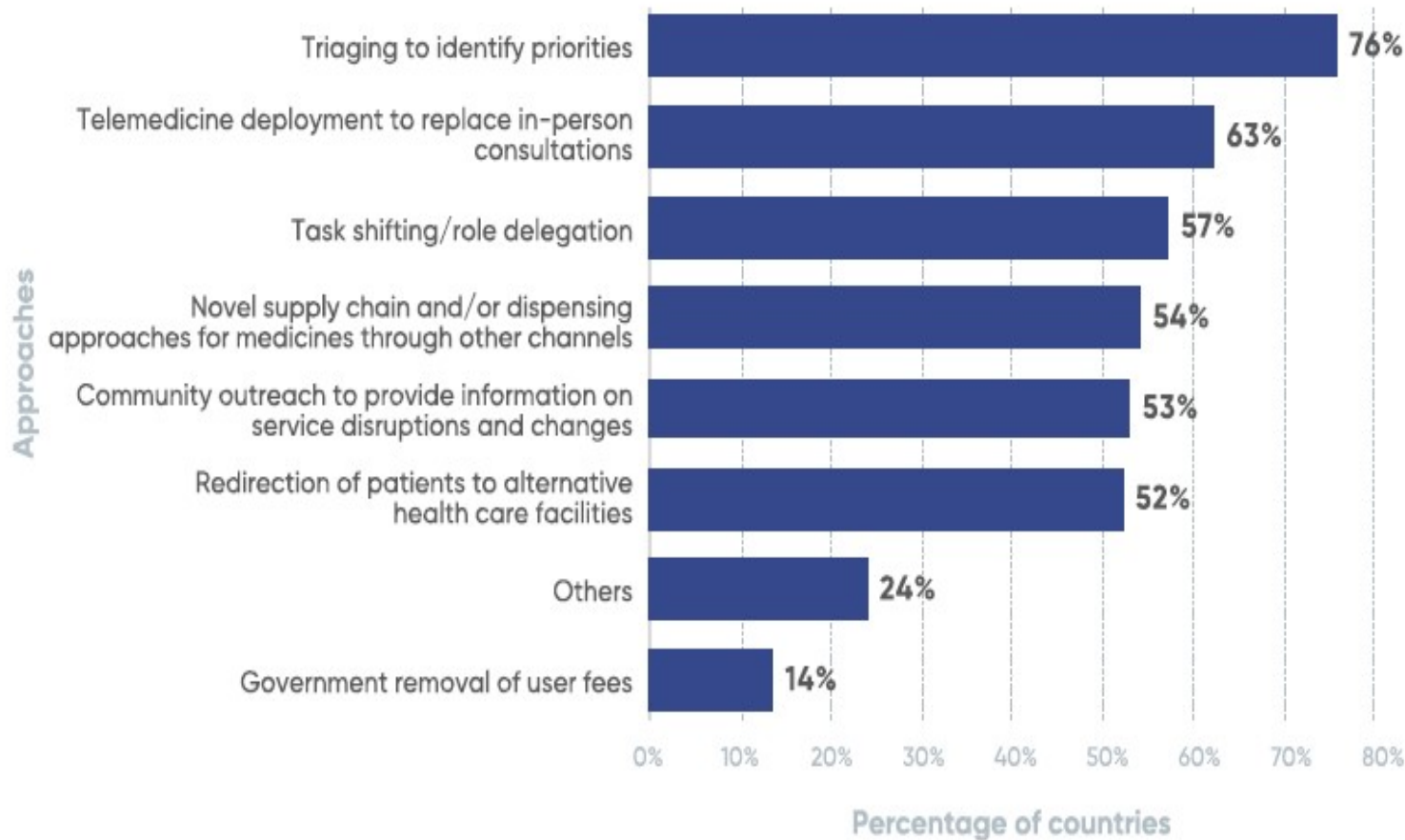


- 80% of countries had a defined essential health services package prior to the outbreak
- 66% of all countries had identified a core set of services to be maintained during the COVID-19 pandemic
- The Eastern Mediterranean Region was the most affected, followed by the African and the South-East Asia Regions. Essential health services in the countries in the European and the Western Pacific Regions were least affected
- **Most frequently disrupted Essential health services** were routine **immunization** services – outreach services (70%) and facility-based services (61%) – **noncommunicable disease diagnosis and treatment** (69%), **family planning and contraception** (68%), **treatment for mental health disorders** (61%), **antenatal care** (56%) and **cancer diagnosis and treatment** (55%).

Countries reporting disruptions (partially or completely) across 25 essential health services

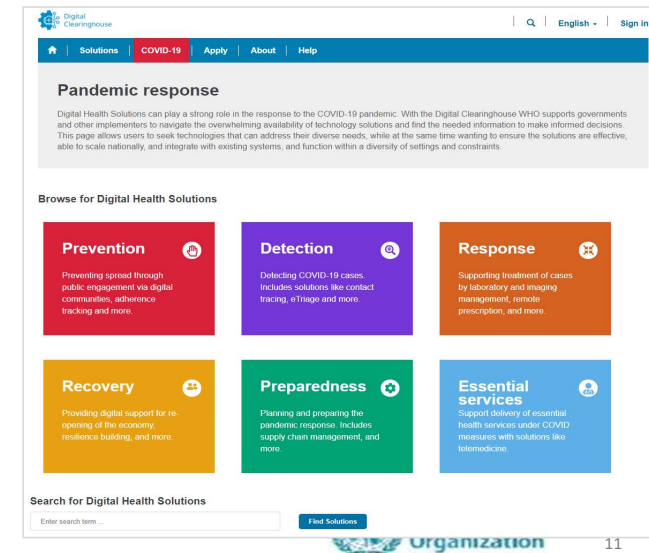
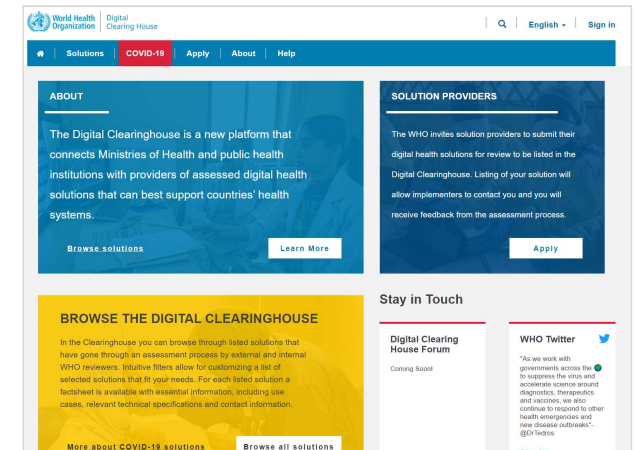


Approaches for overcoming health services disruptions



Digital Health technologies

- Telemedicine
- Global Learning Collaborative Platforms (eg ECHO)
- Electronic, portable health records
- What technologies can be considered public health goods?
- WHO Digital Health Clearing House (A digital platform that connects **Ministries of Health** with **providers of assessed digital health solutions** that can best support **countries' response to COVID-19 and other health domains**)



A new paradigm - (Virtual) Shared Medical Appointments



BENEFITS FOR PATIENTS

- Improved patient engagement
- Higher patient knowledge & satisfaction
- Shorter waits for appointments
- Higher compliance to medications
- Higher follow-up rates over time as patients form bonds with their peers
- Improved clinical outcomes
- Reduced flare-ups and emergency room visits



BENEFITS FOR PROVIDERS

- Higher productivity
- Expanded access
- Reduced cost of no shows
- Reduced long-term patient costs
- Reduced repetition → higher quality interactions
- Greater doctor satisfaction
- Lower environmental footprint

COVID-19 Country Case Studies

- Strong political leadership and successful mobilization of resources using a whole-of-society approach
 - Gain the trust of the public and ensure public compliance with public health measures
 - Whole-of- Government response to the pandemic
- Government cash assistance to people unemployed by the lockdown
- Early activation of a strong response system
- Long-term country investment to strengthen the health emergency response after previous epidemics and essential public health services
- Early contact tracing, quarantine and travel restriction measures
- Assessment of risks (existing and projected) at all levels of healthcare facilities
- Identifying extra 'surge' capacity each healthcare facility needs to cope with COVID-19 and non-COVID-19 related health emergencies



Figure: The message of the King of Bhutan on the website's homepage of the Royal Government of Bhutan.



Figure: mobile COVID-19 testing laboratory Morocco.



COVID-19: THAILAND
Suvarnabhumi International Airport



Doctors and nurses of a major hospital in Viet Nam, Bach Mai hospital, expressed their commitment and solidarity with the rest of the world in the fight against COVID-19. Photo credit: WHO Viet Nam

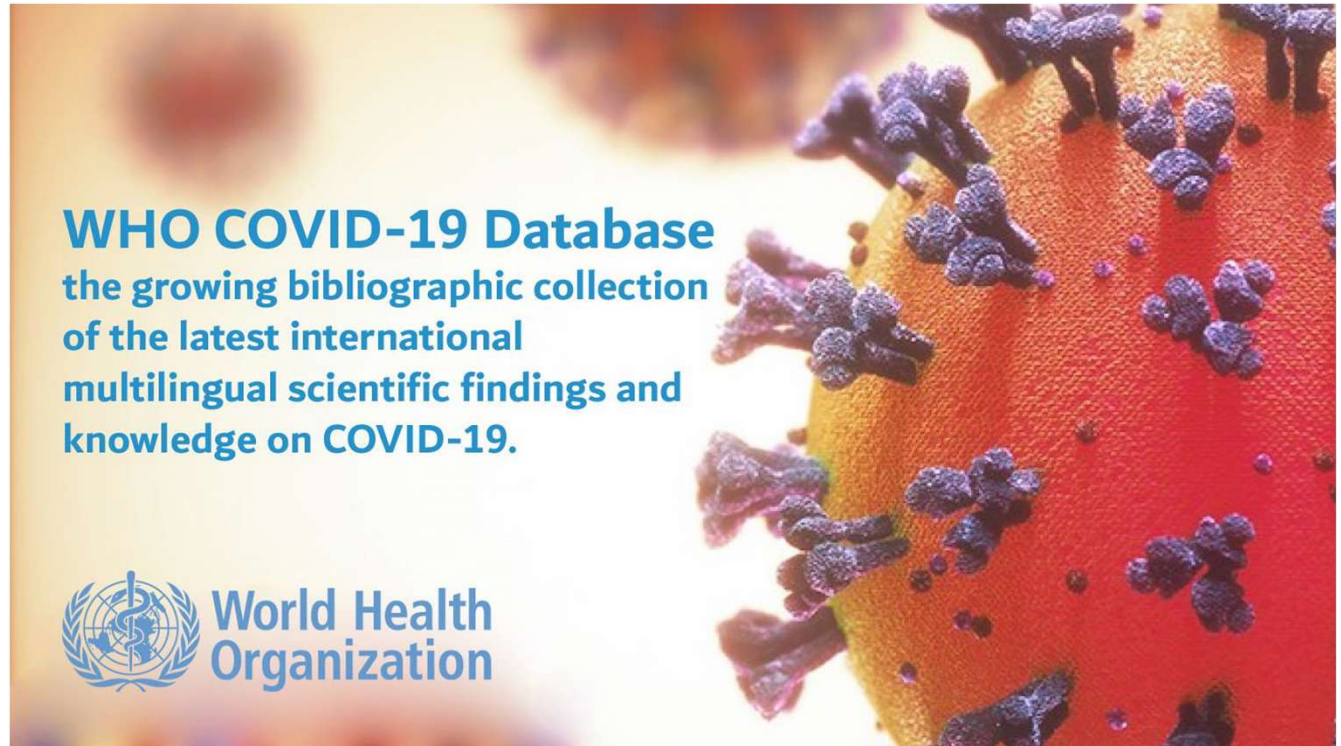


WHO providing on the job support to Seke District RRT team at Beatrice Isolation Centre in Mashonaland East Province. Here, WHO is helping them enter new cases into the ODK platform



WHO COVID-19 Database of scientific publications

- ✓ 90,000 citations and growing since March 2019
- ✓ Extended scope with inclusion of preprints and clinical trials
- ✓ More powerful software for screening and generating backbone of content
- ✓ 3000+ active users/day
- ✓ User feedback on one-stop convenience



<https://search.bvsalud.org/global-literature-on-novel-coronavirus-2019-ncov/>

WHO's normative function: leading policy & technical guidance

➔ Steering policy through:

- HQ-Regional leadership
- 45 Global Health Leaders
- STAG-IH

➔ Convening experts for guidance development:

- Technical guidance published by WHO

- Surveillance
- Clinical management
- Laboratory
- Supply & logistics
- Modeling
- Infection prevention & control

**hundreds
experts
100s calls**

Critical preparedness, readiness and response actions for COVID-19

Surveillance, rapid response teams, and case investigation

Surveillance, rapid response teams, and case investigation

Country-level coordination, planning, and monitoring

Clinical care

Infection prevention and control/WASH

The Unity Studies: Early Investigations Protocols

Essential resource planning

Guidance for schools, workplace and institutions

Risk communications and community engagement

Virus origin/Reducing animal-human transmission

Points of entry/mass gatherings

As of 12 May 2020

Update on published SARS-CoV-2 seroepidemiology

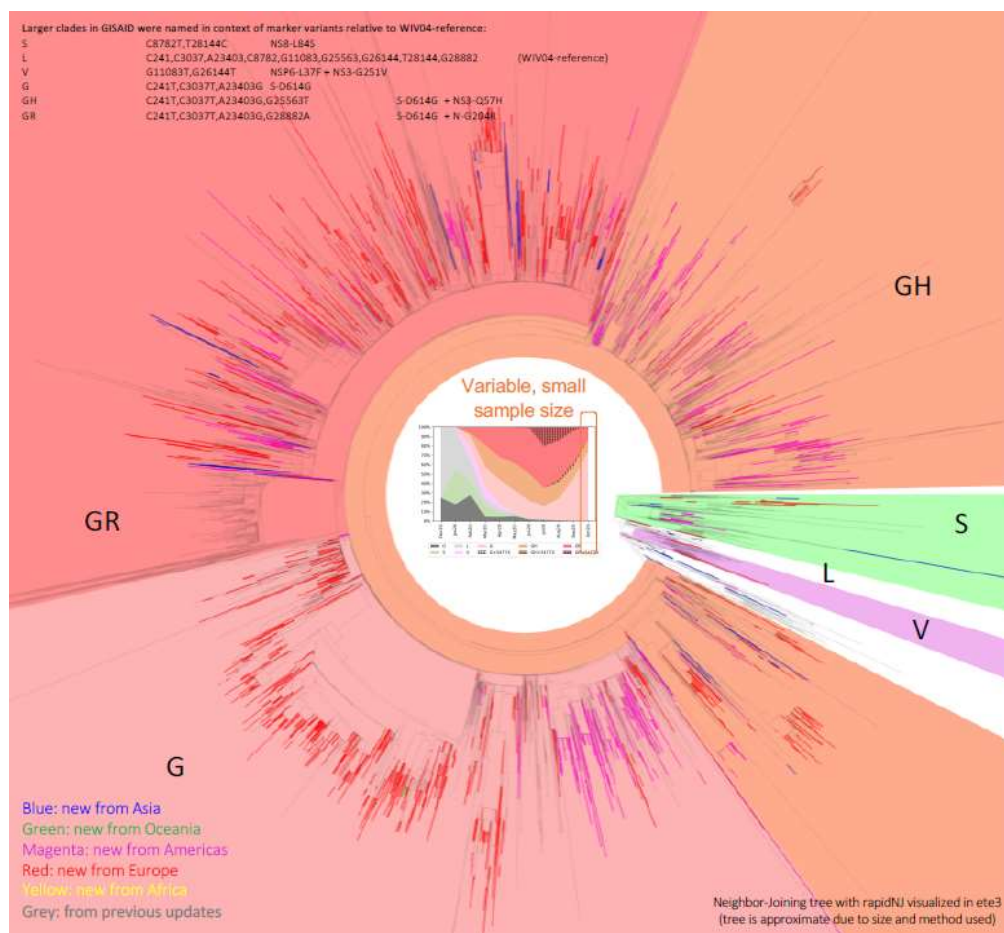
As of end September 2020:

- **>140** peer-reviewed publications and pre-print manuscripts plus **>10** government reports

Available results

- Most study results suggest <10% seroprevalence
- Few studies suggest >20% in areas of higher transmission intensity and/or among frontline workers
- Serial sampling studies show an increasing fraction of the population has been exposed to SARS-CoV-2
- Few studies have stratified results by age or by the same age categories;
 - available studies suggest that younger children appear to have lower rates of infection when compared to adults; adolescents appear to have similar seroprevalence to adults
- Longitudinal cohort and individual studies are limited, so there is currently limited information on persistence of immunity

Full genome tree derived from all outbreak sequences 2020-10-20



Full genome tree derived from all outbreak sequences 2020-10-20

Notable changes:

138,569 full genomes (+2,864) (excluding low coverage, out of 148,257 entries)

Updated clades:

S clade 6,417 (+59)
L clade 4,214 (+33)
V clade 5,258 (+7)
G clade [#S477X] 32,192 [101] (+1,221 [+9])
GR clade [#S477X] 54,064 [7,357] (+887 [+6])
GH clade [#S477X] 32,574 [671] (+644 [+53])
Other clades 3,850 (+13)

We gratefully acknowledge the Authors from Originating and Submitting laboratories of sequence data on which the analysis is based.

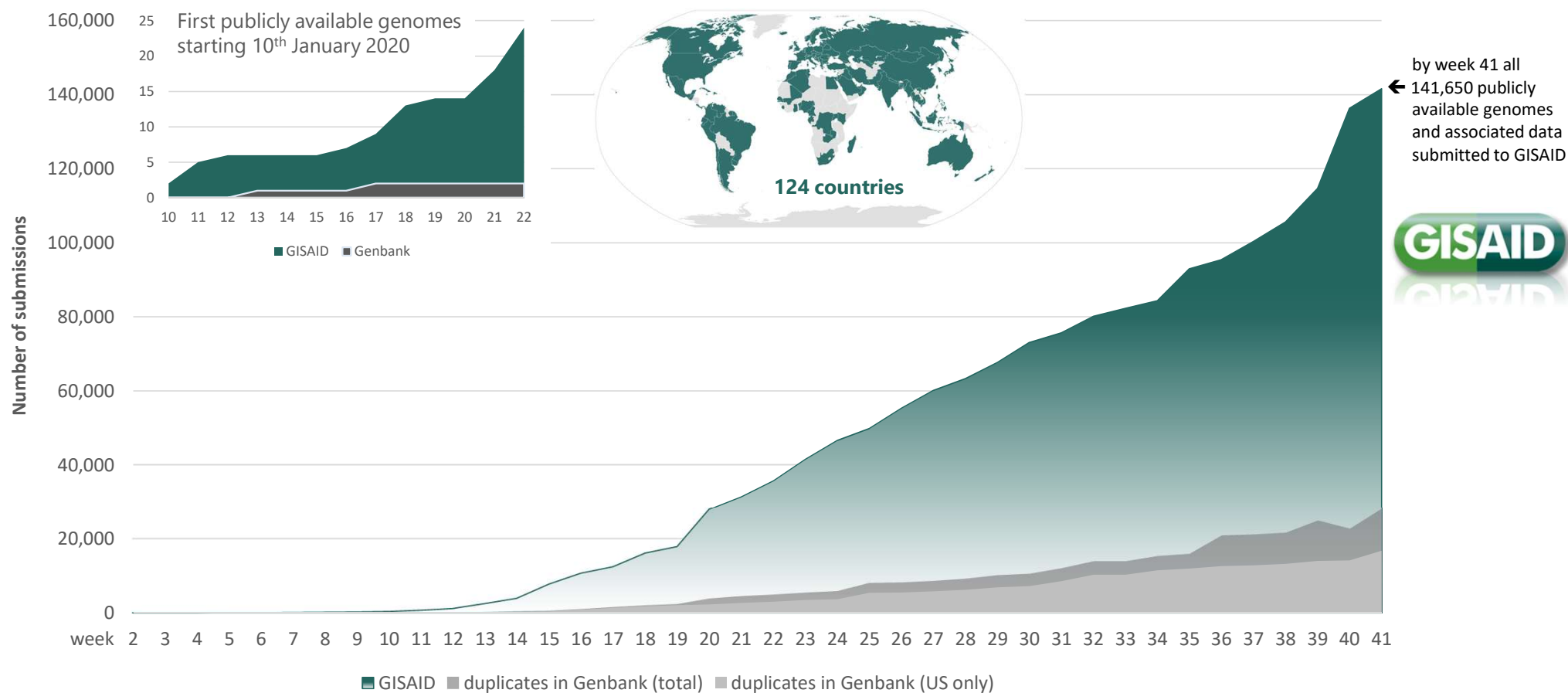


by BII/GIS, A*STAR Singapore



Real-time data sharing is not achieved by governmental Regulations

... it is incentivized by the confidence in transparent sharing mechanisms



Genomic sequence data is important for the response

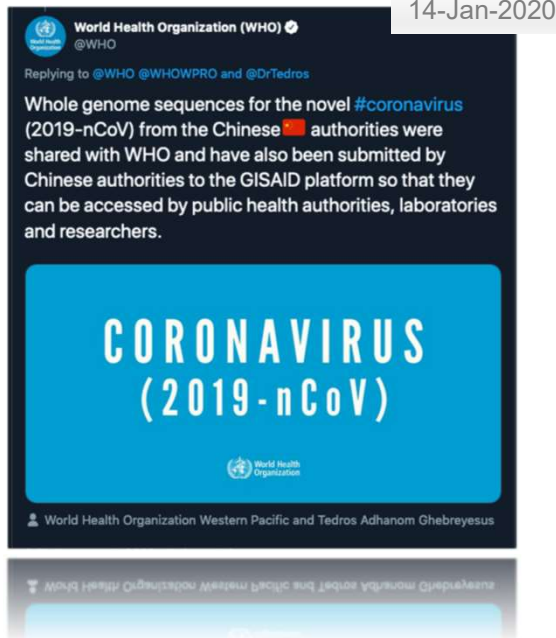
- Development of first diagnostics kits and refinement through ongoing surveillance for mutations
- Identification of potential drug and vaccine targets through repurposing
- Genomic epidemiology allows analysis of the exportation and importation events of viruses between countries, contact-tracing in countries, or identification of nosocomial transmission chains
- Evidence that the virus has not drifted to significant strain difference, with in particular the cell receptor binding pocket appearing stable
- Identification of animal precursors (in bats and pangolins)

GISAID was ready for Disease X ==> newly emerging coronavirus



Viral pneumonia cases reported in central China

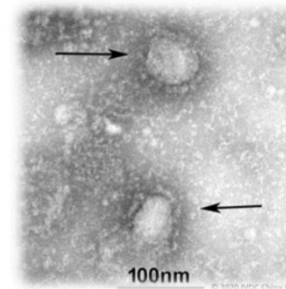
肺炎
Viral pneumonia cases reported in central China



It is a novel coronavirus



It is a novel coronavirus



FASTA

```
>hCoV-19/Wuhan/IVDC-HB-01/2019|EPI_ISL_402119|
ATTAAAGGTTTATACCTTCCAGGTAACAAACCAACCACTTTCGA
AATCTGTGTGGCTGTCACTC
BetaCoV/Wuhan/IVDC-HB-01/2019
EPI_ISL_402119 via GISAID EpiCoV™
```

2020-01-08

Complete genomes sequenced

Less than 48 hours

2020-01-10

Shared with the world via GISAID



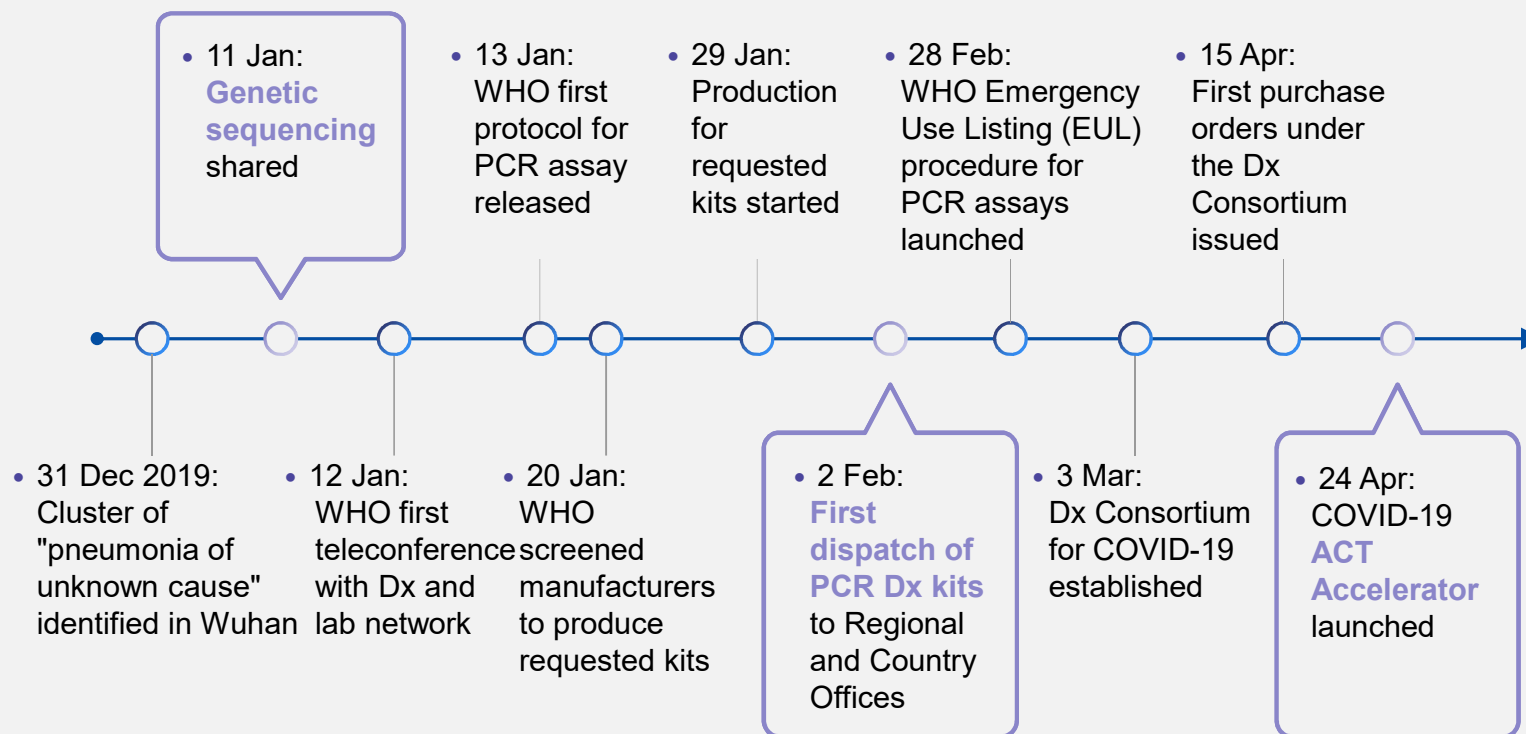
COVID-19 Therapeutics, Diagnostics and Vaccines

ACT Accelerator

- Accelerate the development of diagnostics, therapeutics and vaccines
- Ensure fair and equitable access across the world
- Led by WHO with countries and many global partners
- Covax the vaccine pillar
- Over 200 vaccines in development, 40 in clinical trials
- 60 to 70% of the world's population will need to be immunized
- Public health and social measures needed till 2022

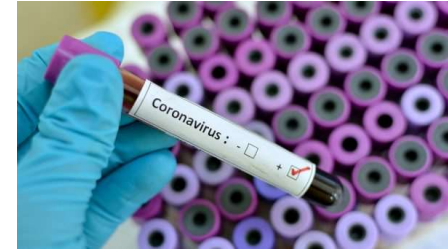


Development of diagnostics for COVID-19 has followed an accelerated timeline

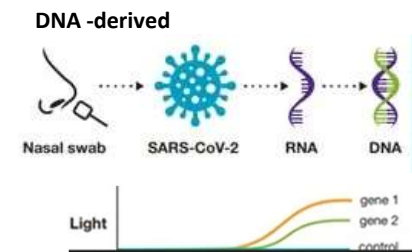


To date: **23 million tests procured** across partners (Global Fund, GDF/StopTB, PAHO, UNDP, Unicef, WHO) with **over 18 million in transit or delivered**

SARS-CoV-2 Diagnostics



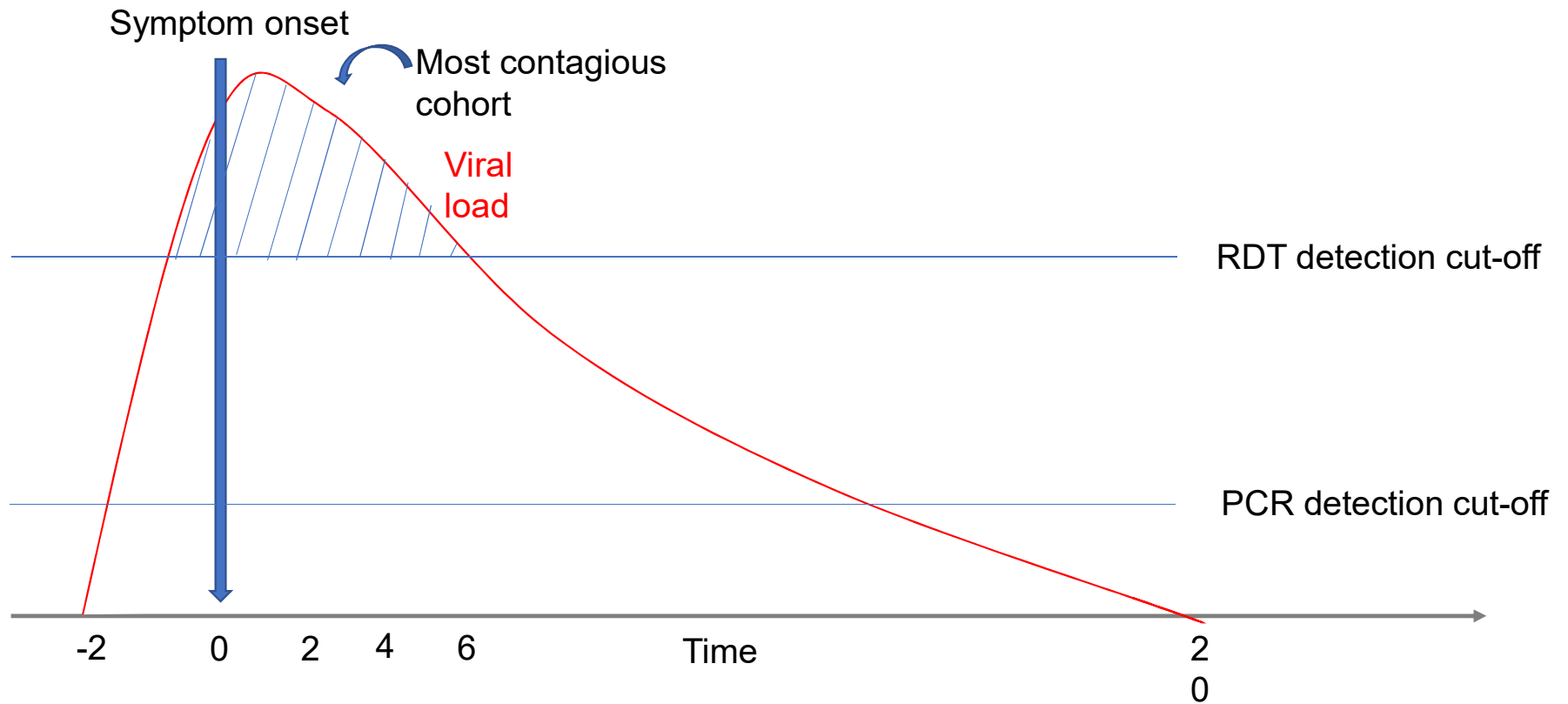
- Nucleic Acid Amplification Test (e.g. rRT-PCR)
 - Reference method for diagnosis of active infection
 - More sensitive than viral culture
 - Limited access globally, especially outside major cities
- Antigen detection (Ag RDT)
 - Simple-to-use lateral flow format, relatively inexpensive, fast
 - Detects most infectious individuals
 - Sensitivity and specificity lower than PCR
- Antibody detection
 - Reliably sensitive only after 2nd to 3rd week of illness
 - Useful in epidemiologic studies, not for acute patient management



Serum, Plasma, fingertip and whole blood



Who can be detected with an Ag RDT?



Repurposed antiviral drugs for COVID-19 Interim WHO SOLIDARITY trial results

**Results submitted to the
New England Journal of Medicine,**

and posted on MedRxiv on October 15, 2020

<https://www.medrxiv.org/content/10.1101/2020.10.15.20209817v1>

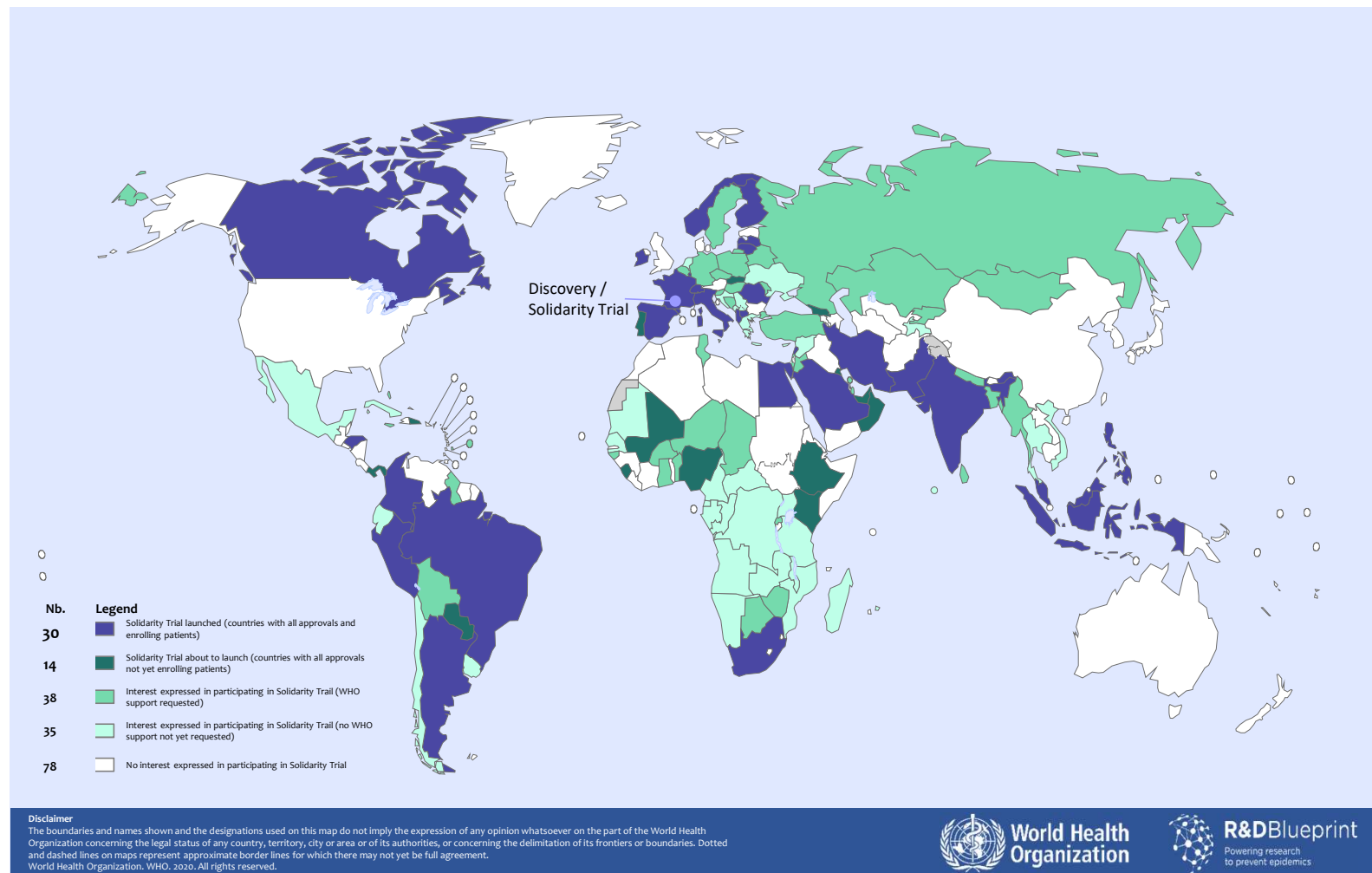
**DISCLAIMER: interim SOLIDARITY trial results does not
necessarily represent the views of the WHO.**

WHO SOLIDARITY randomised trial of 4 drugs for hospitalised COVID

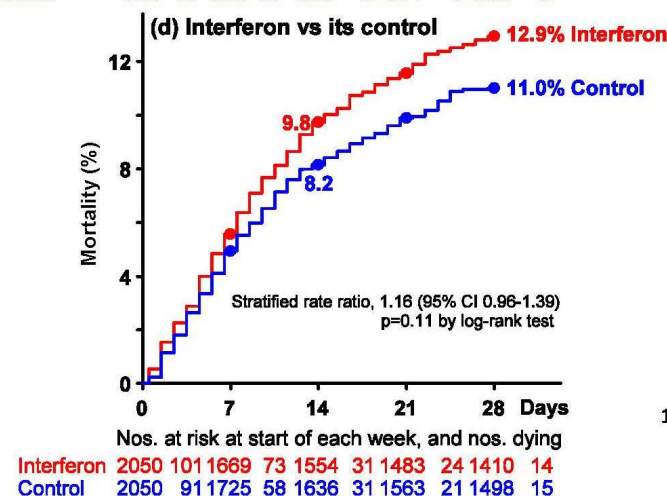
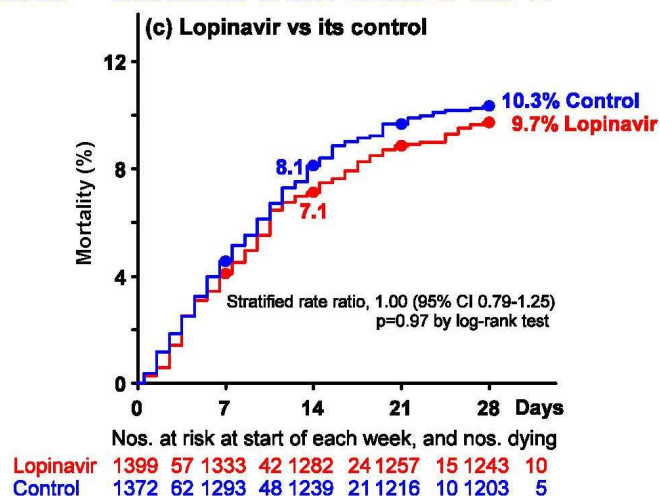
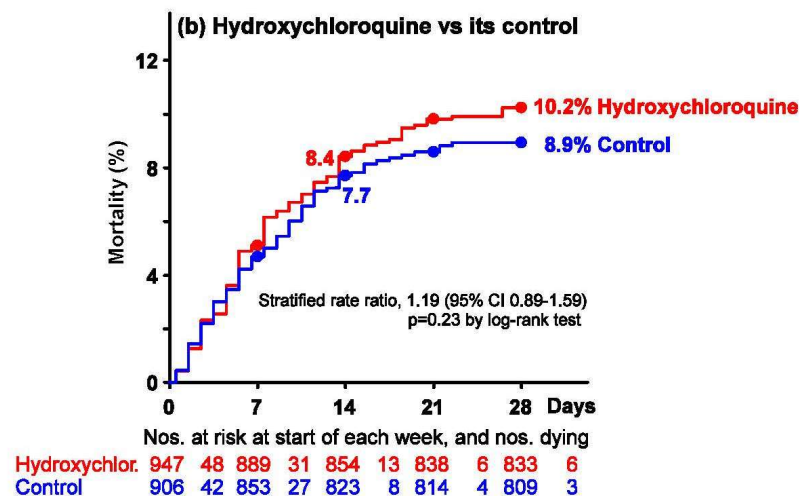
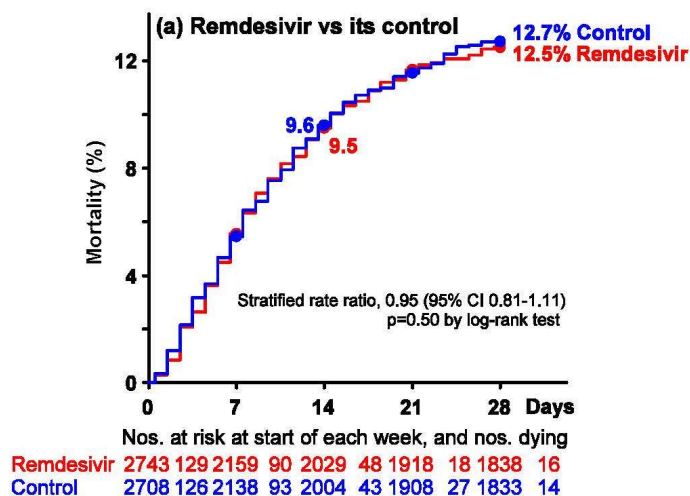
- **Remdesivir (intravenous)**
 - Day 0, 200mg; days 1-9, 100mg.
- **Hydroxychloroquine (oral)**
- **Lopinavir-ritonavir (oral)**
- **Interferon β 1a (mainly subcutaneous)**

WHO Solidarity COVID therapeutics trial

>400 hospitals in **>30** countries (purple: already started) enrolling **>12,000** inpatients



Little effect of the 4 study drugs on 28-day mortality



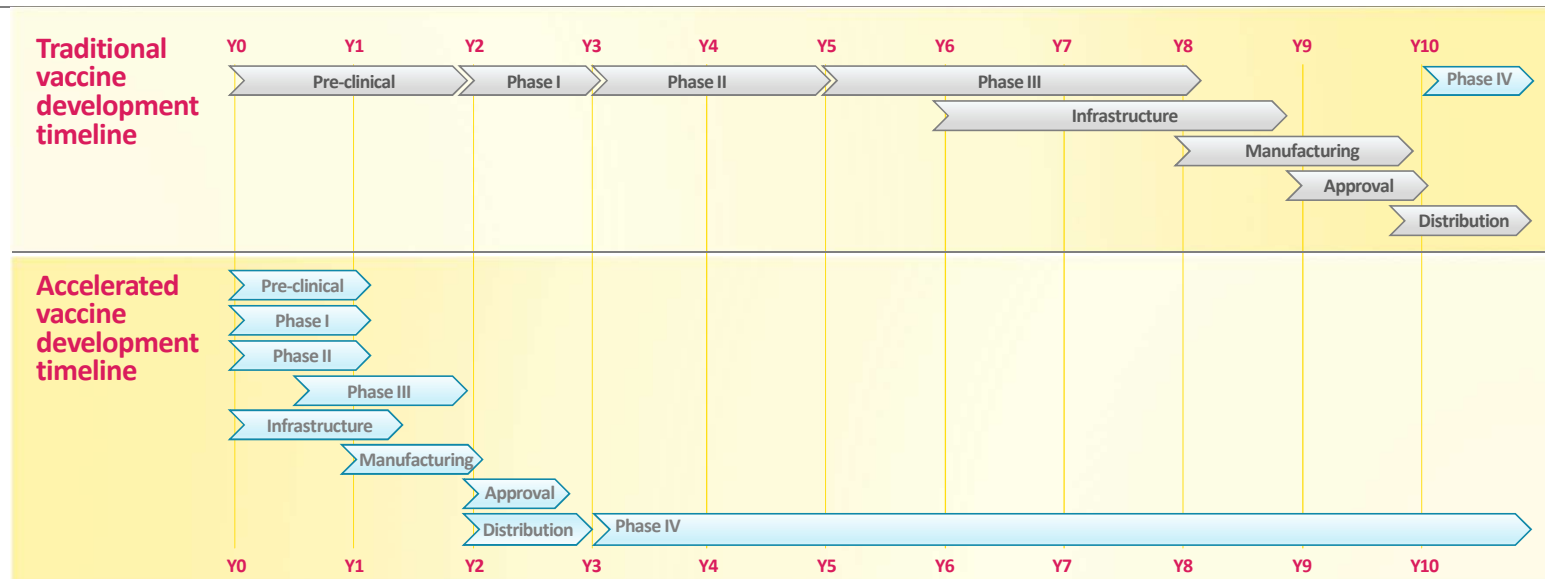
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Solidarity Trials Therapeutics : Conclusions

Combining data from all 4 trials, the Remdesivir vs not death rate ratio (RR) is 0.91 (95% CI 0.79-1.05), $p=0.20$

- This slightly favourable Remdesivir result could have arisen just by chance if Remdesivir does nothing and the real death rate ratio is 1 in all trials.
- Narrower confidence intervals would be helpful (particularly for Remdesivir), but the main need is for better treatments.”
- **There could be a small benefit in lower-risk patients, but not a big one. More evidence is needed, but BETTER DRUGS are needed even more.**

COVID-19 vaccine accelerated development



Normal vaccine development performs each step in sequence
To accelerate COVID-19 vaccine development, **steps are done in parallel**

- All usual safety and efficacy monitoring mechanisms remain in place; such as adverse event surveillance, safety data monitoring & long-term follow-up
- **Phase IV post-marketing surveillance** for side effects is critical and essential

COVID-19 vaccine candidates in Phase III trials

- As of 02 October 2020 there are **42 COVID-19 candidate vaccines** in clinical evaluation of which **10 in Phase III trials**
- There are another **151 candidate vaccines in preclinical** evaluation
- Phase III trials usually require **30,000 or more participants**
- All top candidate vaccines are for **intra-muscular** injection
- Most are designed for a **two-dose** schedule (exceptions with a * in table are single dose)

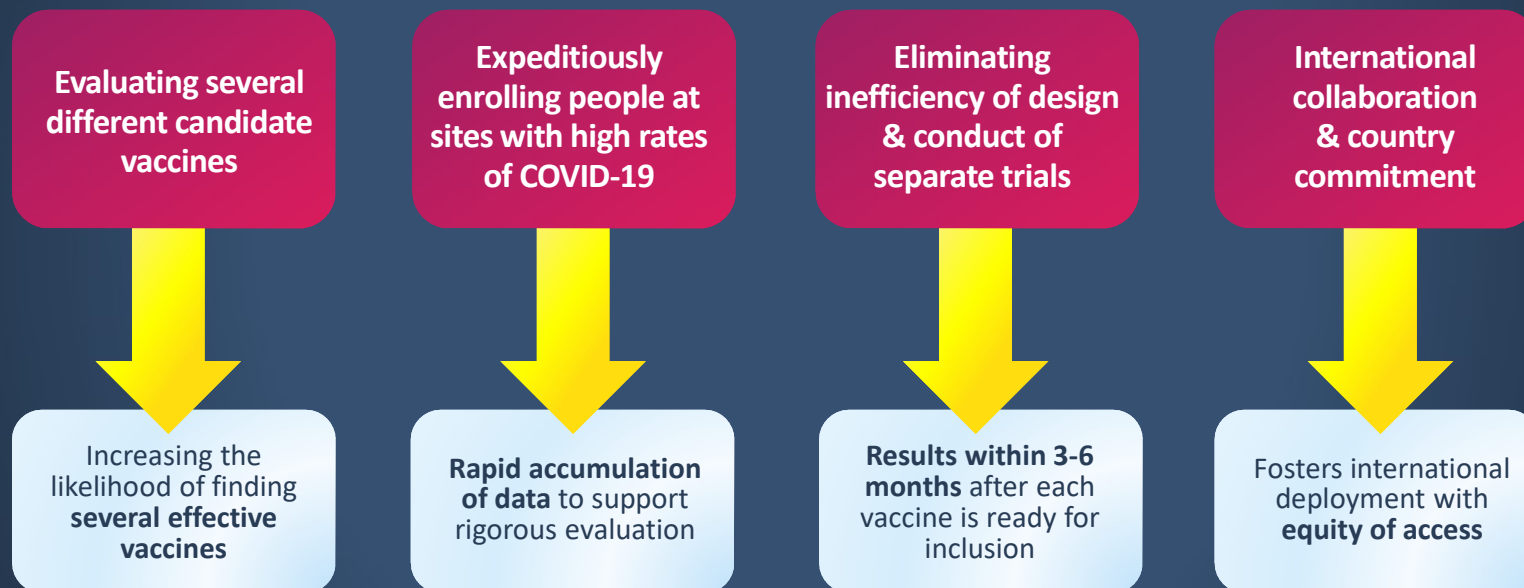
10 CANDIDATE VACCINES IN PHASE III CLINICAL EVALUATION	VACCINE PLATFORM	LOCATION OF PHASE III STUDIES
Sinovac	Inactivated virus	Brazil
Wuhan Institute of Biological Products / Sinopharm	Inactivated virus	United Arab Emirates
Beijing Institute of Biological Products / Sinopharm	Inactivated virus	China
University of Oxford / AstraZeneca	Viral vector *	United States of America
CanSino Biological Inc. / Beijing Institute of Biotechnology	Viral vector *	Pakistan
Gamaleya Research Institute	Viral vector	Russia
Janssen Pharmaceutical Companies	Viral vector	USA, Brazil, Colombia, Peru, Mexico, Philippines, South Africa
Novavax	Protein subunit	The United Kingdom
Moderna / NIAID	RNA	USA
BioNTech / Fosun Pharma / Pfizer	RNA	USA, Argentina, Brazil

<https://www.who.int/publications/m/item/draft-landscape-of-covid-19-candidate-vaccines>

* Single dose schedule

WHO Solidarity vaccine trial

Achieving rapid progress towards global objectives



Key ethical considerations during COVID-19 pandemic

- Build on preexisting guidance to develop guidance materials fit for purpose in COVID-19
- Identify principles for research and their application (SOPs, Solidarity trials, human challenge studies)
- Develop allocation principles for clinical management, vaccines, diagnostics, & therapeutics
- Ethics of AI based applications, contact tracing Apps, immunity passports etc

Ethical Considerations to Guide the Use of Digital Proximity Tracking Technologies for COVID-19 Contact Tracing

Resource Allocation and Priority Setting Policy Brief



Global burden of diseases and injuries, 1990-2009 Findings

- Sustained improvements in health over the past 70 years
- Maternal and child mortality have decreased, as has the burden of TB, HIV, and malaria; All of these were part of the Millennium Development Goals
- But across the world there is rising exposure to crucial risk factors related to the noncommunicable diseases such as diabetes, heart and lung diseases and cancer, including such risks as high blood glucose and particulate air pollution.

Ref: GBD 2019 ([https://www.thelancet.com/journals/lancet/article/PIIS0140-6736\(20\)30925-9/fulltext](https://www.thelancet.com/journals/lancet/article/PIIS0140-6736(20)30925-9/fulltext))

Identified Gaps

- Health systems are ill-prepared for the projected rapid rise in NCDs and disabilities
- Public health has failed to stop the increase in critical risk factors
 - There is a pressing need to tackle wider determinants of health
 - The rise in exposure to key risk factors and deaths from cardiovascular disease indicates a turning point in global life expectancy gains
- The pandemic in context raises difficult questions about the direction of global health:
 - We must pay more attention to the socio-demographic index—which is related to income, education, and fertility rates.

It is not about a single pandemic

As Richard Horton, the editor of *The Lancet*, has written: “The pandemic is not the making of a single coronavirus, but the combination of three epidemics: the virus, the chronic conditions that make people more susceptible to it, and a situation of deepening poverty and inequality. A single pandemic is too simple a narrative to capture this reality. What we’re faced with in Britain,’ he says, is a ‘syndemic’— a synthesis of epidemics.”

The pandemic has shown us that we will only achieve health for all when we address all socioeconomic factors and widespread inequalities.

How are health and healthcare going to be transformed by the current crisis?

- Every government sector now has to be involved in outbreaks
- Early, widespread, accurate communication of information, including WHAT IS NOT KNOWN, is absolutely critical
- Communities must be engaged from the very beginning—bottom up, not top down
- We have to communicate early and often
- We must engage the entire range of experts working on outbreaks, including from the social and behavioural sciences, from the very beginning
- We must prepare now for the next pandemic!



Thank you!

