Monitoring the COVID-19 Pandemic in sub-Saharan Africa: paying attention to health facility admissions and deaths Wednesday 10th June 2020

Chairpersons: Dr Riitta Dlodlo and Dr Jeremiah Chakaya

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Presentation

- 1. **COVID-19:Origins, Global Pandemic, Diagnosis and Treatment.** Leonardo Martinez, Stanford University, School of Medicine
- 2. **Monitoring COVID-19 in health facilities in Africa**Anthony D Harries, The Union, Paris, France, London School of Hygiene & Tropical Medicine, UK

Questions and Answers will follow the presentation.

COVID-19: Origins, Global Pandemic, Diagnosis, and Treatment

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Origin of the COVID-19 pandemic

- First cases likely developed in late November to early December 2019 in Wuhan, China
- First reported case was reported in early January 2020
- Linked epidemiologically to a seafood and wet animal market

BRIEF REPORT

A Novel Coronavirus from Patients with Pneumonia in China, 2019

Na Zhu, Ph.D., Dingyu Zhang, M.D., Wenling Wang, Ph.D., Xingwang Li, M.D., Bo Yang, M.S., Jingdong Song, Ph.D., Xiang Zhao, Ph.D., Baoying Huang, Ph.D., Weifeng Shi, Ph.D., Roujian Lu, M.D., Peihua Niu, Ph.D., Faxian Zhan, Ph.D., Xuejun Ma, Ph.D., Dayan Wang, Ph.D., Wenbo Xu, M.D., Guizhen Wu, M.D., George F. Gao, D.Phil., and Wenjie Tan, M.D., Ph.D., for the China Novel Coronavirus Investigating and Research Team



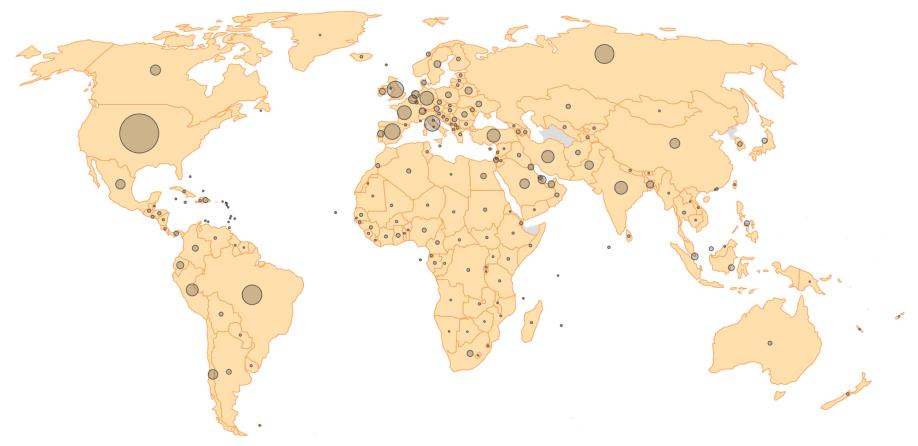
Origin of the COVID-19 pandemic

• By January 7, 2020, scientists isolated the coronavirus

• Full genome sequence data enabled the rapid development of RT-PCR diagnostic tests specific for this novel coronavirus

Global Spread

• By the end of January, cases were confirmed in >25 countries



• Currently, >7 million cases reported on all 6 continents

World Health Organization

• Declared a Public Health Emergency of International Concern on January 30, 2020

• Declared a global pandemic on March 11, 2020

Incubation Period of COVID-19

• Incubation: time from infection to disease onset

• ~ 4-5 days; however, different estimates (range is between 2 and 14 days)

• Among patients, 99% develop symptoms within 14 days

Spectrum of COVID-19 disease

 Asymptomatic and paucisymptomatic disease is common

Mild disease

Severe and critical disease

COVID-19 Diagnosis

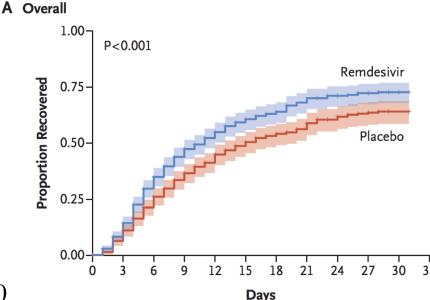
- Nasopharyngeal, oropharyngeal, and nasal swabs are the preferred specimen type
- Some other sampling types (sputum, BAL) are also acceptable under certain circumstances
- Cepheid Xpert platform
- Antibody testing

- Many drugs for treatment being tested
 - Remdesivir
 - Hydroxychloroquine/chloroquine
 - Lopinavir/ritonavir

• Recent positive randomized trial result for Remdesivir (200 mg day 1, 100 mg daily for up to 9 additional days) in hospitalized COVID-19 patients

• Reduced time to recovery (median, 11 vs 15 days)

Reduced mortality
(HR, 0.70; 95% CI, 0.47–1.04)





Hydroxychloroquine

• Observational study (N=1,446) suggests similar mortality risk in hospitalized patients

Trials are needed

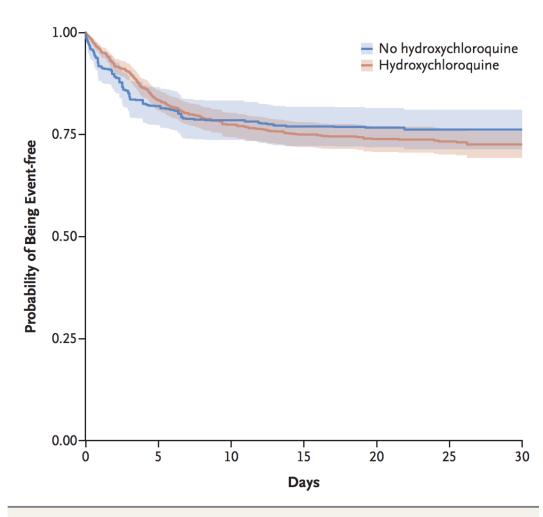


Figure 2. Freedom from Composite End Point of Intubation or Death.

The shaded areas represent pointwise 95% confidence intervals.

- Lopinavir/ritonavir
- Randomized trials

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

MAY 7, 2020

VOL. 382 NO. 19

A Trial of Lopinavir–Ritonavir in Adults Hospitalized with Severe Covid-19

CONCLUSIONS

In hospitalized adult patients with severe Covid-19, no benefit was observed with lopinavir—ritonavir treatment beyond standard care. Future trials in patients with severe illness may help to confirm or exclude the possibility of a treatment benefit.

- No benefit in time to clinical improvement
- Lower mortality (19.2% vs. 25.0%) but low sample size precludes meaningful conclusions

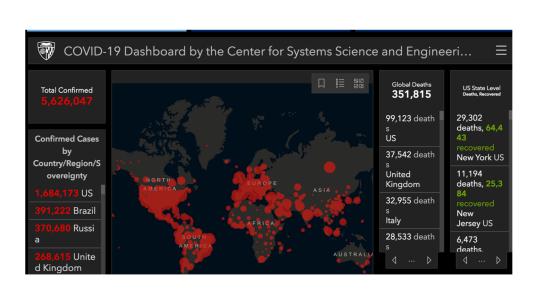
Other therapies under evaluation

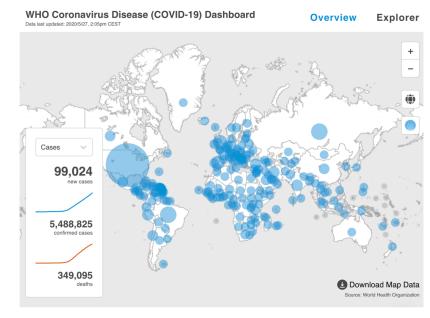
- BCG vaccination
 - BCG has non-specific effects on several diseases other than tuberculosis possibly through 'trained immunity'
 - Several ongoing clinical trials in Australia, Germany, and the Netherlands

Plasma therapy

• Heparin, other antiviral drugs and antiinflammatory drugs

Limitations of Global Case and Mortality Reporting





• Includes only reported diagnosed COVID-19 patients

• Underestimate of true burden; many undiagnosed cases are asymptomatic or paucisymptomatic or never reported

Monitoring COVID-19 in health facilities in Africa

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Case definition of COVID-19

- New onset dry cough
- AND New onset fever
- AND new onset shortness of breath

• Maybe supplement this with alterations in smell and taste [G Spinato et al, JAMA 2020]

Health facility monitoring tool

| Date of Admission | COVID ID no. | Suspected Confirmed | Age | Sex | HIV status: | ART | TB Treatment | Previous TB | HTN DM CVD | Smoker | Given O ₂ | Given Medication ^a | HDU ICU | Mechanical Ventilation | Date of Discharge | DIED | Date of Death |
|----------------------|-----------------|------------------------|-----|-----|----------------|------|-----------------|----------------|------------------|--------|-------------------------|----------------------------------|------------|---------------------------|----------------------|------|------------------|
| 10 June | 1 | S | 50 | M | Neg | NA | No | Yes | No | Yes | Yes | No | No | No | 20 June | No | N/A |
| 10 June | 2 | S | 45 | F | Neg | NA | Yes | No | No | No | No | No | No | No | 15 June | No | N/A |
| 11 June | 3 | С | 65 | M | Neg | NA | No | No | DM | Yes | Yes | Yes | Yes | No | N/A | Yes | 19 Jun |
| 11 June | 4 | С | 62 | M | Neg | NA | No | No | HTN | No | Yes | Yes | Yes | No | 22 June | No | N/A |
| 11 June | 5 | С | 53 | M | Pos | Yes | Yes | No | No | No | No | No | No | No | 18 June | No | N/A |
| 11 June | 6 | S | 25 | M | Neg | NA | No | Yes | No | No | No | No | No | No | 19 June | No | N/A |
| 12 June | 7 | S | 32 | F | Neg | NA | No | No | HTN | No | No | No | No | No | 18 June | No | N/A |
| 12 June | 8 | S | 72 | M | Neg | NA | No | No | No | Yes | Yes | Yes | Yes | No | N/A | Yes | 16 Jun |
| 12 June | 9 | С | 68 | F | Neg | NA | No | Yes | No | Yes | Yes | Yes | Yes | No | N/A | Yes | 22 Jun |
| 13 June | 10 | С | 61 | M | Pos | Y es | No | No | No | No | Yes | No | No | No | 23 June | No | N/A |

Recording Dates

Dates of:

- Hospital admission
- Onset of illness
- Confirmed diagnosis
- Clinical recovery
- Hospital discharge
- Death

Provides information on:

- Duration of illness
- Bed occupancy
- Time to death

Demographics, pregnancy, source of infection

Gender

- Age
- Pregnancy

Source

- Males more affected than females
- Case fatality in China (JAMA 2020):

• Case fatality in New York (JAMA 2020):

Risk factors and co-morbidities

- Hypertension
- Cardiovascular disease
- Diabetes mellitus
- Chronic lung disease
- Asthma
- Cancer
- Other (renal)
- Smoking

• Case fatality China (JAMA 2020):

10% cardiovascular disease; 7% diabetes mellitus; 6% chronic lung disease; 6% hypertension; 6% cancer

• Risk of death in UK (17,000 pts)

increased risk with cardiovascular disease (37%); lung disease (17%); kidney disease (25%)

Co-morbidities and death

Data from one large Italian Hospital:

| • | No comorbidity | 1% of the | deaths |
|---|----------------|-----------|--------|
|---|----------------|-----------|--------|

- 1 comorbidity 26% of the deaths
- 2 comorbidities 26% of the deaths
- 3+ comorbidities 47% of the deaths

Lorenzo G et al, JAMA 2020

Tuberculosis and HIV/AIDS

- BCG at birth
- Previous history of TB
- Currently with TB
- HIV-positive
- On ART
- First-line / second-line
- CPT / IPT

Comments:

- BCG protective? Clinical trials underway
- Previous TB chronic respiratory disease and cardiac disease
- Active TB increased risk of COVID-19 (China)
- PLHIV immune-suppressed
- ARV drugs LPV/r said to possibly work but clinical trial in China showed no benefit (NEJM 2020)

Presenting symptoms

- Fever
- Night sweats
- Cough dry, blood-stained
- Chest pain
- Shortness of breath
- Headache
- Other "COVID Toes"



Alterations in smell and taste

130 patients interviewed:

- 64% with altered sense
- More common in women
- Median score 4 (max = 5)
- 1/3 had blocked nose
- ³/₄ at same time/after first symptoms

- virus invades CNS
 through olfactory
 system and replicates
 in olfactory bulb
- nasal epithelial cells have highest expression of ACE2 receptors



Physical examination

• One of the key things is to measure height and weight and obtain BMI (Weight/height²)

• BMI of 25-30 = overweight BMI > 30 = obese

• UK: Obesity associated with 37% increase risk of death

Medical interventions in hospital

Oxygen +/- CPAP

Cough

Day 0

LPV/r; remdesivir

Pneumonia

Day 7

Chloroquine (hydroxy-CQ)

Respiratory failure

Day 14

Steroids – anti-inflammatories

Antibiotics

Cytokine storm
Clotting disorder
Multi-organ failure

Day 21

Heparin/aspirin

Death

Further specialised services

High dependency unit

Intensive care unit

Mechanical ventilation

• ICU services (dialysis)

New York City
Mechanical ventilation
mortality:
18-65 years = 76%

 $>65 \ years = 97\%$

Progress

Clinically recovered

Hospital discharge

[readmission]

Absconded

Hospital death

1.Daily reporting: cross-sectional analysis

| Date | Number of new cases (suspected and confirmed) | Number of new deaths | Number of cumulative cases (suspected and confirmed) | Number of cumulative deaths |
|---------|---|----------------------|--|-----------------------------|
| 10 June | 4 | 1 | 4 | 1 |
| 11 June | 6 | 2 | 10 | 3 |
| 12 June | 10 | 2 | 20 | 5 |
| 13 June | 6 | 2 | 26 | 7 |
| 14 June | 4 | 0 | 30 | 7 |

2. Monthly reporting: cohort analysis

| Year | Monthly cohort | Number admitted with COVID | Number discharged /absconded | Number died | Number still in hospital | Date of report |
|------|----------------|----------------------------|------------------------------|----------------|--------------------------|----------------|
| 2020 | January | 50 | 40 | 10 (20%) | 0 | 1 March |
| 2020 | February | 60 | 48 | 10 (17%) | 2 | 1 April |
| 2020 | March | 80 | 65 | 13 (16%) | 2 | 1 May |
| 2020 | April | 80 | 60 | 15 (19%) | 5 | 1 June |
| 2020 | May | 100 | 60 | 25 (25%) | 15 | 1 July |

3.Six-month reporting on COVID: Risk factors for death in COVID admissions

| Category | Variable | Total COVID n | Died n (%) | RR (95% CI) |
|--------------|-------------|------------------|---------------|----------------|
| Gender | Female | | | reference |
| | Male | | | > 1.0 |
| Age in years | <60 | | | reference |
| | ≥60 | | | >1.0 |
| Comorbidity | Normal BMI | | | reference |
| | Obese BMI | | | >1.0 |
| | No diabetes | | | reference |
| | Diabetes | | | >1.0 |
| Smoking | No | | | reference |
| | Yes | | | ?? |
| Ventilation | No | | | reference |
| | Yes | | | >1.0 |

Conclusion

• Standardised and regular health facility monitoring and reporting

• Sentinel surveillance in hot spots / high-risk groups:

care homes for the elderly congregate settings – e.g., prisons health care workers

Poll Question

Questions?

The Union



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THANK YOU

Register now for next week's webinars: Thursday 18 June, 14:00 - 15:00 CEST

COVID-19 and TB: Personal perspectives

This open discussion will focus in particular on the mental health challenges of dealing with TB and COVID-19 and issues around the stigma which so many survivors of both diseases face. Other topics will include how the TB response has

been affected by COVID-19 and community responses to this.

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