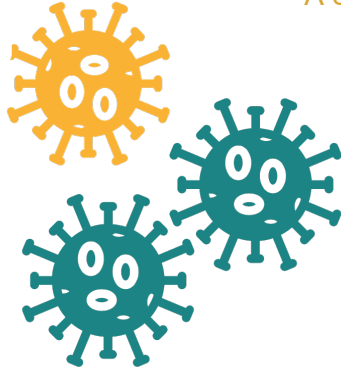


Modelling COVID-19 epidemics

A course organized by ISBE-NL, ELIXIR-LU, and EOSC-Life



Biology & Epidemiology

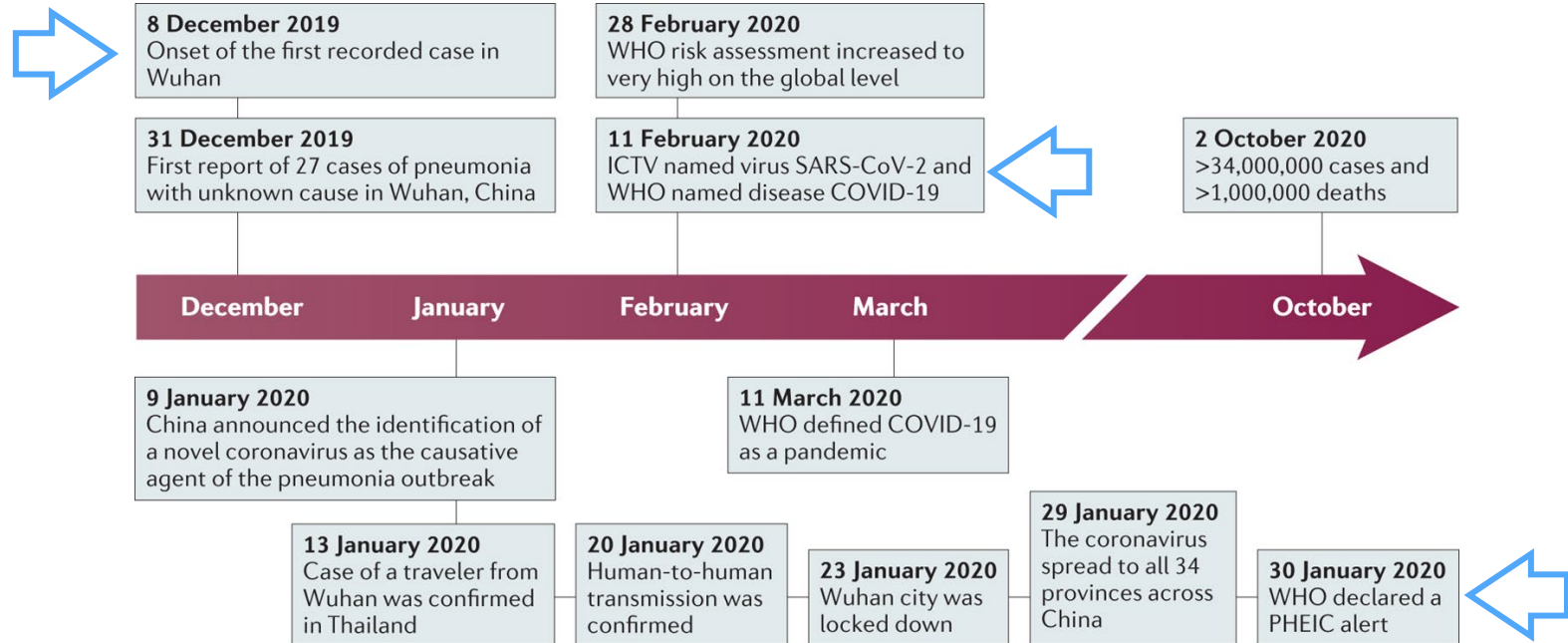
Module I

November 30- December 2, 2020

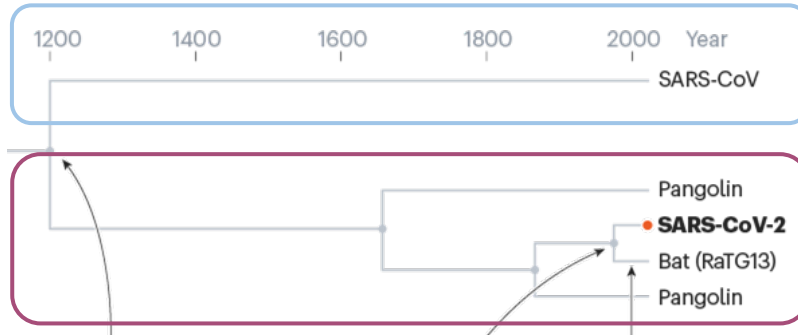
Stefan Astorolo & Hans WESTERHOFF



Timeline of COVID-19 outbreak



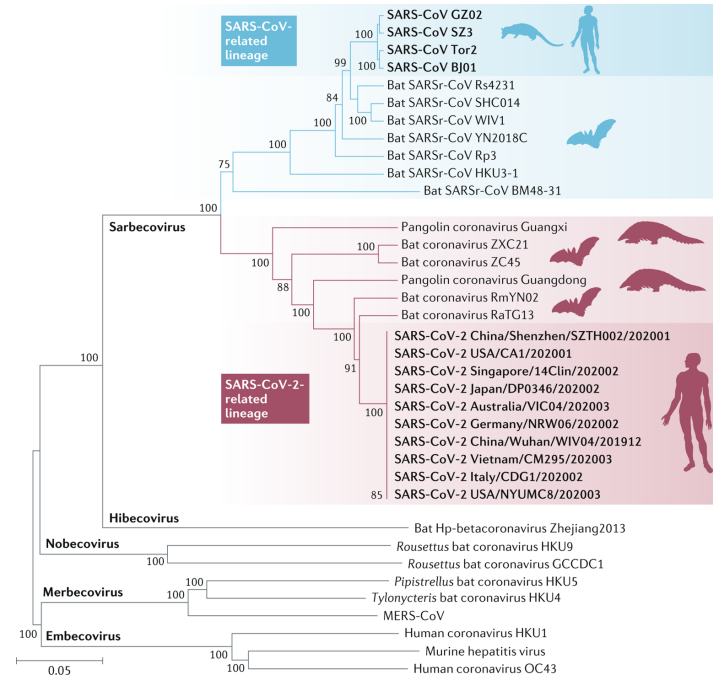
January 31, 2020 first sequencing SARS CoV2



The SARS-CoV-2 lineage might have separated centuries ago from that of SARS-CoV, the cause of SARS.

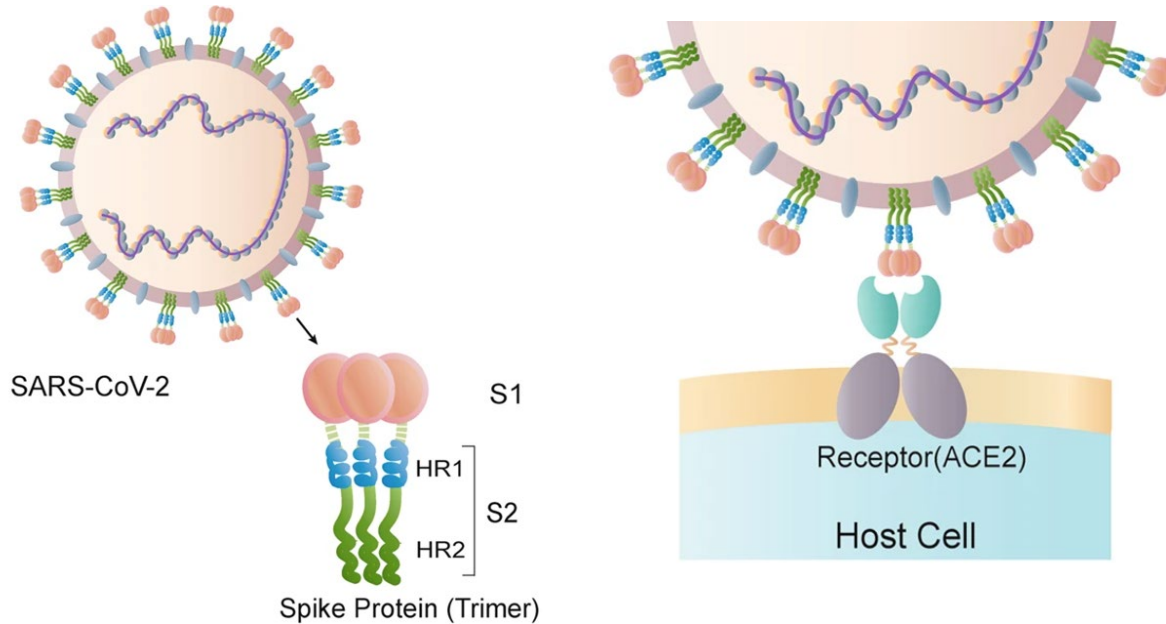
A coronavirus found in a bat (RaTG13) is the closest known relative of SARS-CoV-2. They might have split 40–70 years ago.

The lineage leading to RaTG13 apparently lost the specialized binding domain that helps SARS-CoV-2 infect human cells.



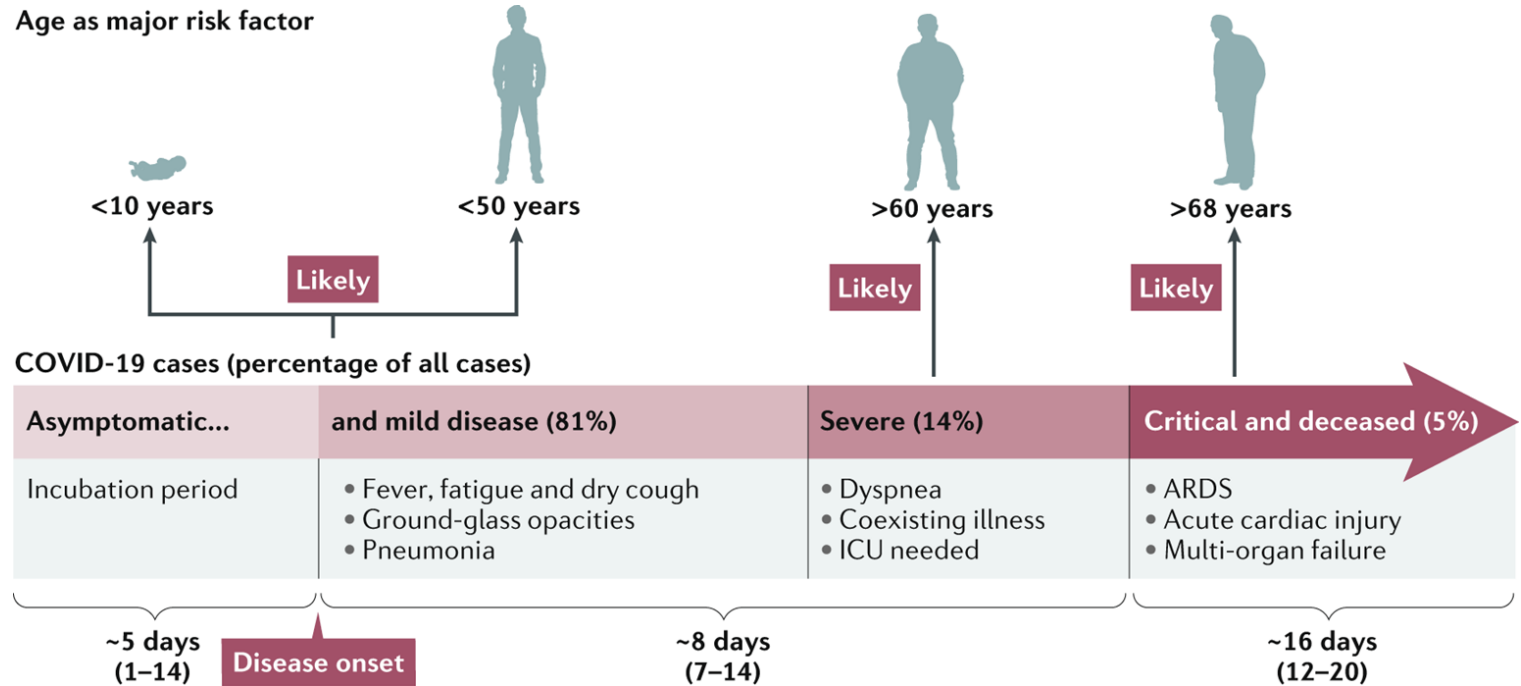


Features of SARS CoV2 (virus structure)



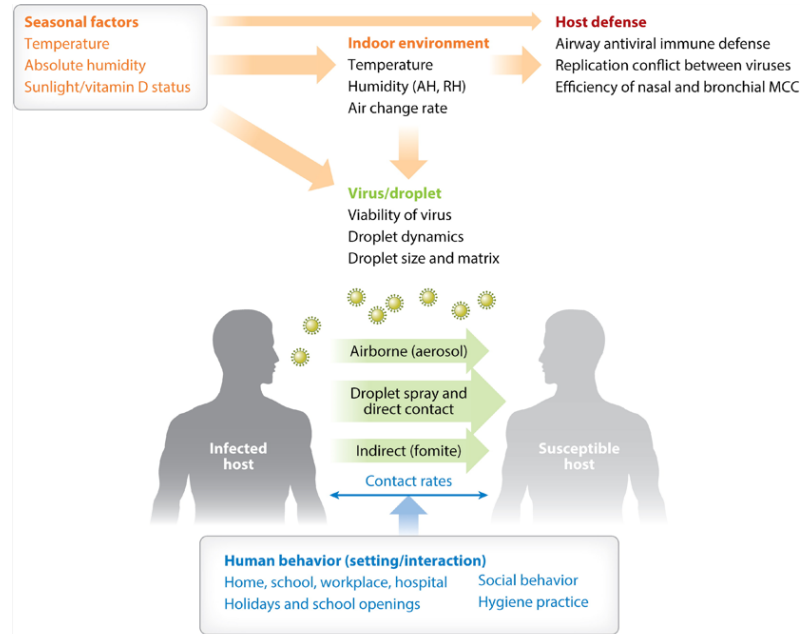
Clinical features of COVID - 19

Age as major risk factor





Transmission routes



Moriyama M, et al, 2020.
Annu. Rev. Virol. 7:83–101



Obvious measures to prevent spreading

Personal Protective measures:



→ Hands hygiene



→ Cough and Sneeze into
your elbow or a tissue.



→ Wear masks and PPE

Environmental measures:



→ Frequently clean used
surface, clothes and
objects



→ Minimize sharing objects



→ Ensure appropriate
ventilation



Social distancing

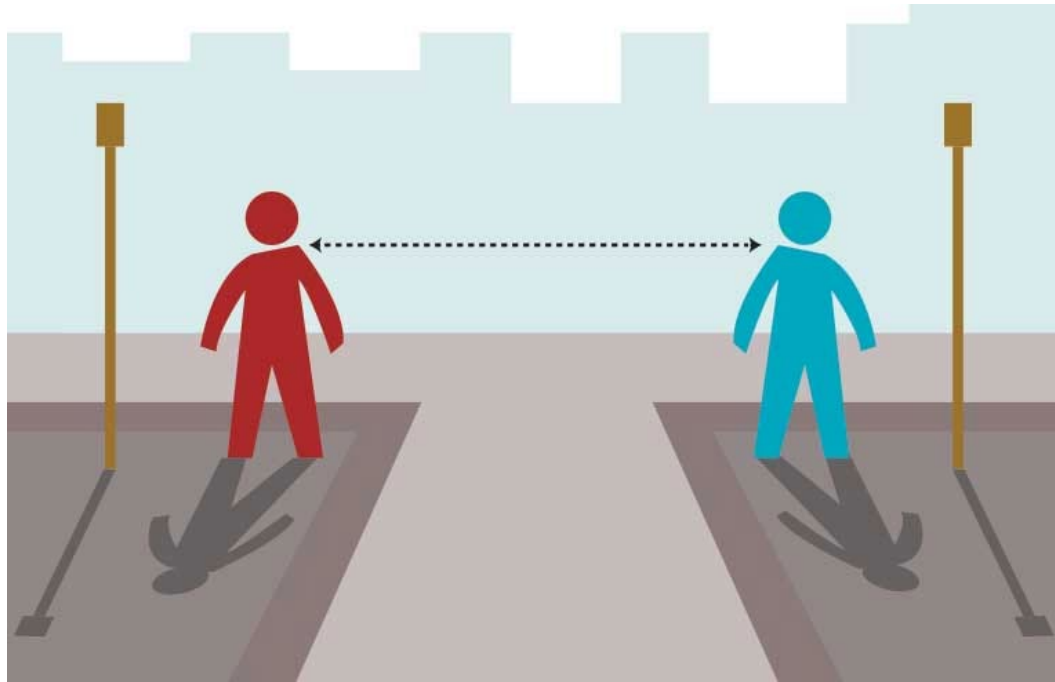


Figure from [safetyandhealthmagazine.com](https://www.safetyandhealthmagazine.com)



COVID-19 testing basics

There are two different types of tests – **diagnostic tests** and **antibody tests**.

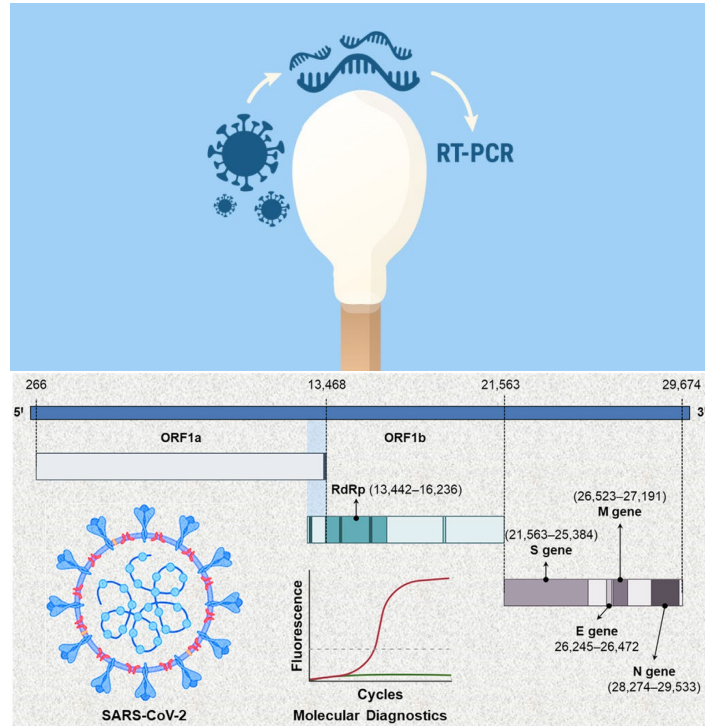
	Virus RNA	Host immunity	Virus protein
How the sample is taken...	Nasal or throat swab (most tests), Saliva (a few tests)	Nasal or throat swab	Finger stick or blood draw
How long it takes to get results...	Same day (some locations) or up to a week	One hour or less	Same day (many locations) or 1-3 days
Is another test needed...	This test is typically highly accurate and usually does not need to be repeated.	Positive results are usually highly accurate but negative results may need to be confirmed with a molecular test.	Sometimes a second antibody test is needed for accurate results.
What it shows...	Diagnoses active coronavirus infection	Diagnoses active coronavirus infection	Shows if you've been infected by coronavirus in the past
What it can't do...	Show if you ever had COVID-19 or were infected with the coronavirus in the past	Definitively rule out active coronavirus infection. Antigen tests are more likely to miss an active coronavirus infection compared to molecular tests. Your health care provider may order a molecular test if your antigen test shows a negative result but you have symptoms of COVID-19.	Diagnose active coronavirus infection at the time of the test or show that you do not have COVID-19



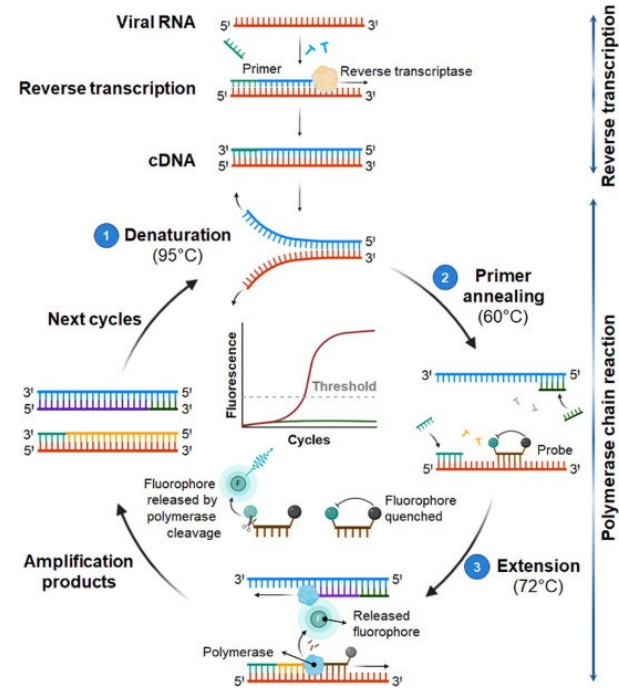
“gargle” testing in Vienna



Molecular test (RT-PCR)



Afzal 2020, *Journal of Advanced Research*





- there are ~ 150 kits ... available
- The majority close to 100% specificity
- Still there are false negatives ...

Timing in testing

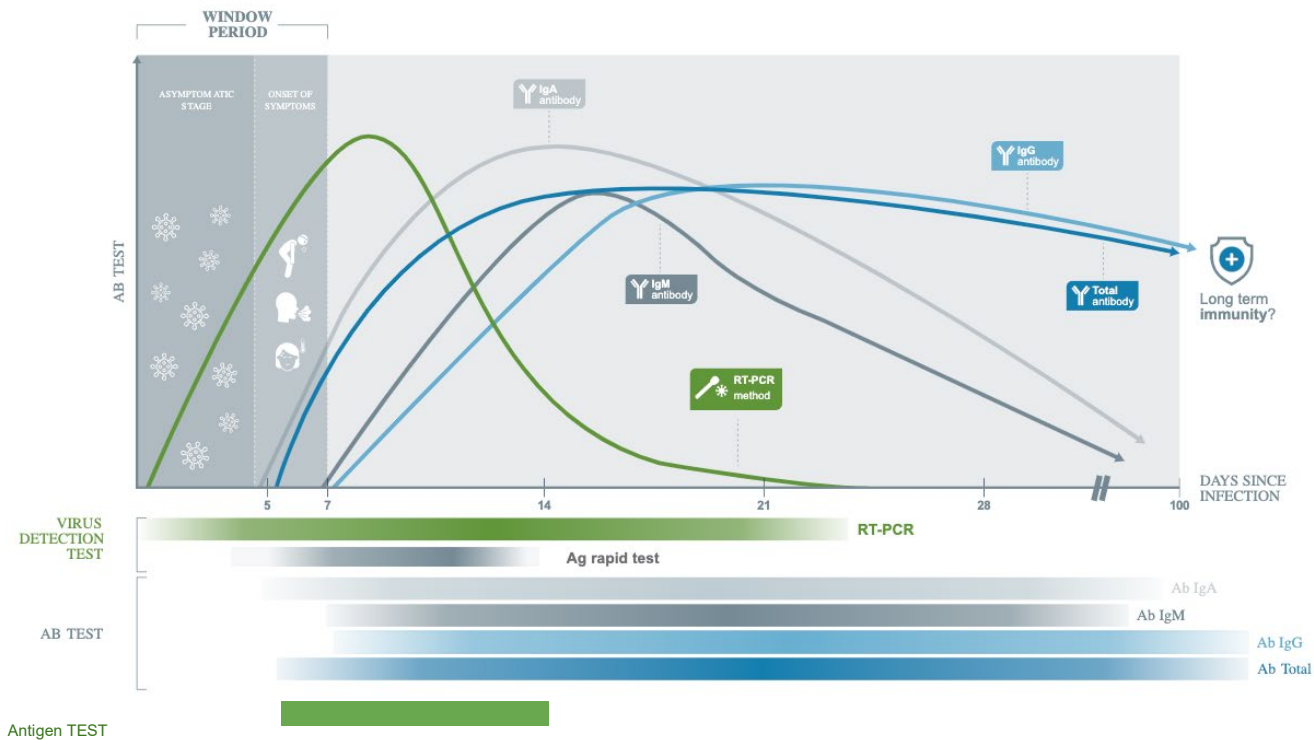
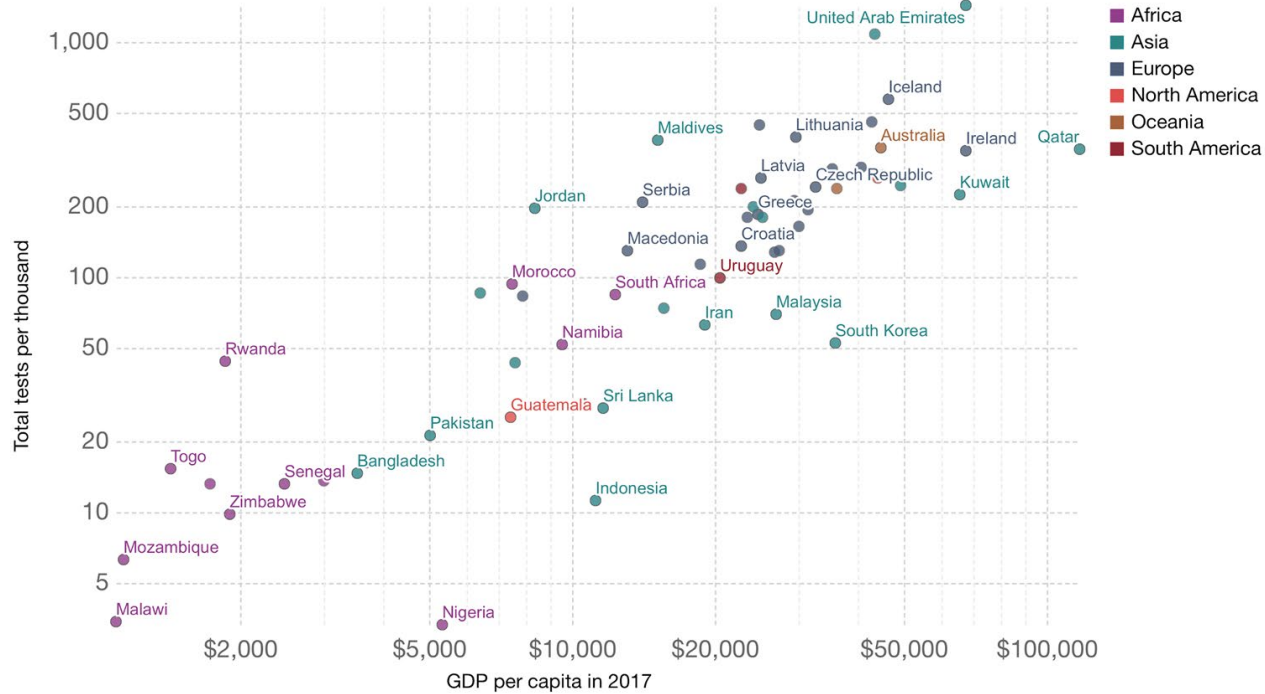


Figure from <https://www.synlab.com/news-publications/sars-cov-2/antigen-tests-for-sars-cov-2-detection>

Why it is so important to develop *cheapest* tests?





R_0 : *reproduction number*



Figure from <https://www.bbc.com/news/health-52473523>



R_0 : *reproduction number*



What is R_0 : reproduction number ?

- First used almost a century ago in demography, R originally measured the reproduction of people — whether a population was growing or not.
- In epidemiology, the same principle applies, but it measures the spread of infection in a population. If R is two, two infected people will, on average, infect four others, who will infect eight others, and so on.



Figure from <https://www.bbc.com/news/health-52473523>



What R_0 *can* and *can't* tell us about managing COVID-19

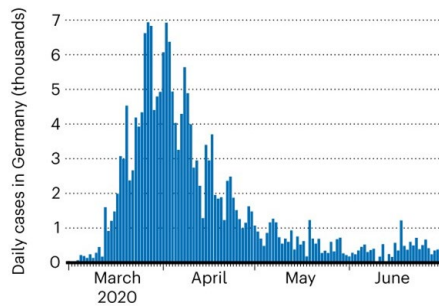
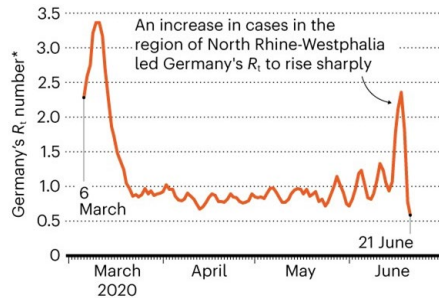


Figure from <https://www.nature.com/articles/d41586-020-02009-w>



What R_0 *can* and *can't* tell us about managing COVID-19

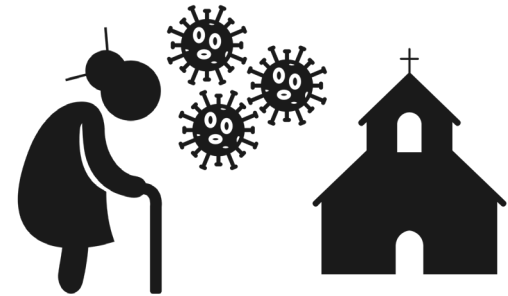
Regional Outbreak



*Latest data 26 June (5-day delay).

©nature

superspreaders



Shincheonji Church of Jesus in Daegu, South Korea



The *fallacy* of herd immunity



(© Laurinson Crusoe/Shutterstock)



The *fallacy* of herd immunity



→ What is herd immunity?

“Herd immunity happens when a virus can’t spread because it keeps encountering people who are protected against infection. “You don’t need everyone in the population to be immune — you just need enough people to be immune,”



(© Laurinson Crusoe/Shutterstock)



The *fallacy* of herd immunity



Figure from <https://www.bbc.com/news/uk-51677846>



A cemetery in Manaus, Brazil, in June.

<https://www.nature.com/articles/d41586-020-02948-4>



The cost of herd immunity



~ 747 million

1-3% deaths



~ 11 million



Our World In Data



Our World Data → Coronavirus Pandemic (COVID-19)

They built 207 country profiles which allow you to explore **the statistics on the coronavirus pandemic for every country in the world** .

Each profile includes **interactive visualizations** , **explanations** of the presented metrics, and the details on the **sources of the data** .

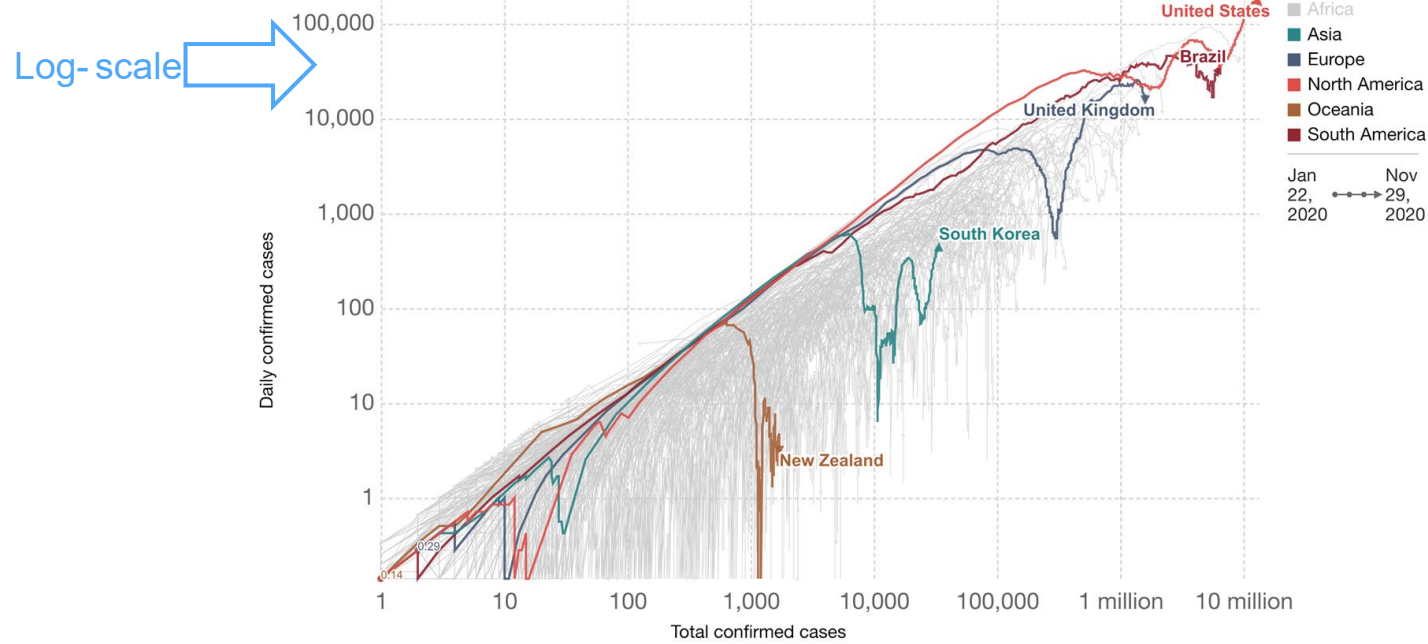
Every country profile is **updated daily** .

Every profile includes four sections:

1. **Deaths**
2. **Testing**
3. **Cases**
4. **Government responses**



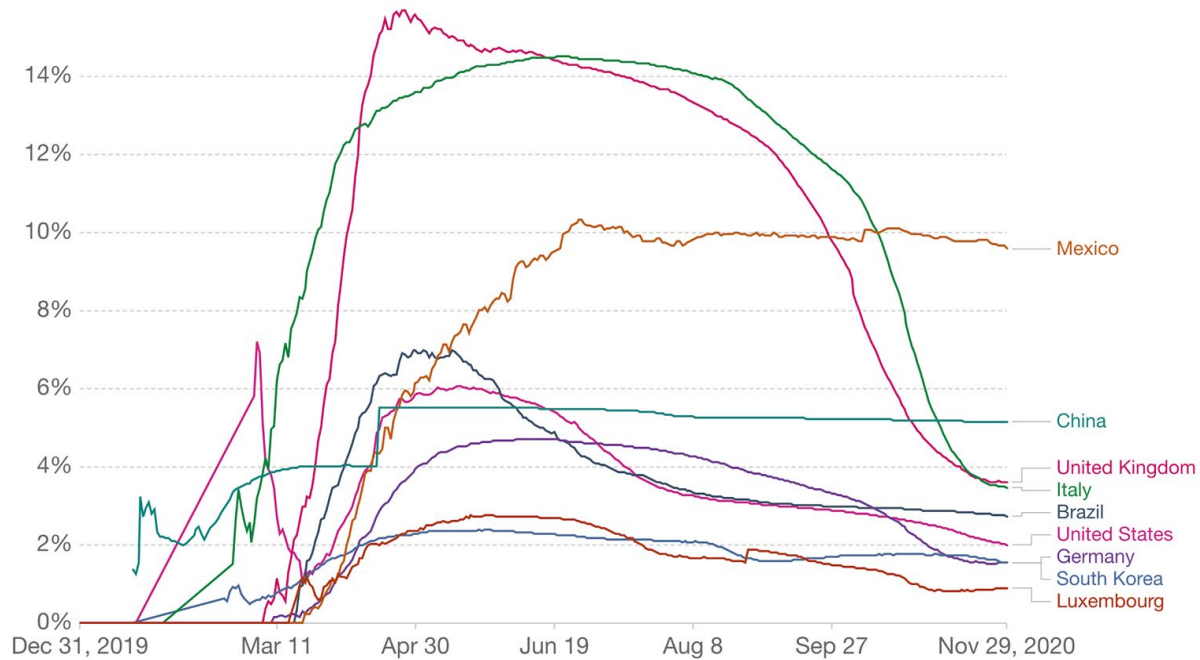
When did countries bend the curve?



Source: European CDC – Situation Update Worldwide – Last updated 29 November, 10:06 (London time), Our World In Data
OurWorldInData.org/coronavirus • CC BY



Case fatality rate



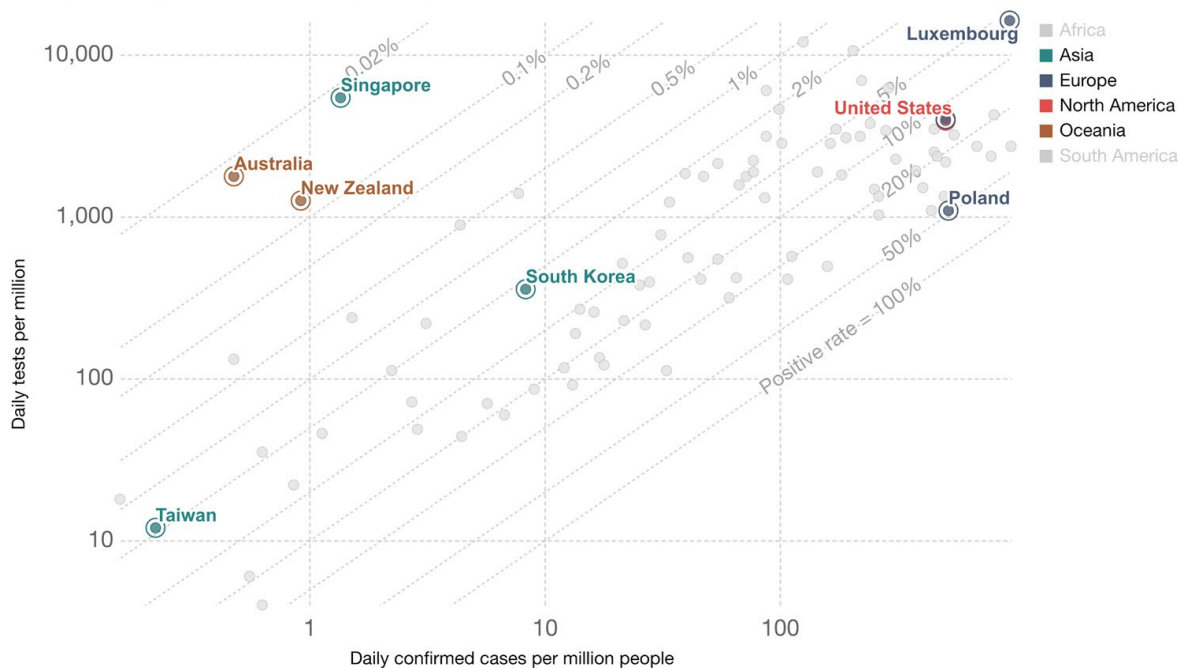
Source: European CDC – Situation Update Worldwide – Last updated 29 November, 10:06 (London time)

CC BY

Positivity rate

COVID-19: Daily tests vs. Daily new confirmed cases per million

The figures are given as a rolling 7-day average.



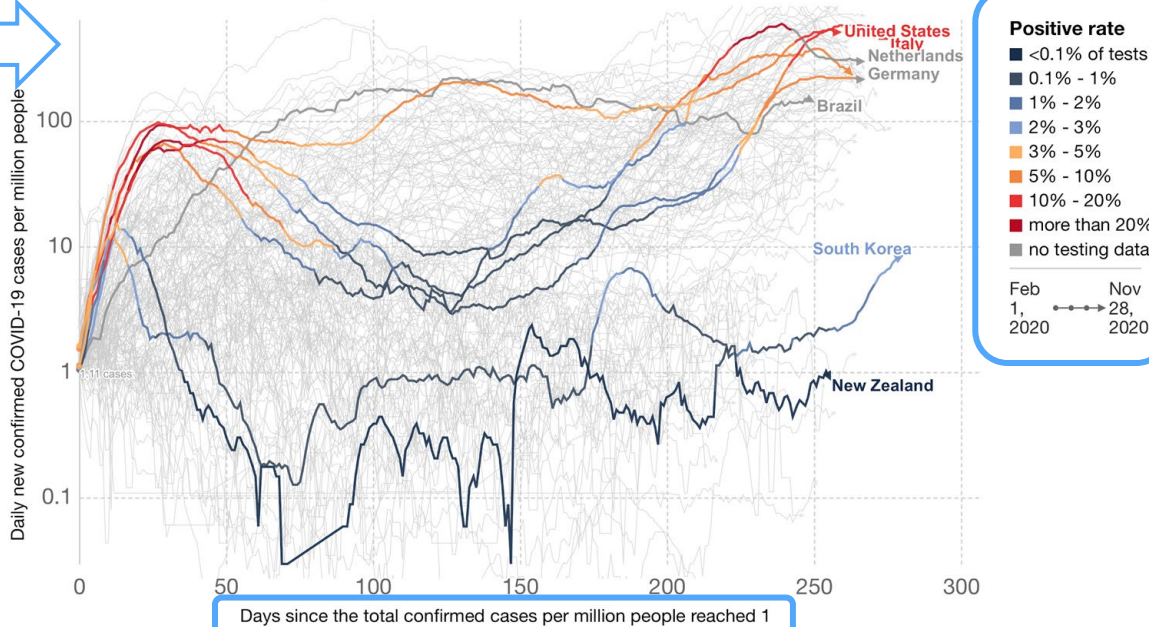
Daily (confirmed) cases

Daily new confirmed COVID  per million people

Shown is the rolling 7-day average. The number of confirmed cases is lower than the number of actual cases; the main reason for that is limited testing.

Our World in Data

Log-scale 



Source: European CDC – Situation Update Worldwide – Last updated 28 November, 10:06 (London time), Official data collated by Our World in Data
 CC BY



Vaccines & Therapeutics





Vaccines & Therapeutics



Therapeutic Drugs



Vaccines

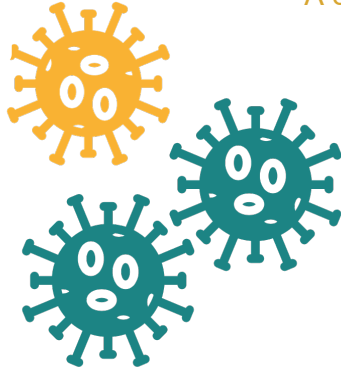


Categories

1. Inactivated virus
2. Live attenuated virus
3. Protein Subunit
4. DNA-Based
5. RNA-Based → [BNT162b2 \(Pfizer and BioNTech\)](#)
6. Replicating Viral Vector
7. Non-Replicating Viral Vector → [Sputnik V](#)
8. Virus-like particle → [AZD1222 \(Astrazeneca\)](#)
9. Other vaccine

Modelling COVID-19 epidemics

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Policies of COVID-19

Module I

November 30- December 2, 2020

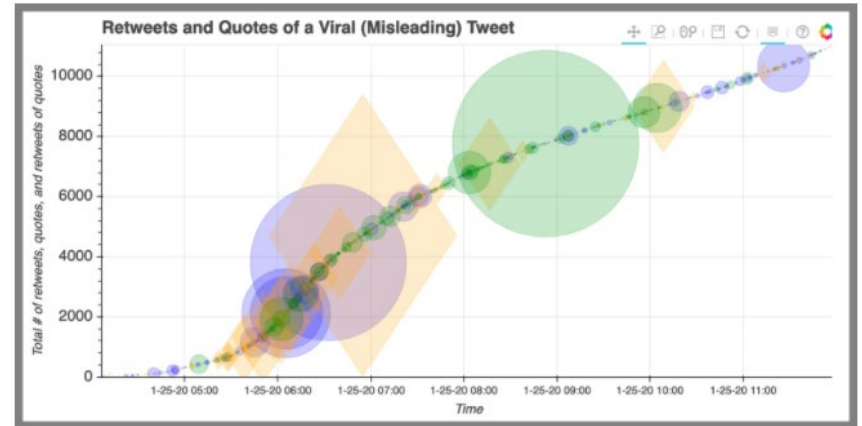
Stefania Astrologo & HAns WESTERHOFF



The scientist' duty

Provide data, facts, scientific knowledge, tests, ... vaccine

All to support *rational* policy making



University of Washington professor Kate Starbird used a database of tweets about Covid-19 to create this chart showing how retweets (blue circles), quotes (orange diamonds), or retweets of quotes (green circles), boosted a tweet sharing inaccurate scientific claims about the novel coronavirus. COURTESY OF KATE STARBIRD



Just a cold!



Brazil's President Jair Bolsonaro



The scientist' task

Provide data, facts, scientific knowledge, tests, ... vaccine

All to support *rational* policy making



The world is seeing many *crises* ...



Hunger
Poverty
Cancer
Diabetes
Global warming
War
Covid-19
...



Still perspectives are sobering ...

Current World Population

7,825,597,952

[view all people on 1 page >](#)

TODAY	THIS YEAR
Births today 234,548	Births this year 122,665,931
Deaths today 98,469	Deaths this year 51,498,097
Population Growth today 136,079	Population Growth this year 71,167,835

worldometers.info/coronavirus

Normal death rate: $\frac{112618 \cdot 366}{7825172174} \cdot 100 =$
0.53 %/year: **112,000** per day ←

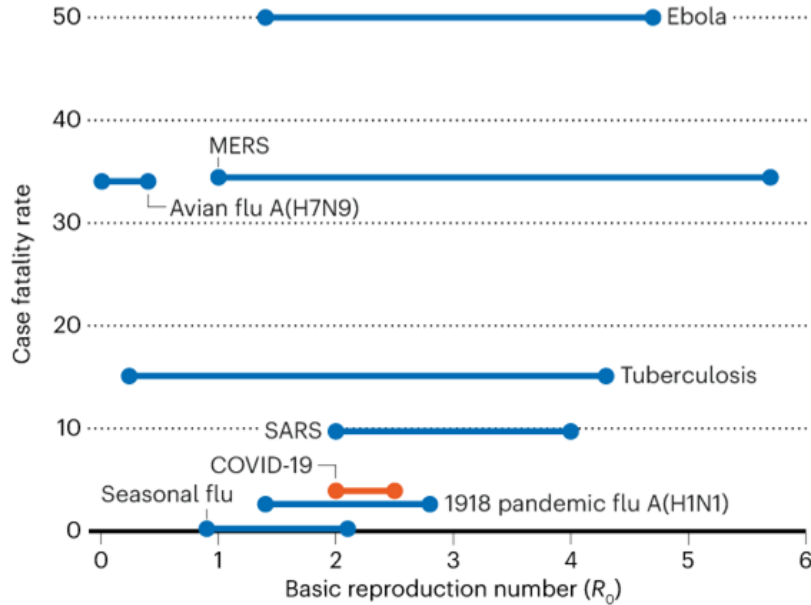
Corona death rate:
 $\frac{1303830}{7825172174} \cdot 100 = 0.017 \% =$
one thirtieth of a normal year
7 months; per day 9,333 ←

Hunger:
847,418,234 Undernourished people in the world
769,803,040 Obese people in the world
19,973 People who died of hunger today ←

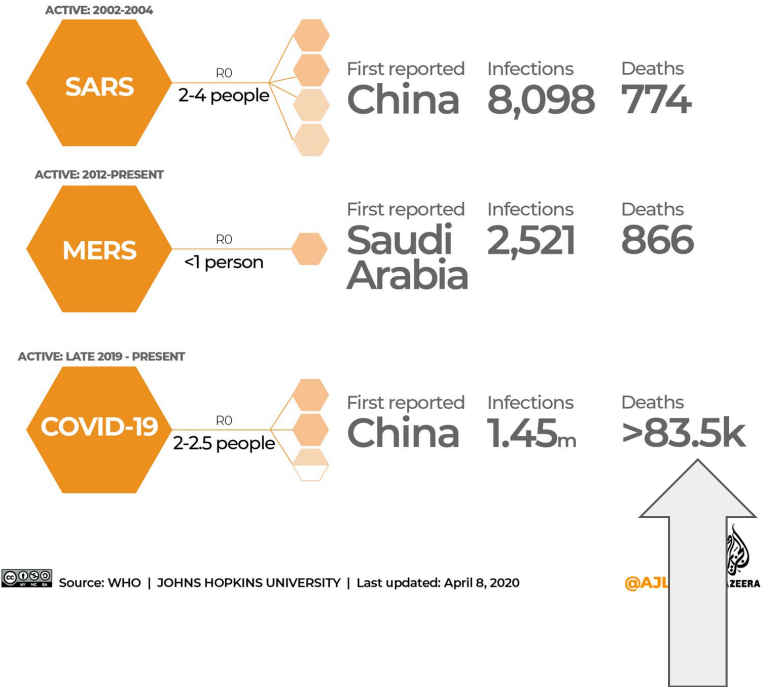
Well, what is the Corona problem then?

- Exponential growth with doubling time of 3.5 days
- Herd immunity is around 60%
- 1.865395 million active cases as of 13 November to world population of 7.825172131 billion = 12 doubling times, i.e. 6 weeks.
- **By the end of the year 60% of the world population could have been infected.**
- Fatality rate will be higher than USA (2%), assume 3%
- Then: **120 Million deaths between now and the end of the year.**
- 4 times higher rate for 2020 than normal.
- **50 times higher rate than normal per day.**
- Swamping of all facilities
- **And this time it is also the rich who die**

COVID-19 vs other 'influenzas'



©nature





Just a cold!



Brazil's President Jair Bolsonaro

Just er influenz

Last updated: November 15, 2020, 11:57 GMT

Coronavirus Cases:

54,443,746

[view by country](#)

Deaths:

1,320,154



Recovered:

37,960,169

worldometers.info/coronavirus

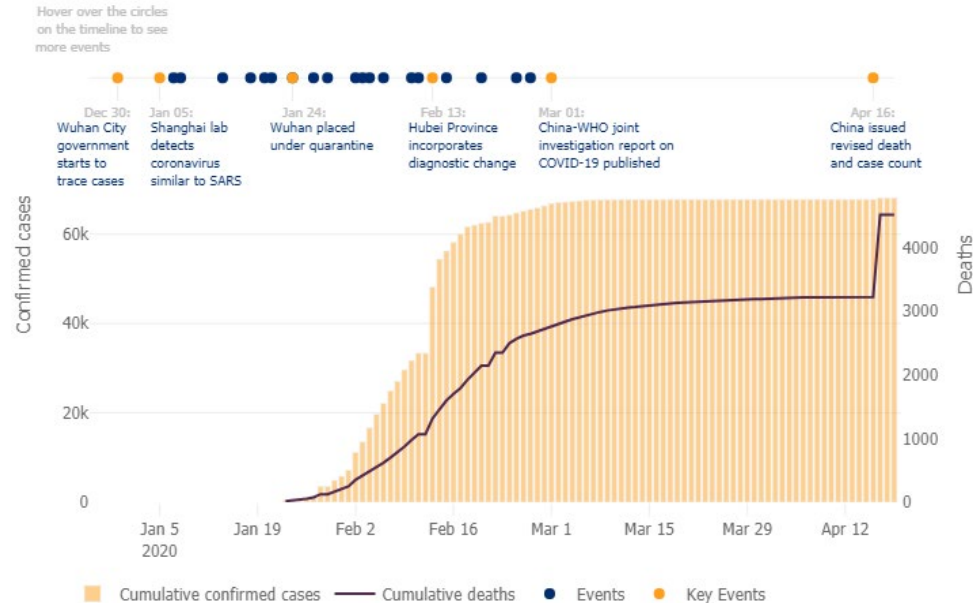
More policy fallacies

- ❖ 'There are no cases in my country' (Bolsonaro)
- ❖ 'We just have to wait for herd immunity' (Boris J.)
- ❖ I am healthy and strong so I won't get it (Bolsonaro)
- ❖ RIVM, Netherlands:
 - 'Mouth masks' merely promote the epidemic
 - 1.5 meter distance should suffice
 - Children do not contract the disease, hence the schools can be kept open
 - Nurses in old-age homes can keep working if they are non symptomatic
- ❖ The virus is only transmitted through droplets in air
- ❖ We can save the economy by preventing lockdown
- ❖ The people will not accept lockdown
- ❖ We can just wait and see and adjust our measures as the epidemic worsens



Can rash government policy *help*?

It correlated in **China** _{CN} and **New Zealand** _{NZ}. This may be sufficient evidence of a cause-effect relation?

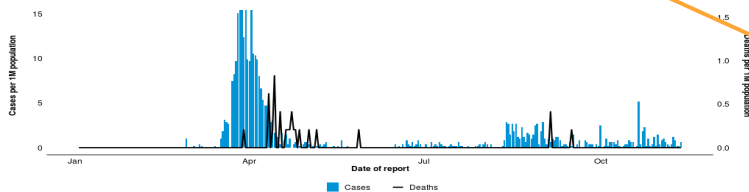


<https://coronavirus.jhu.edu/data/hubei-timeline>

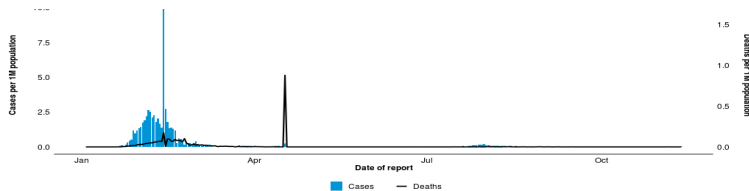
Comparing countries one finds *two scenarios*:

1. Single wave, then flares (China, New Zealand, Taiwan)
2. First, second and third wave (Belgium, Netherlands, USA)

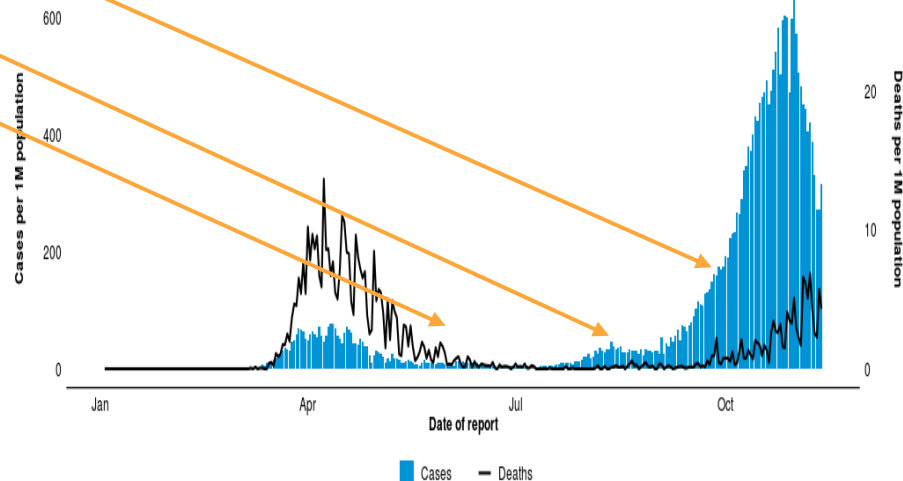
New Zealand



China



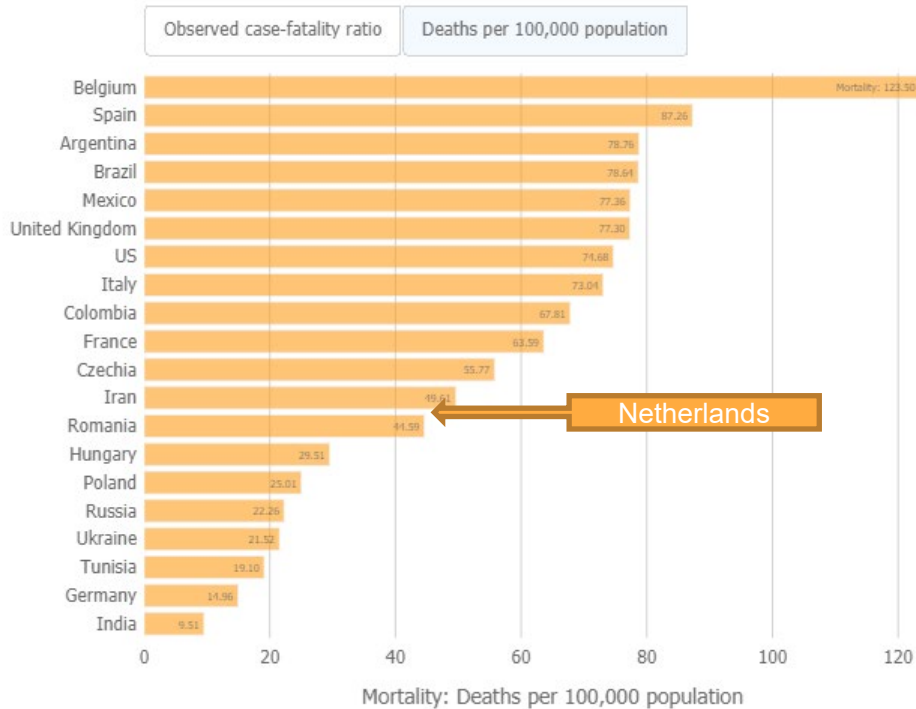
Netherlands





The Ill-performing countries on top

Why these countries?



Large differences not just due to:

- Genes
- Religiousness
- Public health quality
- GNP per capita
- Difference in virus strains
- Political system
- Being an island
- People's government/rules obedience?

Perhaps due to:

- Language (??)
- Disinterested government?
- Elastic government: Trump/Johnson versus Merkel/Ardern
- Amicable social behaviour?
- Privatized health care?

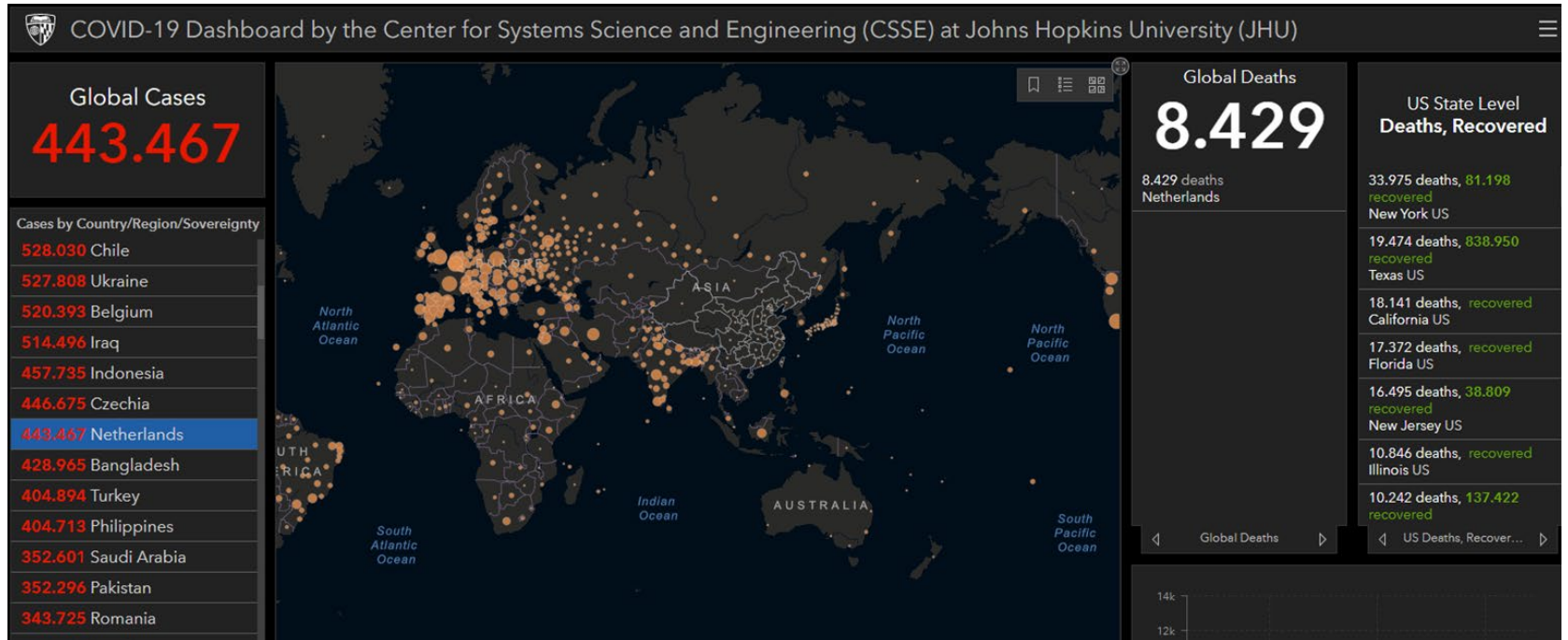
The correlations between policies and persistence of the epidemic is still not clear.

Topic for projects next week?



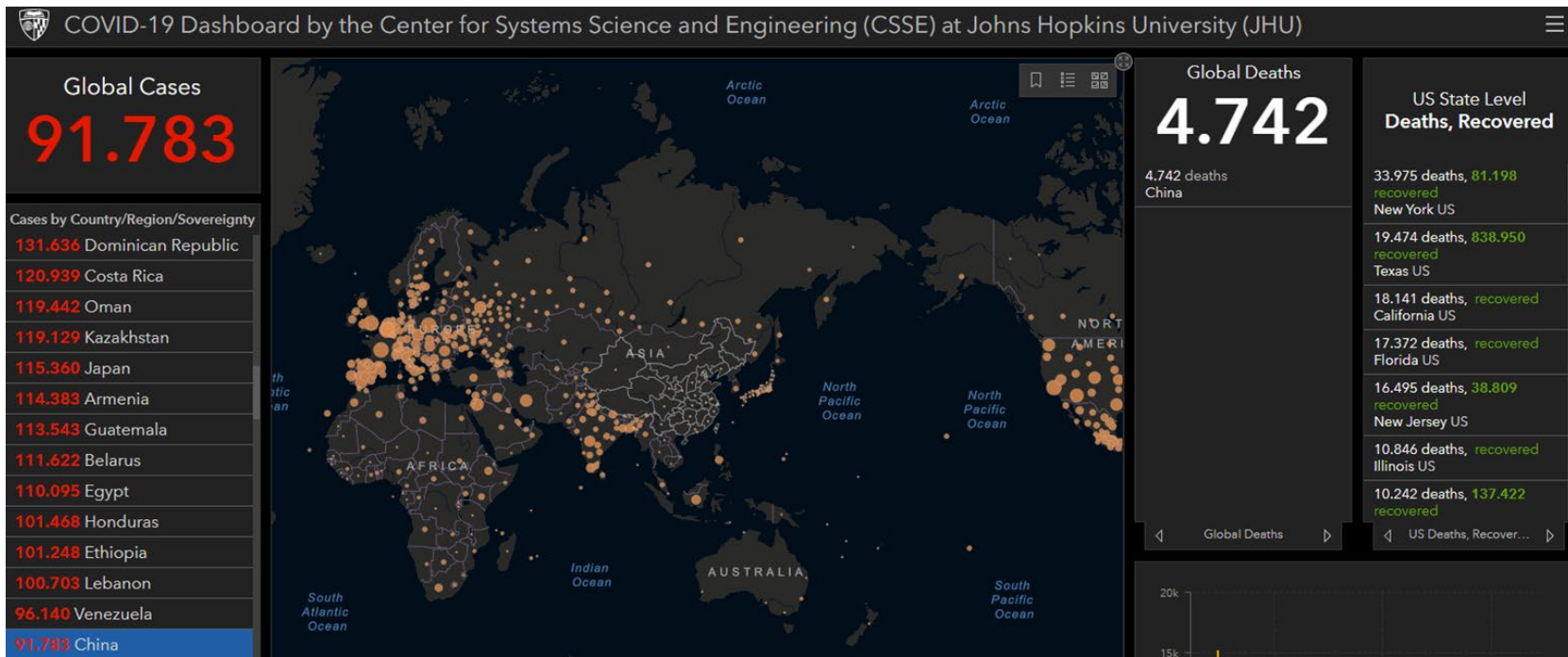
The Netherlands and Corona

(20201113 15h25)



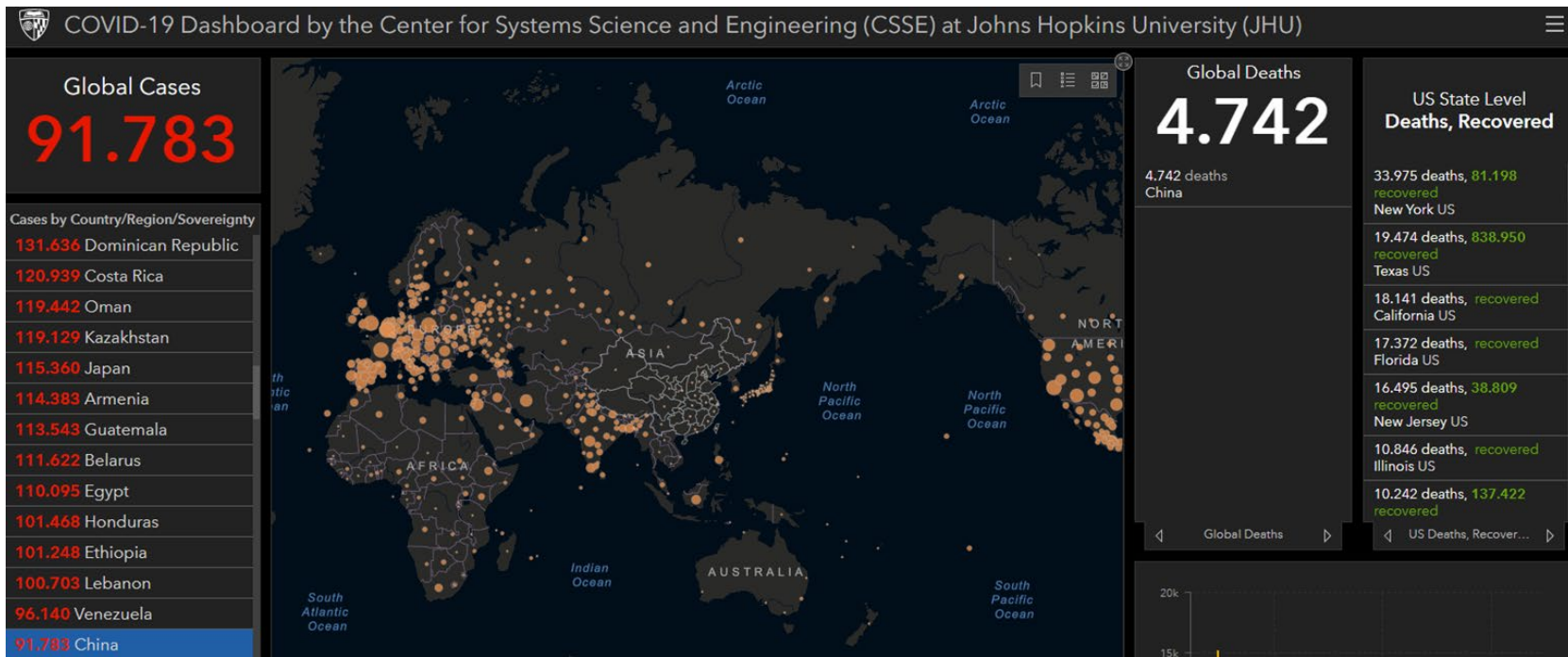
China: where it originated and 60 times more populous: worse or better ?

(20201113 15h25 CET)



China: where it originated and 60 times more populous: > 60 times better !

(20201113 15h25 CET)





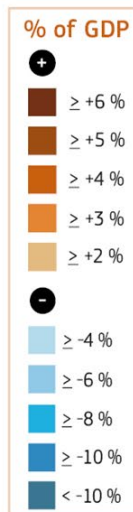
Economic impact



- **Influenza:** The total annual economic burden of influenza epidemics in the United States across all age groups was \$90 billion per year.
- My estimate **for COVID-19:** lockdown by 10% for 6 months: **1.2 trillion** for USA, i.e. some **15 times influenza**

Projected economic *consequences*

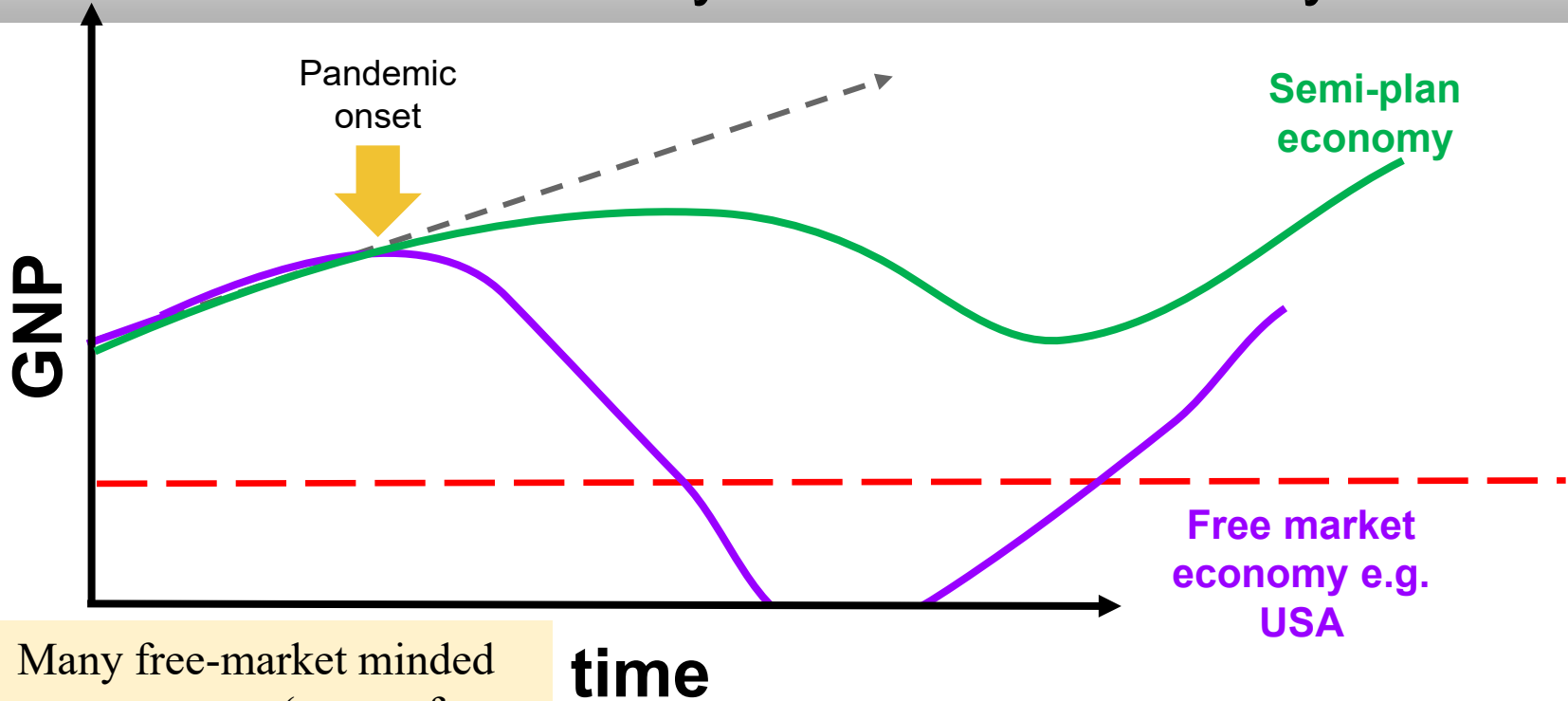
Growth map



Source: European Economic Forecast, Autumn 2020



Plan economy vs market economy



Many free-market minded governments (except for Trump's) used the plan economy option ...

Transmission routes

→ Corona virus secreted from:

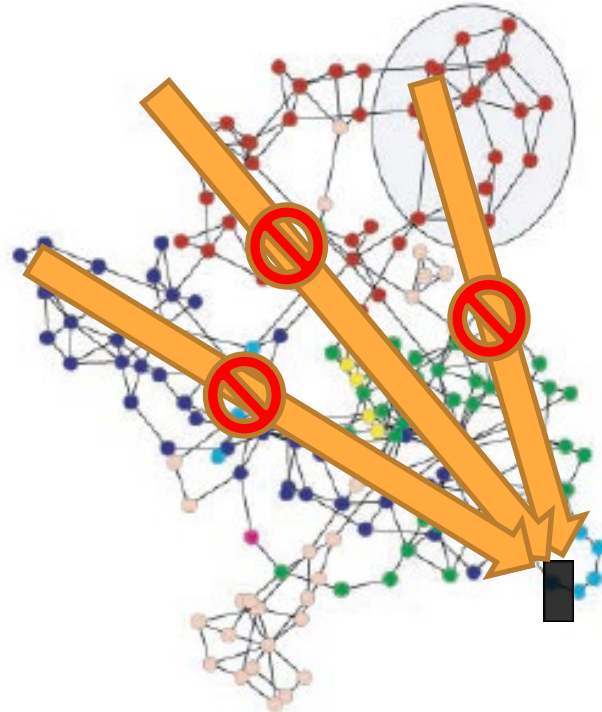
- ◆ Lungs
- ◆ Throat
- ◆ Intestines
- ◆ Urine
- ◆ Post mortem material

→ Hence transfer through:

- ◆ Coughing, shouting, singing: droplets
- ◆ Speaking breathing in close space: aerosols

***Which of these is the one that should be dealt with?
'Which is the rate limiting step?'***

Lesson 1 from Systems Biology



Diseases tend to have multiple (co-) causes

**Both
Simultaneous
and
Alternative causes**

**Multiple causation: all factors
matter**

Impaired function





References: in file

Websites:

- ourworldindata.org/coronavirus
- worldometers.info/coronavirus
- biorender.com/covid-vaccine-tracker (slide)

Scientific articles:

- [Profile of a killer: the complex biology powering the coronavirus pandemic](#)
- [A guide to R- the pandemic's misunderstood metric](#)

Conclusions

The epidemic is complex (many factors, nonlinear)

Policy making is all too often irrational

Can we support this by understandable, open-science, modelling?

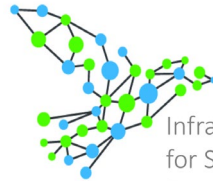
Let us see during this course and thereafter



Thank for your attention and thanks



Rebecca Ludwig



Infrastructure
for Systems Biology
Europe **Netherlands**
ISBE.NL



Alexey Kolodkin
Roland Kraus
Veronica Codoni



This project has received funding from The European Union's Horizon 2020 research and innovation programme under grant agreement No 824087