On the retrospective analysis of COVID-19 macroscopic epidemiological data

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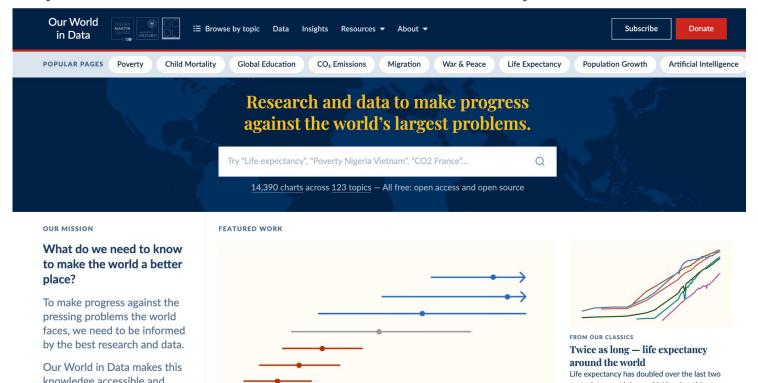
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- Clustering of countries
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- The case of Israel
- The origin of the virus
- The next pandemic

Motivation

- After five years we can look back, can't we?
- Posing question rather than giving answers
- Preparedness for coming pandemics
- The war on virus

Data source

Mostly the "Our World in Data" site of the University of Oxford

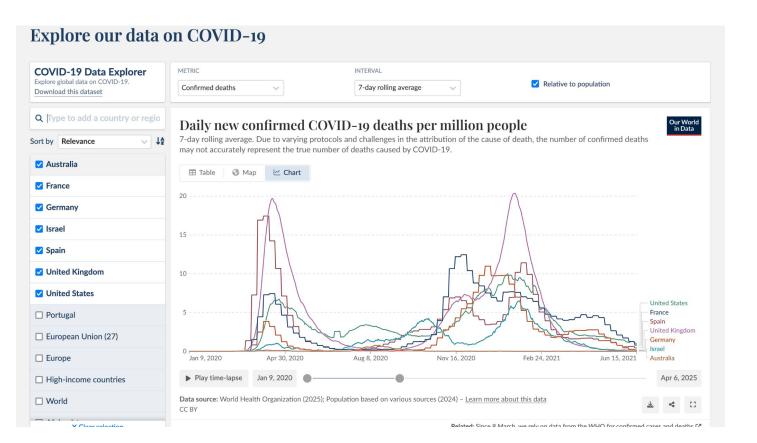


The first wave

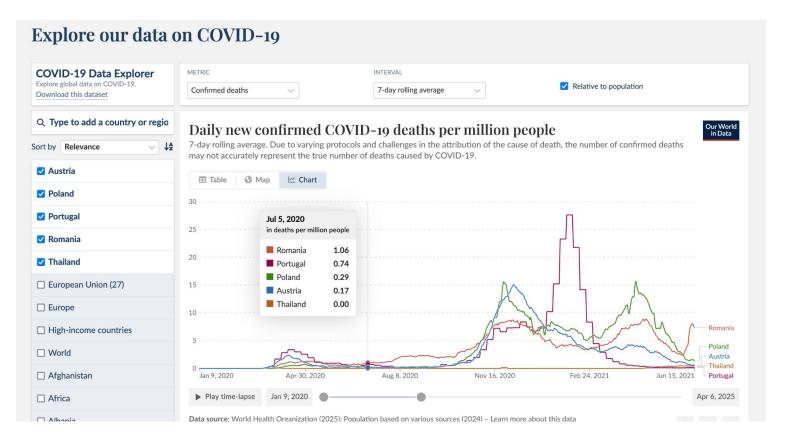
The first wave (mortality)

- In the spring of 2020 several countries suffered (strong) mortality peaks,
- Others didn't
- synchronization of peaks seemed independent of geographical distance

first wave



first wave



First wave

In January 2020, the World Health Organization introduced a disease code specific for COVID-19

Some countries did use this codification since the early stages of the pandemic, others lagged behind

In some countries, COVID-19 cases were paid more than other respiratory diseases, the hospital administration was rewarded for using the COVID-19 disease code

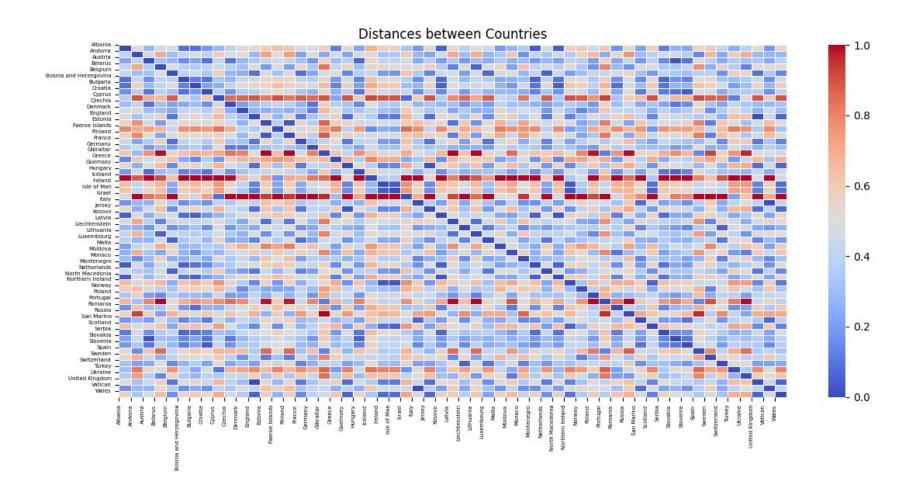
Propagation of the virus in Europe in 2020

Propagation of the virus in Europe in 2020

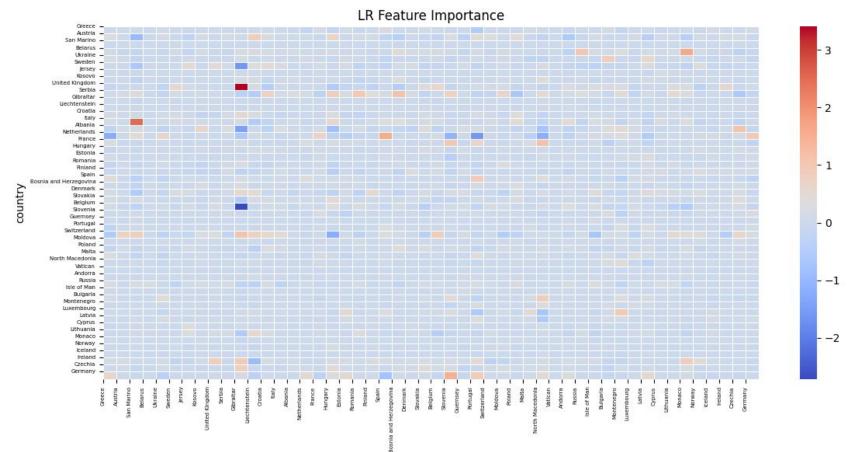
We have considered the mortality in European countries

We correlate similarity measures among mortality time series with distances among country capital cities

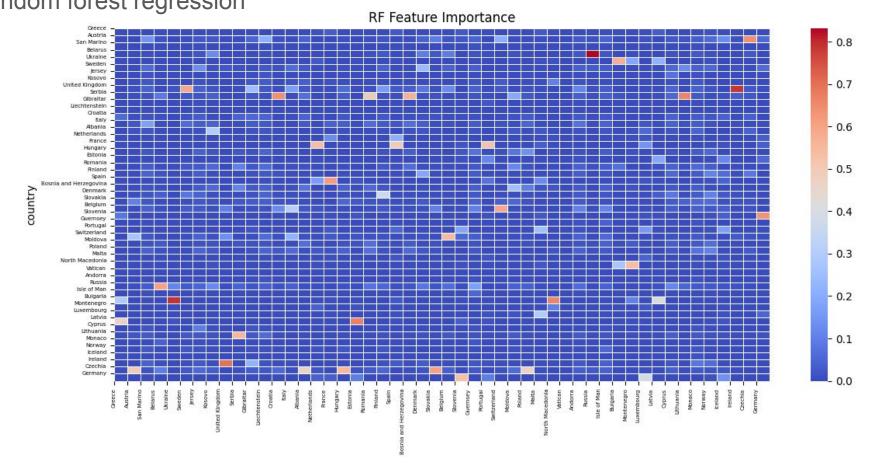
The idea is that more distant countries should be less similar



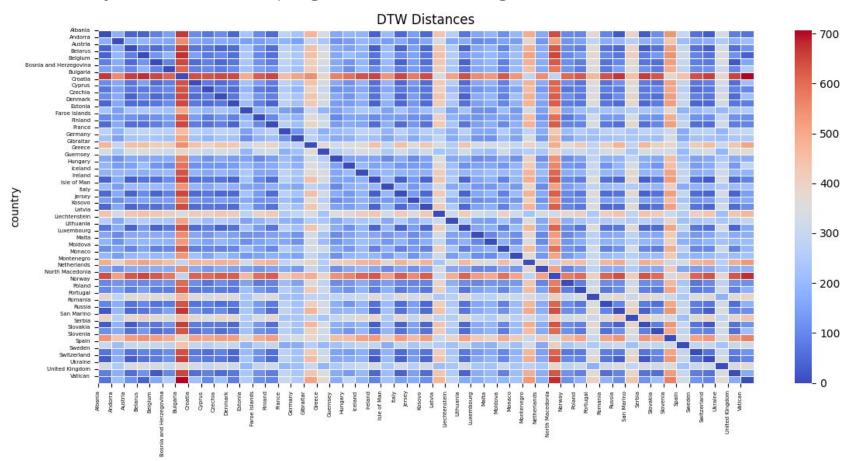
significance of each country when trying to predict deaths in other countries by Linear regression



significance of each country when trying to predict deaths in other countries by random forest regression



dynamic time warping distances among death time series



Propagation of the virus in Europe in 202

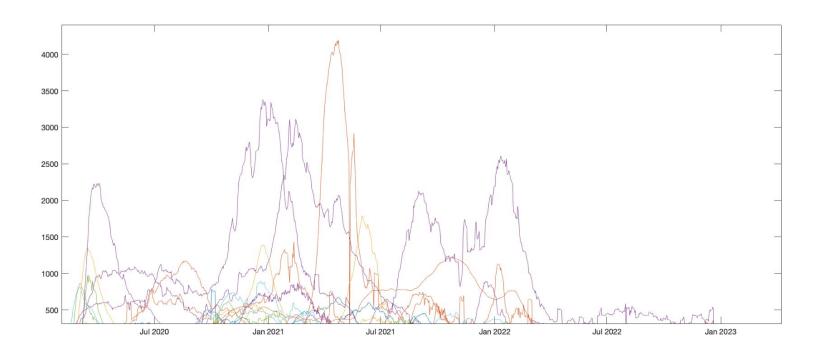
- Pearson's Correlation Coefficients found against geographical distance:
 - o DTW distance r ≈ -0.60
 - Random Forest regression importance r ≈ -0.50
 - Linear regression importance r ≈ -0.35
- Conclussion: there is no influence of geographical distance on the similarity of the death time series
- Is there any viral diffusion process?

Clustering of countries

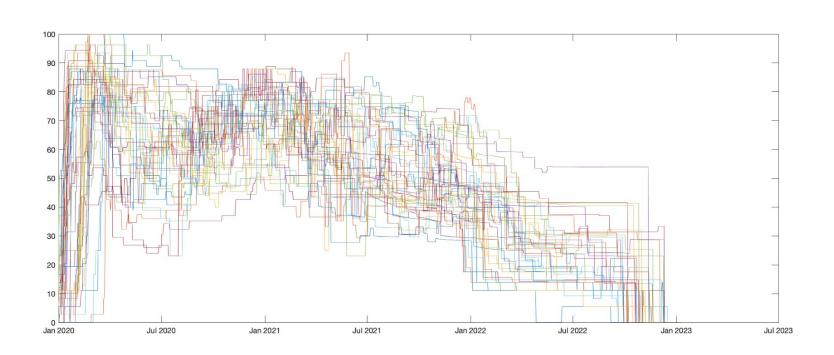
Clustering of countries

- We compute the Dynamic Time Warping (DTW) similarity among the main epidemiological variables of countries
 - death time series
 - stringency index
 - vaccination
 - we select countries with the highest relative accumulative death toll
- We carry out hierarchical clustering of DTW distances
- we find relevant clusters of countries with similar policies and results

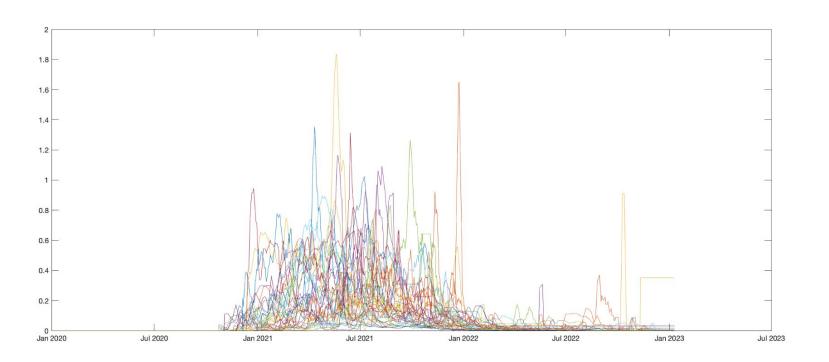
death time series of selected countries

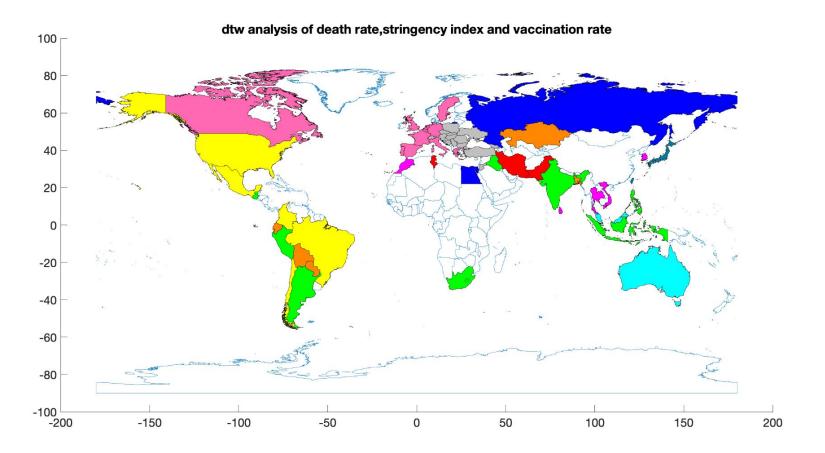


stringency index time series of selected countries



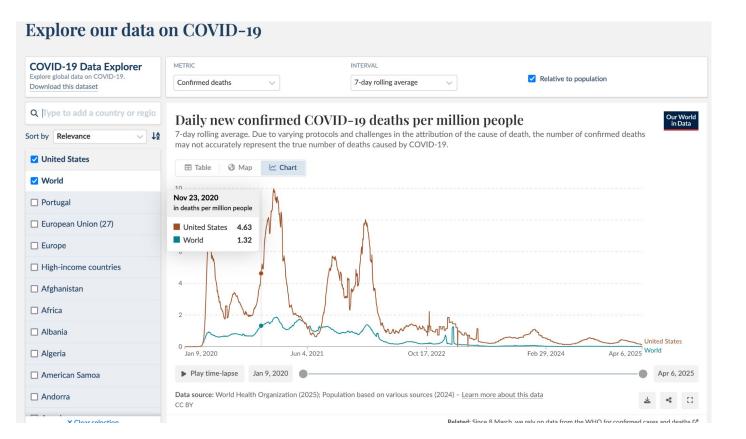
vaccine doses time series of selected countries





The case of USA

The case of USA

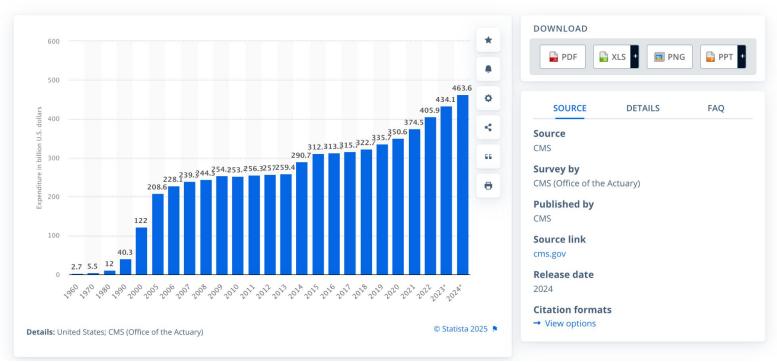




Health, Pharma & Medtech > Pharmaceutical Products & Market

Prescription drug expenditure in the United States from 1960 to 2024

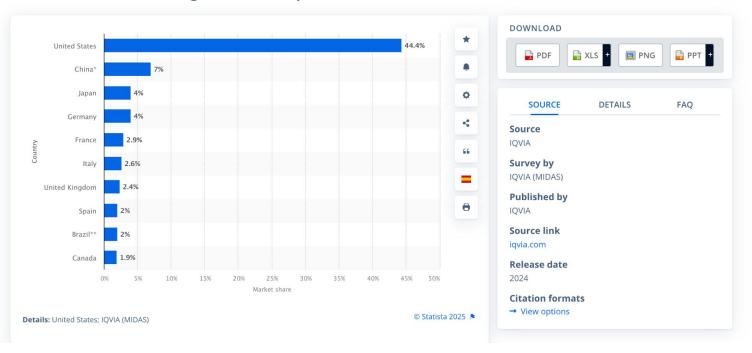
(in billion U.S. dollars)





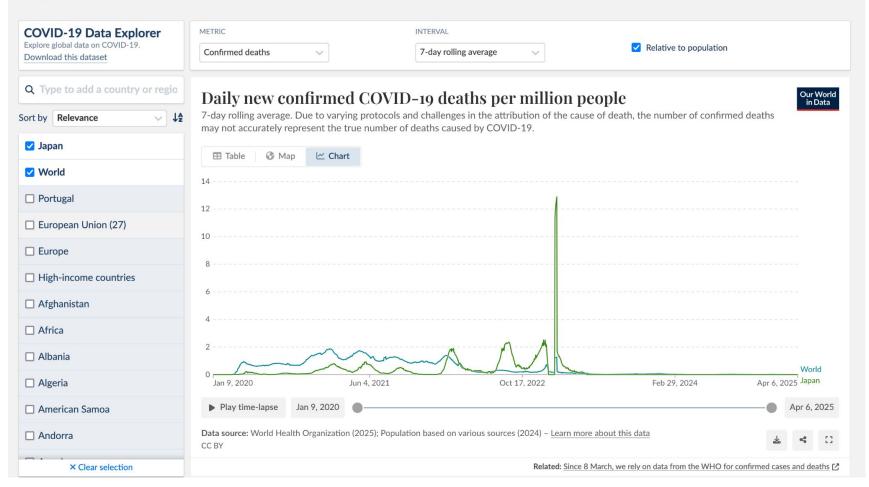
Health, Pharma & Medtech > Pharmaceutical Products & Market

Market share of leading 10 national pharmaceutical markets worldwide in 2023

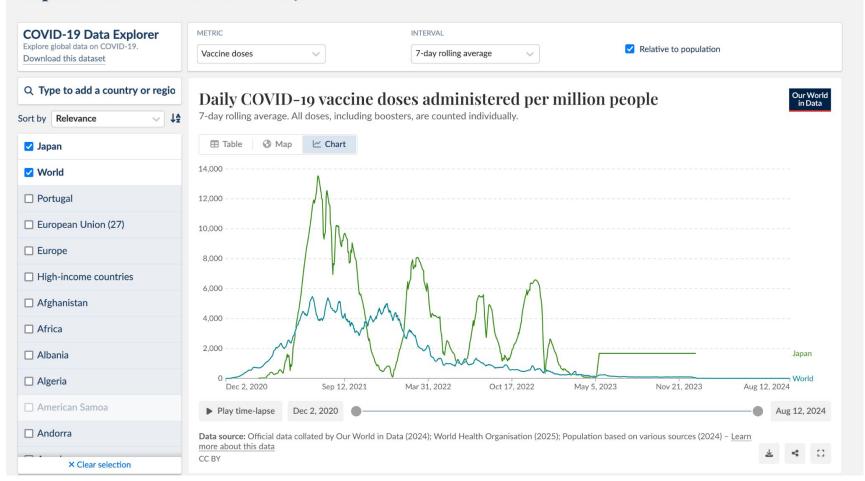


The case of Japan

Explore our data on COVID-19



Explore our data on COVID-19

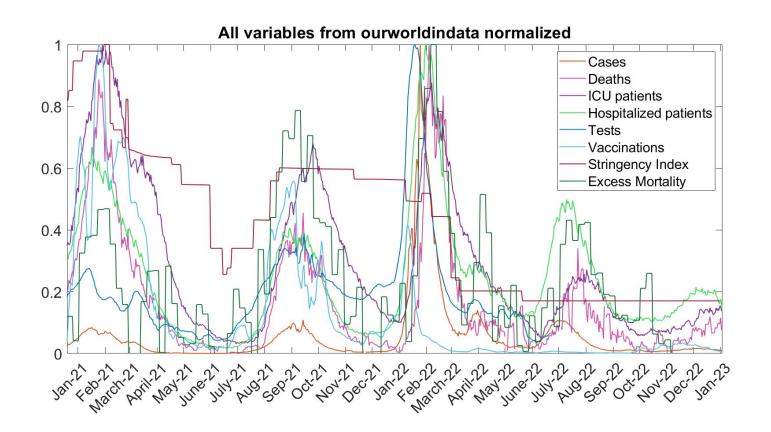


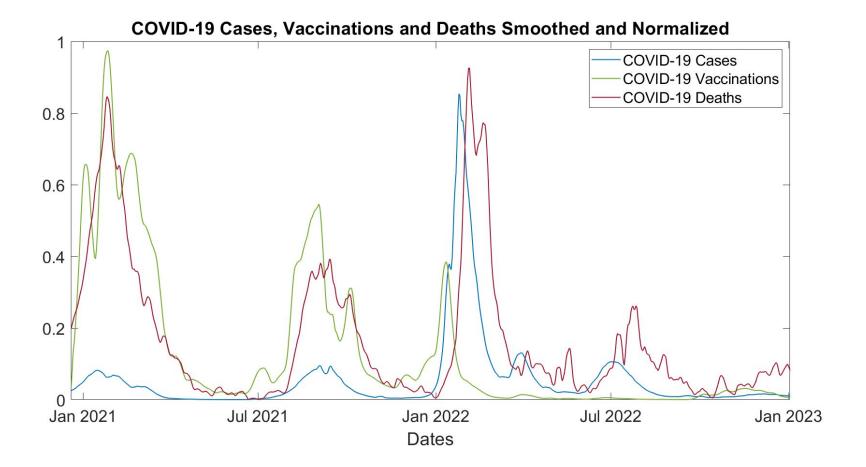
The case of Israel

The case of Israel

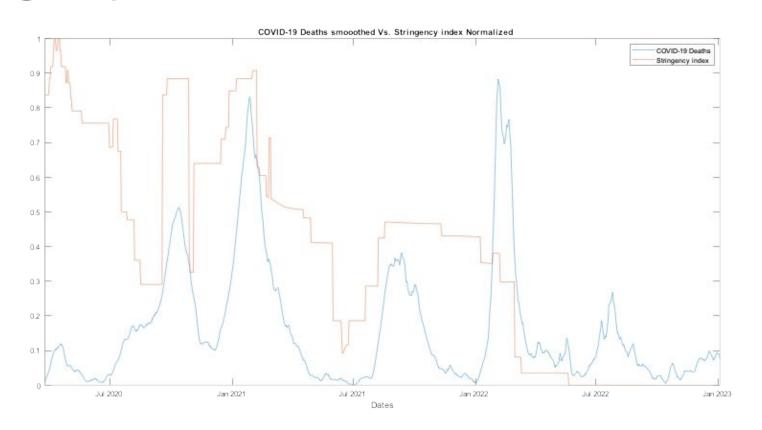
- Israel was the showcase of Pfizer-Biontech vaccine
- Israel was in the forefront of non-pharmaceutical measures

The case of israel

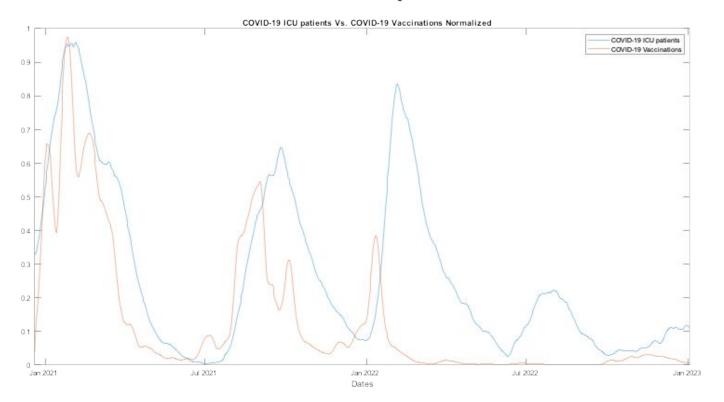




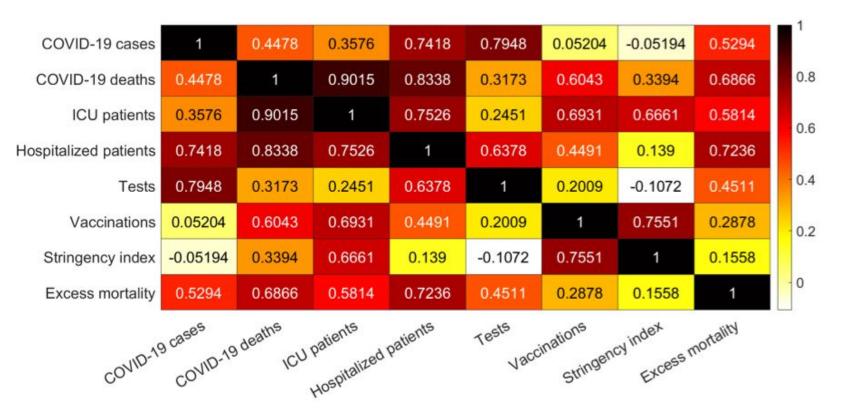
Stringency index, deaths • Normalized to [0,1]



Vaccination doses, ICU patients Normalized to [0,1]



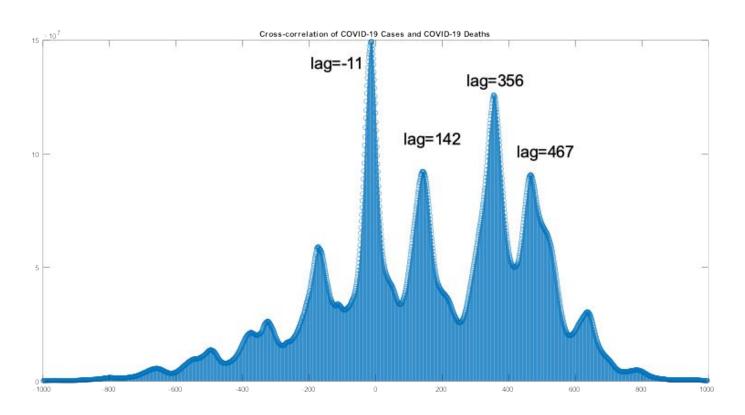
Overall Correlation



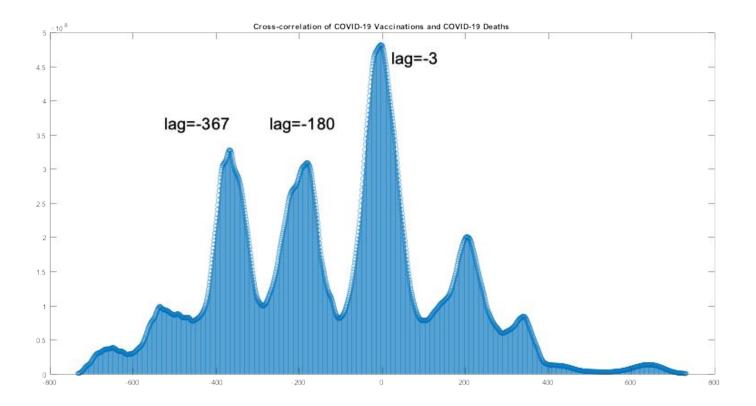
Overall signal correlations 2021-2022

- Non-surprising positive correlations:
 - Deaths cases, Deaths ICU patients
- Surprising positive correlation
 - Deaths Vaccination
- Surprising null correlation
 - Stringency index deaths and cases

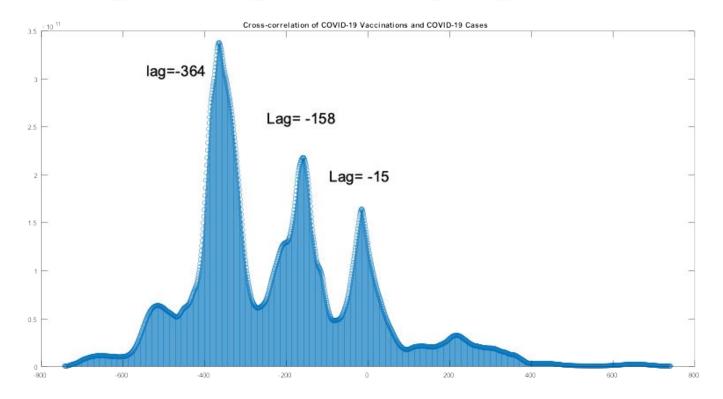
xcorr(new_cases_smoothed, new_deaths_smoothed)



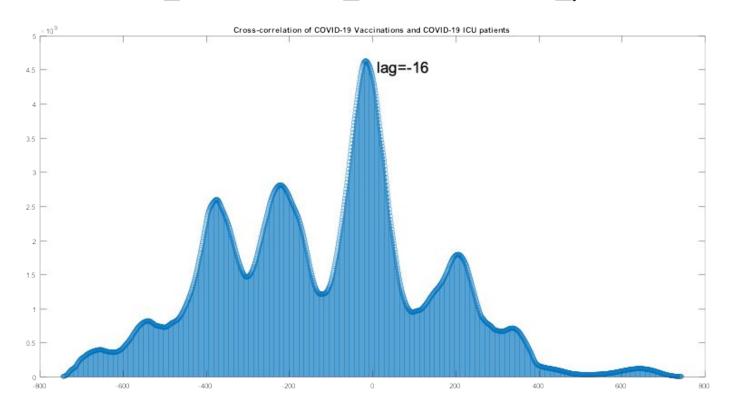
xcorr(new_vaccinations_smoothed, new_deaths_smoothed)



xcorr(new_vaccinations_smoothed, new_cases_smoothed)



xcorr(new_vaccinations_smoothed, icu_patients)



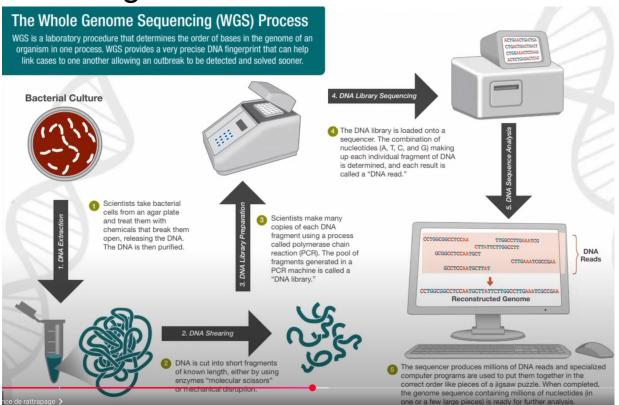
Results from cross-correlation

- Non surprising
 - Cases match/predict deaths after 11 days
- Surprising
 - Vaccination doses match/predict ICU after 16 days
 - Vaccination doses match/predict deaths after 3 days
 - Vaccination doses match/predict cases after 15, 158 days
 - Maximal at 365 days??

https://www.nature.com/articles/s41586-020-2008-3



sample size
n=1
analysis based on a
sample of
pulmonary tissue of
a person with
Pneumonia



the sample biological material is the basis for a sequencing process, whose critical component is a software looking for the **most likely** sequence similar to a given pattern: i.e. SARS-COV-1

56 millions de chaînes de 150 nucléotides

Erreurs de réplication :

- substitutions
- déletions
- insertions

=> 300 000 erreurs

Erreurs de mesure

=> 8,4 millions d'erreurs

Main section

Renewed de novo assembly of published sequence data

To repeat the de novo assembly, we downloaded the original sequence data (SRR10971381) from 27/01/2020 on 11/30/2021 using the SRA tools [19] from the Internet. To prepare the paired-end reads for the actual assembly step with Megahi (v.1.2.9) [20], we used the FASTQ preprocessor fastp (v.0.23.1) [21]. After filtering the paired-end reads, 26,108,482 of the original total of 56,565,928 reads remained, with a length of about 150 bp. A large proportion of the sequences, presumably a majority of those of human origin were overwritten by the authors with "N" for unknown and therefore filtered out by fastp. This is to be regarded as problematic. In the sense of scientificity, since not all steps can be retraced or reproduced. For the elaborate contiguence and the default setting.

Some uncertainties and errors in the process

Two sequencing softwares were applied to find the "most likely" sequence similar to SARS-COV-1

Trinity

1 million de séquences Max : 11 760 nucléotides

Megahit

400 000 séquences

Max: 30 474 nucléotides



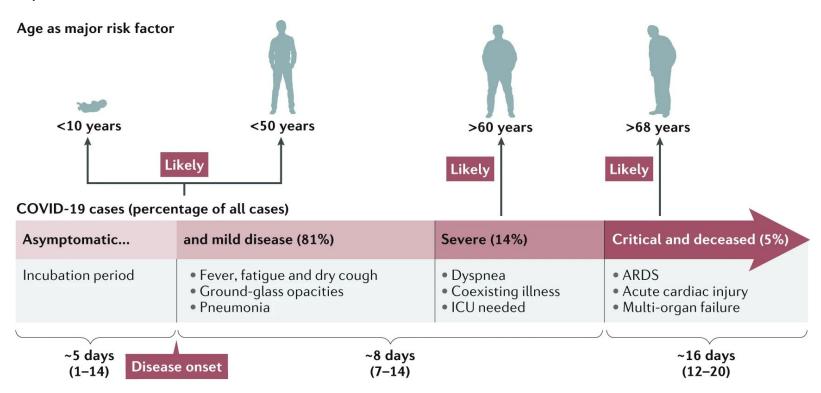
the chosen one

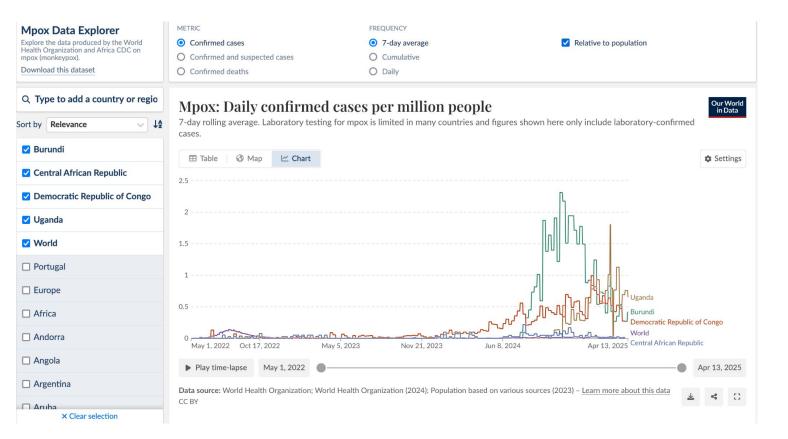
The sequence published in Nature has never been replicated by any researcher in the world

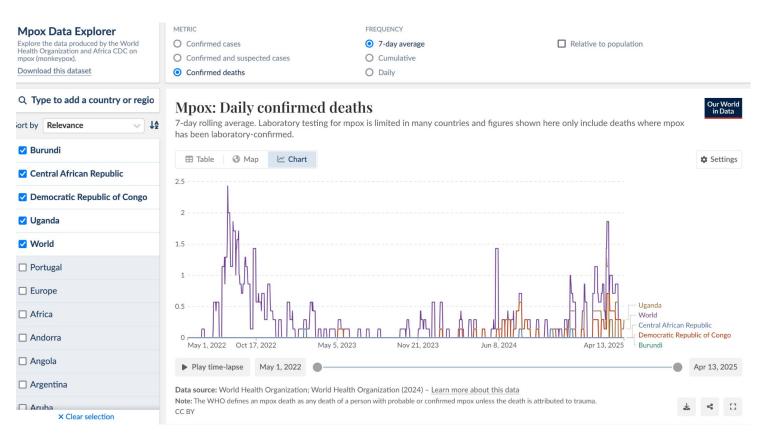
More than 6 M variants

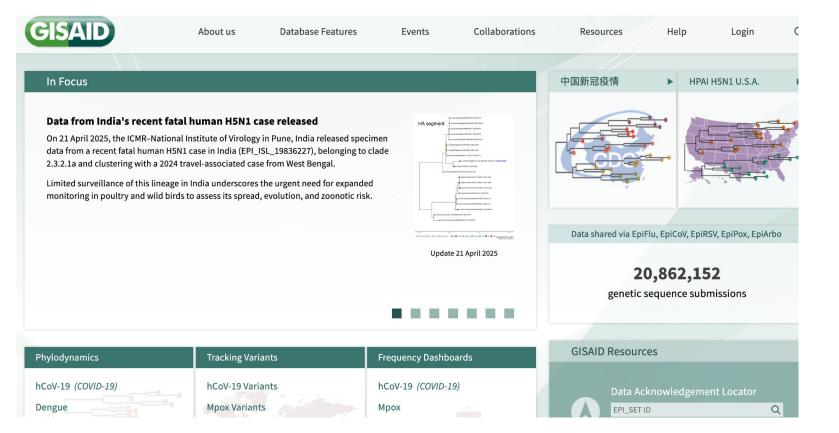
It has been corrected in the GISAID (https://gisaid.org/) repository

https://www.nature.com/articles/s41579-020-00459-7









H5N1 Bird Flu continues to take its toll in the United States

Clade 2.3.4.4b of the highly pathogenic avian influenza (HPAI) virus causing outbreaks in wild and domestic birds around the world continues to spread in dairy cows, poultry and other animals across the United States. Since April 2024, the U.S. CDC confirmed 70 human cases through genome sequence analysis. Data in some of these cases showed the amino acid substitution NA-S247N known to slightly reduce susceptibility to the neuraminidase inhibitor oseltamivir in laboratory tests. In one case, a different change in the polymerase acidic (PA) protein was detected.

On 19 March 2025, the CDC released specimen data from the first human case in Ohio. he specimen sequence belongs to genotype D1.3. Previously the CDC released data from the first human fatality involving a patient from Louisiana that had been exposed to non-commercial backyard poultry and wild birds.