Вебинар. Международная лаборатория исследований населения и здоровья. Высшая школа экономики. 9 апреля 2020 г.

COVID-19: количественная оценка пандемии и ее воздействия на уровень смертности

COVID-19: quantification of the pandemic and its impact on the level of mortality

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Covid-19 cases and deaths: comparability of the data



Covid-19 pandemic

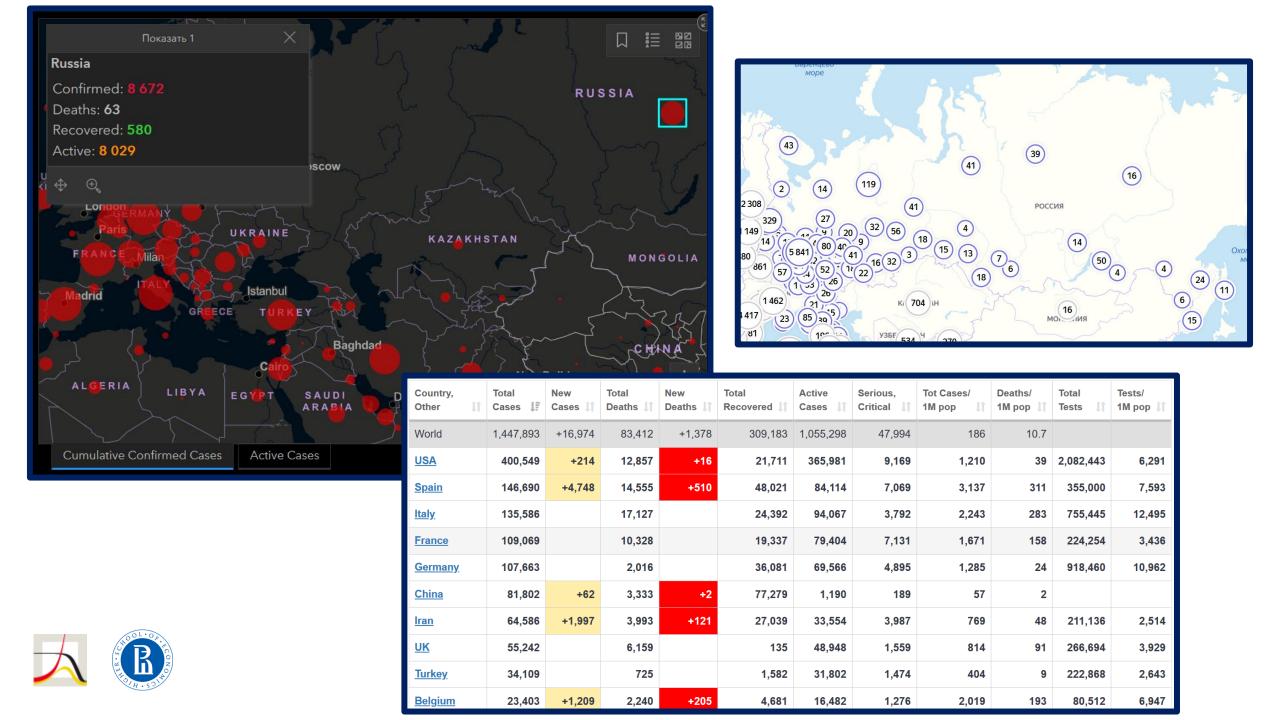
31 December 2019 – the cases of pneumonia of unknown cause reported to WHO by China

7 January 2020 – a novel coronavirus (2019-nCoV -> SARS-CoV-2) was identified

11 March 2020 – WHO announces a pandemic

First pandemic since 2009 (influenza A (H1N1))

First pandemic caused by a coronavirus



Our World COVID-19: Total confirmed cases vs. Total confirmed deaths, Apr 6, 2020 in Data The number of confirmed cases is lower than the number of total cases. The main reason for this is limited testing. The grey lines show the corresponding case fatality rates, CFR (the ratio between confirmed deaths and confirmed cases). LOG Africa 10,000 France Asia United Kingdom Europe China North America Belgium Germany Oceania Switzerland Brazil Total confirmed deaths due to COVID-19 South America 1,000 Denmark 0.25% Algeria • India Egypt Peru 100 Our \ in [Case fatality rate of the ongoing COVID-19 pandemic **Czech Republic** Argentina The Case Fatality Rate (CFR) is the ratio between confirmed deaths and confirmed cases. Australia San Marino Finland During an outbreak of a pandemic the CFR is a poor measure of the mortality risk of the disease. We explain this in Andorra detail at OurWorldInData.org/Coronavirus Croatia United Arab Emirates 🔂 Add cou Bangladesh 10 Singapore Italv 12% Bahrain Qatar Sint Maarten (Dutch part) 🐠 0 France Monaco Georgia Costa Rica Sudar 10% Mauritania Gabon Liechtenstein Kuwait 1,000 10,000 4 10 100 8% Total confirmed cases of COVID-19 Source: European CDC - Situation Update Worldwide - Last updated 6th April, 12:00 (London time) 6% Sweder 4% China

2%

0%

Feb 25, 2020

Source: European CDC - Situation Update Worldwide - Last updated 6th April, 12:00 (London time)

Mar 16, 2020

Mar 26, 2020

Mar 6, 2020

South Korea Germany

Apr 6, 2020



Key indicators

Spread of the disease:

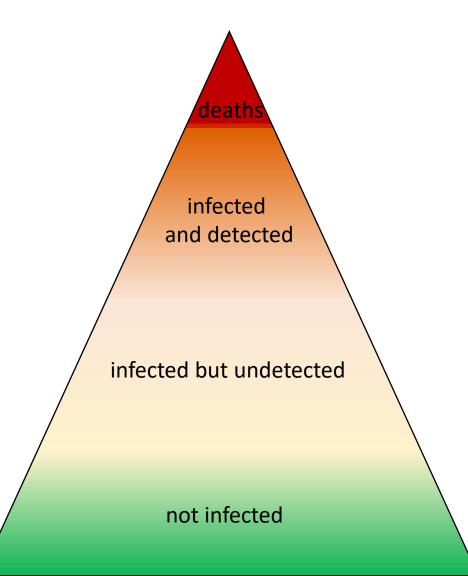
How many are infected in the population? How it evolves in time?

Lethality:

How many of those infected will die? How it depends on age and sex?

Mortality:

How many deaths will appear in the population due to the disease outbreak? How will it affect mortality and life expectancy trends?







WHO definition (for Global surveillance purposes):

Confirmed case A person with laboratory confirmation of COVID-19 infection, irrespective of clinical signs and symptoms.

Colorado. U.S.

Positive cases include people who tested positive, as well as cases where epidemiological investigation has determined that there is a high likelihood that an untested individual has COVID-19 due to their symptoms and close contact with someone who tested positive for COVID-19. The number of untested positive individuals included in the total positive sum is a very small portion of the total positives reported.



Testing capacities

Three months since the new virus was identified

Countries had to ramp up their testing and laboratory capacities in a short time

A huge gap between countries

"intensified COVID-19 molecular testing has led to shortages of molecular testing reagents globally for COVID-19 and for other molecular diagnostics" [WHO, 21 March 2020]

WHO Testing strategy recommendations

Countries that have not yet reported cases Countries dealing with sporadic cases Countries dealing with clusters of cases

- All suspected cases

Suspect case

- a) Acute respiratory illness AND a travel in a location reporting community transmission
- b) Acute respiratory illness AND contact with Covid-19 case
- c) Severe acute respiratory illness AND requiring hospitalization AND in the absence of alternative diagnosis

Countries dealing with community transmission

Testing constraints should be anticipated, and prioritization will be required

Different testing strategies might be needed within the same country





Testing strategies

Testing only those who fits the criteria (decision is made by doctors):

- Most of the countries

Testing all interested:

- Iceland. 32623 samples tested as of April, 8 (NUHI + deCode Genetics voluntary screening)

Russia. Commercial labs in large cities offer tests for SARS-CoV2, positive or inconclusive tests are sent to governmental laboratories to be checked

- U.S. Drive-thru testing sites. For those with symptoms (including self-reported). Currently closed (?)

How criteria change in time. Example of Germany

21.01.2020

Clinical, radiological or histopathological evidence of pneumonia + stay in risk area (China, Wuhan, Hubei Province) Acute respiratory symptoms + Covid-19 contact

23.01.2020

Clinical or radiological evidence of acute infection of the lower respiratory tract + stay in risk area

Acute respiratory symptoms + Covid-19 contact

10.02.2020

Non-specific general symptoms or acute respiratory symptoms + Covid-19 contact Acute respiratory symptoms + stay in risk areas



How criteria change in time. Example of Germany

26.02.2020

Non-specific general symptoms or acute respiratory symptoms + Covid-19 contact Acute respiratory symptoms + stay in risk areas Acute respiratory symptoms + stay in regions with Covid-19 cases Clinical or radiological evidence of viral pneumonia without alternative diagnosis

24.03.2020

Acute respiratory symptoms + Covid-19 contact

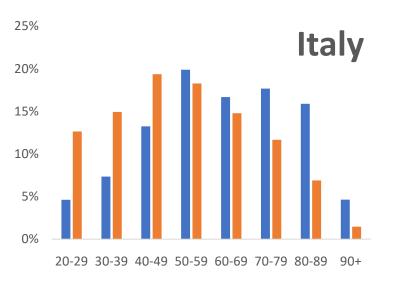
Clinical or radiological evidence of viral pneumonia with increased pneumonias in nursing homes/hospitals

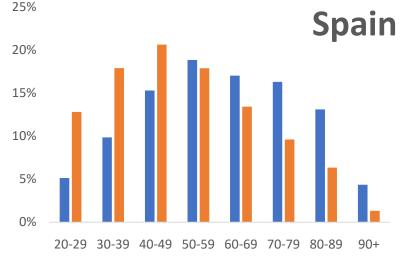
Clinical or radiological evidence of viral pneumonia without alternative diagnosis Acute respiratory symptoms + work in nursing, medical practice or hospital

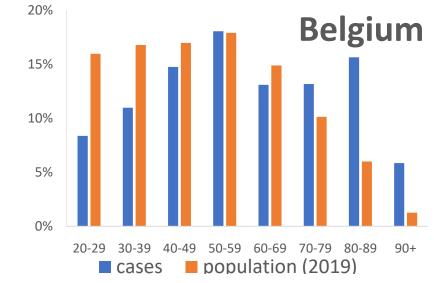
+ belonging to a risk group

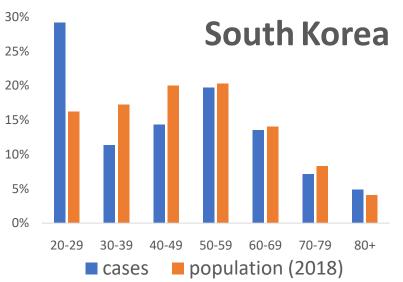
Acute respiratory symptoms w/t known risk factors - COVID-19 diagnostics only with sufficient test capacity

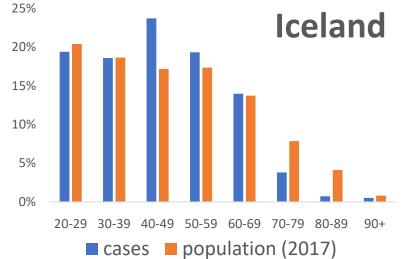
Do testing criteria produce different bias for different age groups?

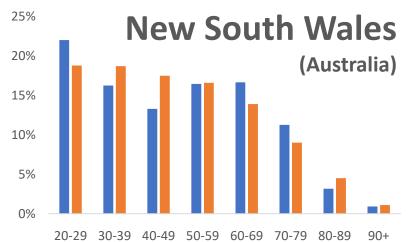


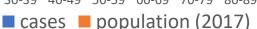










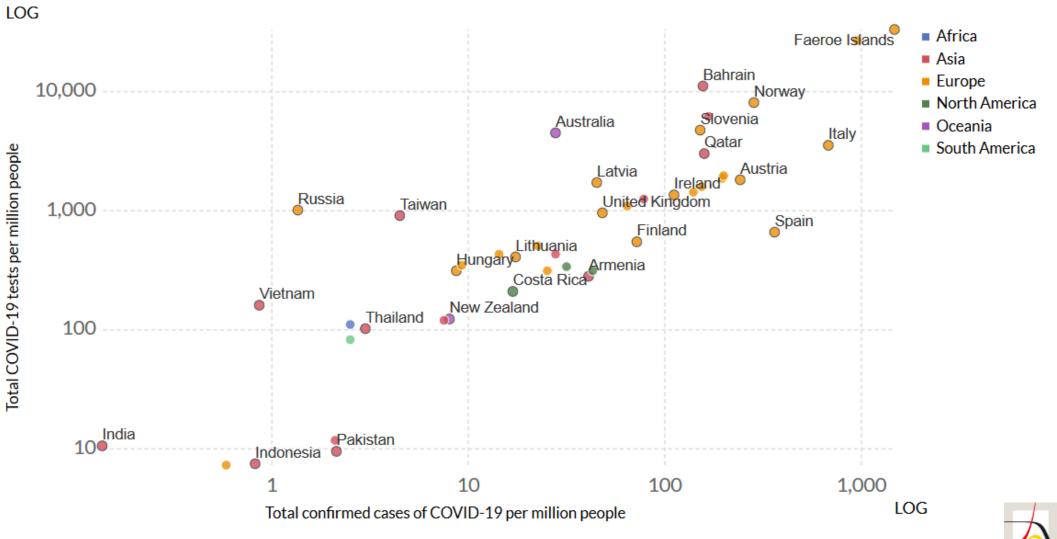


COVID-19 data as of 20 March: Tests conducted vs. Total confirmed cases per million people

Our World in Data

Data collected by Our World in Data from official country reports.

For some countries the number of tests corresponds to the number of individuals who have been tested, rather than the number of samples.



Source: Our World in Data based on official sources Note: Data for the US corresponds to estimates from the COVID-Tracking Project

Post-mortem testing (Influences both: cases and deaths)

U.S.

Postmortem collection of the swabs to be tested for the presence of SARS-CoV-2 in persons under investigation for Covid-19. Can be adapted by state and local health departments depending on the situation [CDC]

Ideally, testing for COVID–19 should be conducted, but it is acceptable to report COVID–19 on a death certificate without this confirmation if the circumstances are compelling within a reasonable degree of certainty [CDC]

England and Wales

If death can be due to COVID-19 (criteria on possible COVID-19 case in living is applied), the decision on post-mortem collection of swabs or further postmortem examination should be taken on a case-by-case basis depending on the needs of the coroner and other relevant parties [The Royal College of Pathologists]

For example, if before death the patient had symptoms typical of COVID-19 infection, but the test result has not been received, it would be satisfactory to give 'COVID-19' as the cause of death, tick Box B and then share the test result when it becomes available. In the circumstances of there being no swab, it is satisfactory to apply clinical judgement [Office for National Statistics]



Post-mortem testing (Influences both: cases and deaths)

Norway

A post-mortem test for COVID-19 should be performed if a patient dies of acute respiratory tract infection of unknown cause in a healthcare institution [Norwegian Institute of Public Health]

Germany

"We don't consider post-mortem testing to be a deciding factor. We assume that patients are diagnosed before they die" [RKI to AFP, 20.03.2020]

Those who had not been tested for COVID-19 during their lifetime but who are suspected of having died of COVID-19 can be examined for the virus postmortem - i.e. after their death. [*BR24, 21.03.2020, according to RKI Spokesperson*]

The RKI says those who were not tested for Covid-19 in their lifetime but are suspected to have been infected with the virus "can" be tested after death, but in Germany's decentralised health system this is not yet a routine practice [The Guardian, 22 March 2020]

Reporting deaths. Dying with or dying from?



Italy

"I would like to point out that these are not deaths "by" coronavirus. These are people who have died and among the various pathologies they also had the coronavirus" [A. Borelli, Head of the Italian Civil Protection Agency, 10 March 2020]

"...we put all the deaths to the numerator without that maniacal attention to the definition of the causes of death that have for example French and Germans, who before attributing a death to the Coronavirus perform a series of checks and evaluations that even in some cases had led to the removing deaths from the list" [W. Ricciardi, the EACHR member, consultant of the Italian Ministry of Health on the Covid-19, 10 March 2020]

"The way in which we code deaths in our country is very generous in the sense that all the people who die in hospitals with the coronavirus are deemed to be dying of the coronavirus. On re-evaluation by the National Institute of Health, only 12 per cent of death certificates have shown a direct causality from coronavirus, while 88 per cent of patients who have died have at least one pre-morbidity - many had two or three" [W.Ricciardi, The Telegraph, 21 March 2020]



U.S.

"There are other countries that if you had a pre-existing condition, and let's say the virus caused you to go to the ICU [intensive care unit] and then have a heart or kidney problem. Some countries are recording that as a heart issue or a kidney issue and not a COVID-19 death. [...] The intent is ... if someone dies with COVID-19 we are counting that" [D. Brix, the response coordinator for the White House coronavirus task force, 7 April 2020]



Germany

"Bei uns gilt jemand als Corona-Todesfall, bei dem eine Corona-Infektion nachgewiesen wurde." [L.Wieler, RKI President, 20.03.2020]

Newspaper BR24

Der #Faktenfuchs hat beim RKI nachgefragt, wer nun genau als Corona-Todesfall zählt.

Das RKI zählt laut Angaben einer Sprecherin als Corona-Todesfälle alle Menschen, die mit einer COVID-19-Erkrankung in Verbindung stehen.

Dazu gehören erstens Menschen, die direkt an der Erkrankung gestorben sind ("gestorben an"). Und zweitens Patienten mit Grundkrankheiten, die mit COVID-19 infiziert waren und bei denen sich nicht klar nachweisen lässt, was letzten Endes die Todesursache war ("gestorben mit"). [according to RKI Spokesperson]

Iceland



"A foreign tourist who yesterday came to the Health Care Institution of North Iceland with a serious illness turned out to be infected with the virus that causes the COVID-19 illness. The man died shortly after he arrived at the institution. The cause of death is not known, but his symptoms were not typical for COVID-19. The cause of death is not yet known. [Office of the Medical Director of Health, 17 March 2020]

"Two of the people diagnosed with COVID-19 in Iceland have passed away." [Office of the Medical Director of Health, 24 March 2020]

Russia

"Причиной смерти стала массивная двусторонняя тромбоэмболия легочной артерии. Несмотря на то что пациентка была госпитализирована с начинающимися признаками пневмонии и у нее при жизни подтвердился положительный результат анализа на коронавирусную инфекцию, причиной смерти стала не пневмония, а оторвавшийся тромб. На момент смерти изменений в легких не было. Тяжелые сопутствующие хронические заболевания подтвердились, среди них сахарный диабет, цереброваскулярная болезнь, ишемическая болезнь сердца" [О. Зайратьянц, главный Патологоанатом Москвы Dahilova, Shkolnikov, Jdanov. HSE Webinar: COVID-19: Quantification

Covid-19 in medical death certificate



New ICD-10 codes for COVID-19

- •U07.1 COVID-19, virus identified
- •U07.2 COVID-19, virus not identified

o Clinically-epidemiologically diagnosed COVID-19

o Probable COVID-19

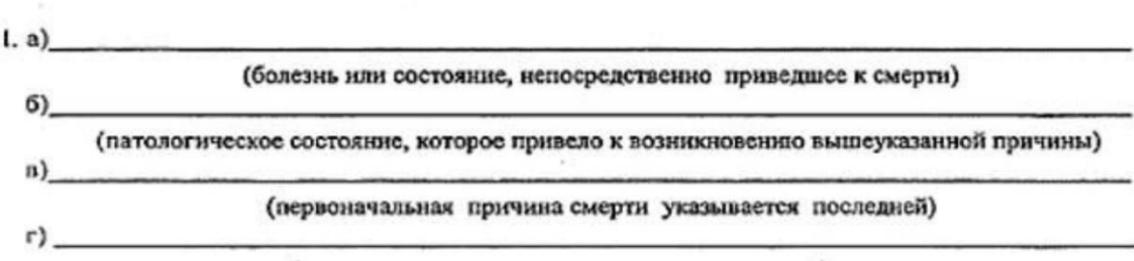
o Suspected COVID-19

Mortality Coding of COVID-19 with ICD-10

Both categories, U07.1 (COVID19, virus identified) and U07.2 (COVID19, virus not identified) are suitable for cause of death coding. [...] COVID-19 is reported on a death certificate as any other cause of death, and rules for selection of the single underlying cause are the same as for influenza (COVID-19 not due to anything else). For recording on a death certificate, no special guidance needs to be given. The respiratory infection may evolve to pneumonia that may evolve to respiratory failure and other consequences. Potentially contributing comorbidity (immune system problem, chronic diseases...) is reported in part 2, and other aspects (perinatal, maternal...) in frame B, in line with the rules for recording. A manual plausibility check is recommended for certificates where COVID-19 is reported, in particular for certificates where COVID-19 was reported but not selected as the single underlying cause of death.

WHO. COVID-19 coding in ICD-10, 25 March 2020

10. Причины смерти:



(внешняя причина при травмах и отравлениях)

 Прочие важные состояния, способствовавшие смерти, но не связанные с болезнью или патологическим состоянием, приведшим к ней, включая употребление алкоголя, наркотических средств, психотропных и других токсических веществ, содержание их в крови, а также операции (название, дата)



np

Influenza in medical death certificate



ICD. Volume II.

- I (a) Acute myocardial infarction
 - (b) Atherosclerotic heart disease
 - (c) Influenza

After applying Rule I and then Modification Rule C, Acute myocardial infarction should be coded as an underlying cause of death.

CDC. ICD-10-Mortality Manual 2a

- I (a) 1. Coronary occlusion
 - (b) 2. Diabetes, chronic, severe
 - (c) 3. Hypertension and arteriosclerosis
 - 4. Renal disease
- II 5. Influenza, 1 week

Code to coronary occlusion (I219) by applying Selection Rule 2

How the concept of "deaths with" is different from our usual cause-of-death mortality estimates?

Influenza deaths in England, 2016-2017 season

25000

20000

15000

10000

5000

0

Underlying cause of Underlying+Contributory Estimated influenzadeath associated deaths

Deaths with

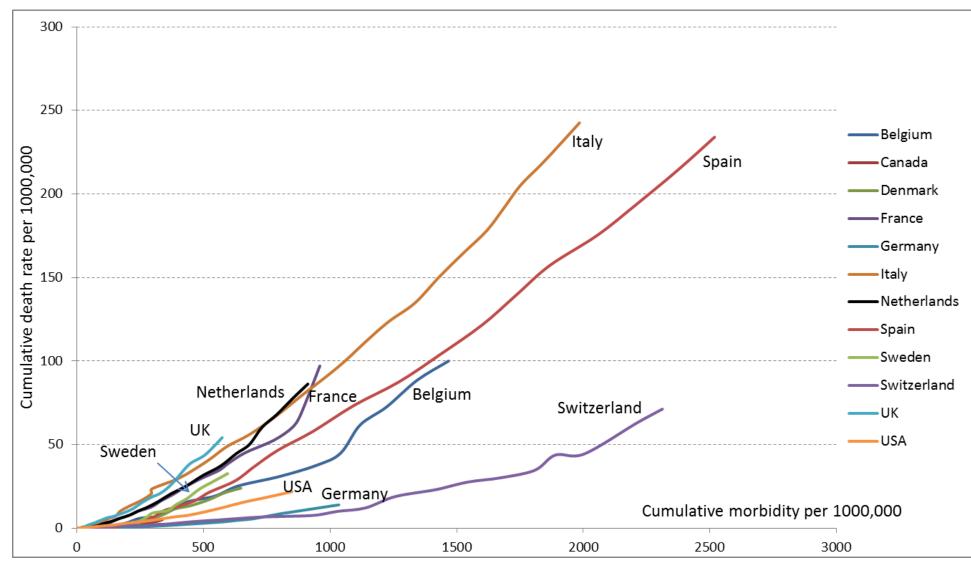


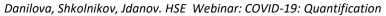
Danilova, Shkolnikov, Jdanov. HSE Webinar: COVID-19: Quantification



An alternative method of estimation: excess mortality from all causes by week or month

Cumulated COVID-19 incidence vs. cumulated COVID-19 mortality: huge variation in countries' trajectories per 1 mln. (1.03-4.04.2020)

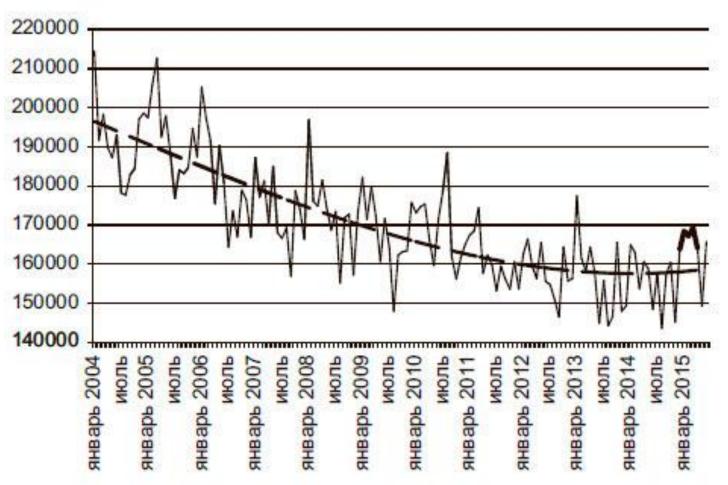








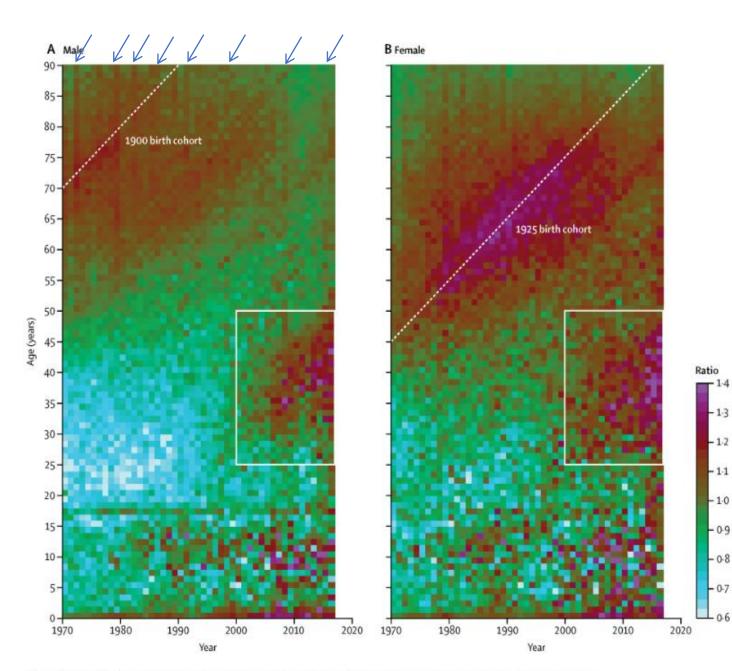
An old idea: assessment of mortality fluctuations across months (weeks, days)



Andreev & Shkolnikov 2016

http://www.demoscope.ru/weekly/2016/0683/analit01.php

Tracks of flu on the mortality surface: England and Wales vs. median mortality in other high income countries

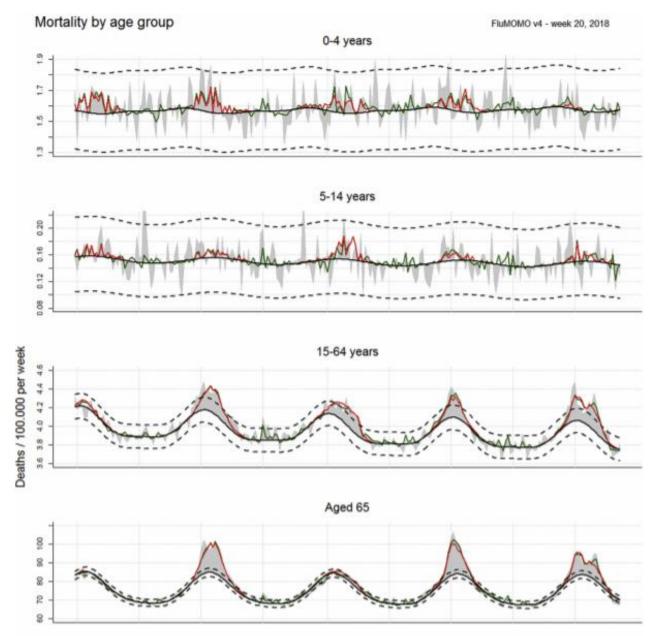


Leon, Shkolnikov, Jdanov 2019

https://pubmed.ncbi.nlm.nih.gov/31677776/

Figure 2 Mortality rate ratios in England and Wales relative to the median for 22 comparator countries by age, year, and sex

EuroMOMO project: an analysis of weekly mortality in 24 countries and within-country regions of Europe



Nielsen et al. 2019

http://www.euromomo.eu/methods/pdf/european excess

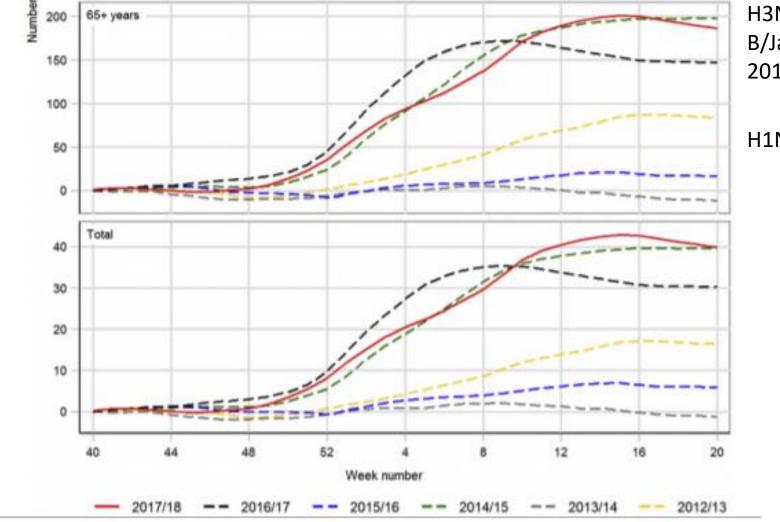
mortality 2017 18.pdf

Danilova, Shkolnikov, Jdanov. HSE Webinar: COVID-19: Quantification



Deadly influenza epidemics of the 2010s





H3N2: 2014-15, 2016-17 B/Jamagata&H3N2: 2017-18

H1N1: 2012-13, 2015-16

Nielsen et al. 2019

http://www.euromomo.eu/methods/pdf/european excess

mortality 2017 18.pdf



How it applies to time series of 2020

Trends in COVID-19 deaths in England and Wales: NHS vs. ONS



Office for National Statistics

Statistical bulletin

Deaths registered weekly in England and Wales, provisional: week ending 27 March 2020

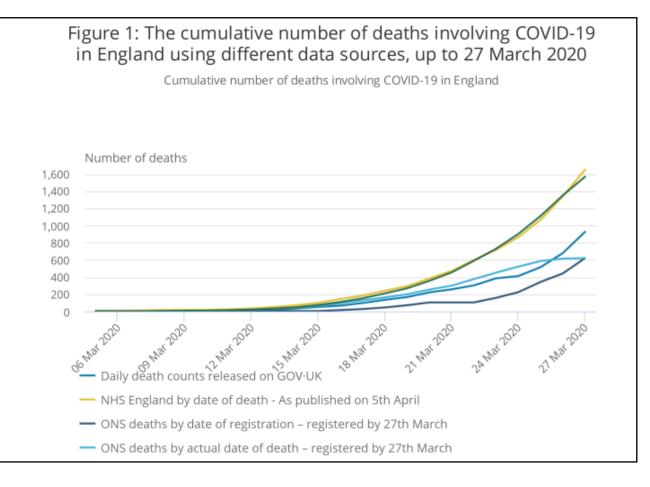
Provisional counts of the number of deaths registered in England and Wales, including deaths involving the coronavirus (COVID-19), by age, sex and region, in the latest weeks for which data are available.

Release date:

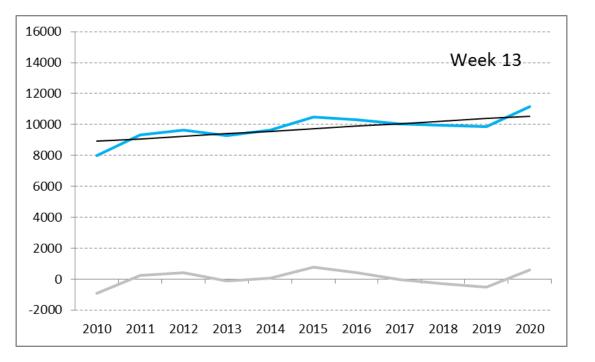
7 April 2020



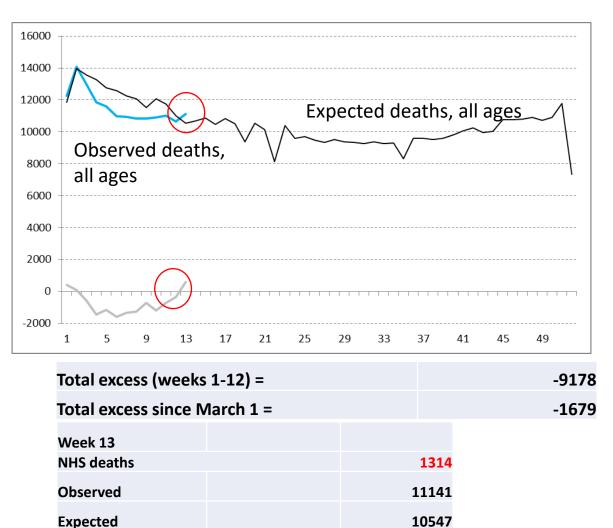
Contact: Sarah Caul health.data@ons.gov.uk +44 (0)1633 456 490 Next release: 14 April 2020



Assessment of weekly deaths on data from England and Wales 5 (for weeks 1 to 13 (March 27) (Method 1)



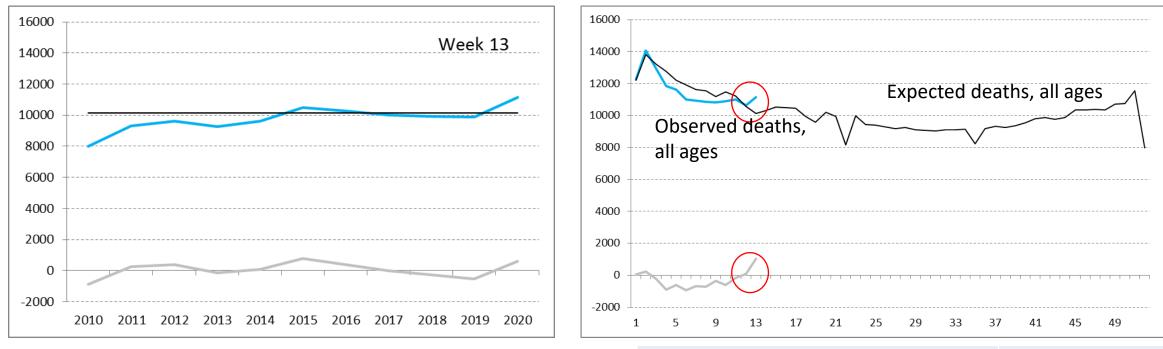
Expected Deaths 1 (Week N, age, 2020) = = Linear trend_extrapol (Week N, age, 2010-2019)



594

Observed-Expected

Assessment of weekly deaths on data from England and Wales for weeks 1 to 13 (March 27) (Method 2)

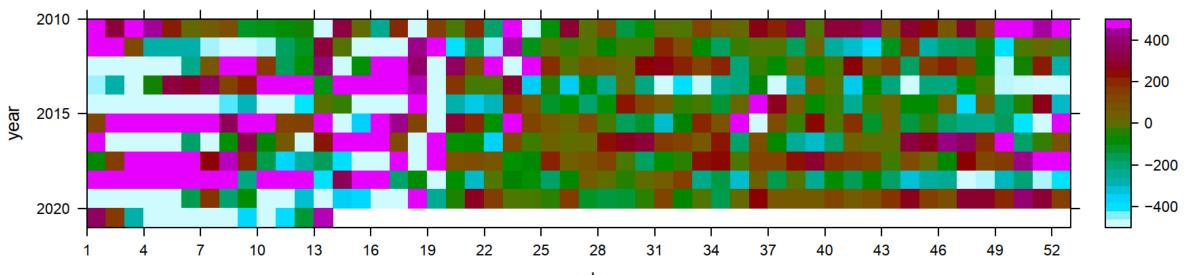


Expected Deaths 2 (Week N, age, 2020) = = Average (Week N, age, 2015-2019)

Total excess (weeks 1-12) =	-3806	
Total excess since March 1 =	-1069	
Week 13		
NHS deaths	1314	
Observed	11141	
Expected	10123	
Observed-Expected	1018	

Maps of excess mortality: England and Wales, weeks 1 to 13, all ages



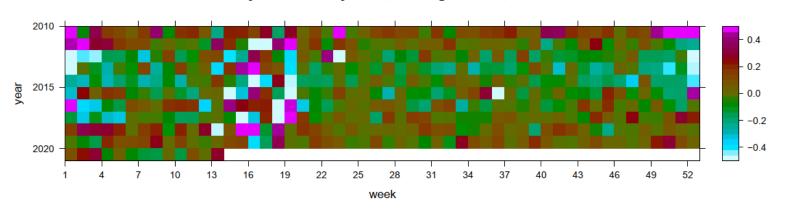


fluctuation in deaths (absolute numbers) by week, E&W, all ages

week

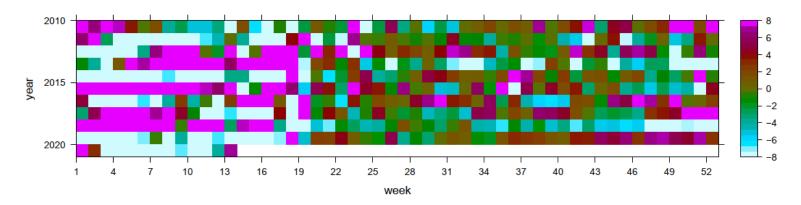
Maps of excess mortality: England and Wales, weeks 1 to 13, age groups 45-64 and 85+





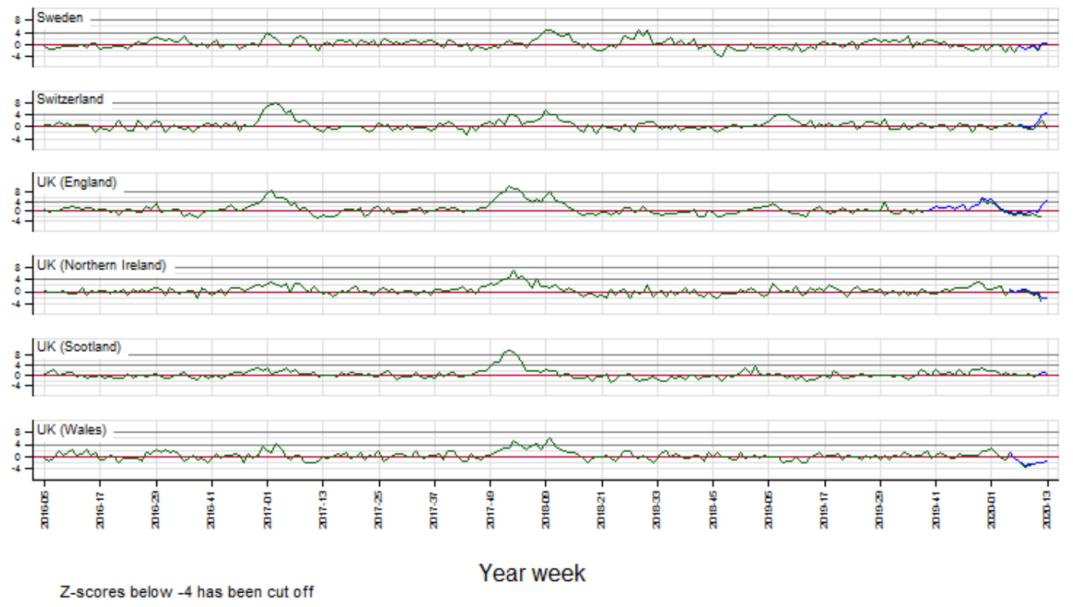
mortality fluctuation by week, E&W, ages 45-64, both sexes

mortality fluctuation by week, E&W, ages 85+, both sexes



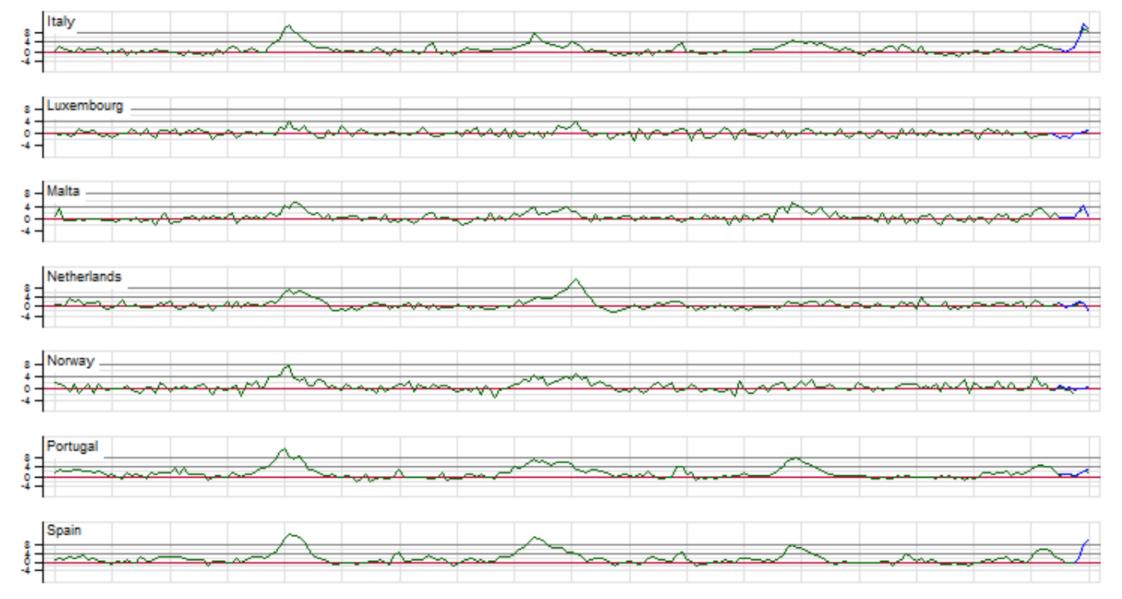


Weekly deaths by country by EuroMOMO (1)





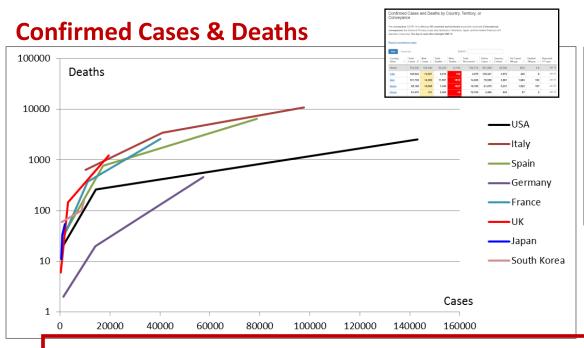
Weekly deaths by country by EuroMOMO (2)



Danilova, Shkolnikov, Jdanov. HSE Webinar: COVID-19: Quantification

Спасибо за внимание!

Other types of data to quantify the pandemic



Quality data from special sites: Diamond Princess cruise ship

Age Range	clFR	cCFR	Hospitalisation-to-death Distribution
All ages combined	0.91% (0.11% - 4.3%)	1.9% (0.60% - 4.3%)	Non-truncated (Figure 1A)
	1.2% (0.39% - 2.7%)	2.3% (0.75% - 5.3%)	Truncated (Figure 1B)
70+	7.3% (3.0% - 14%)	14% (6.0% - 27%)	Non-truncated (Figure 1A)
	9.0% (3.8% - 17%)	18% (7.3% - 33%)	Truncated (Figure 1B)

 Table 1: clFR and cCFR estimates calculated using the reported case and death data on the Diamond Princess cruise ship [2]. Correction

 was performed using equation (1) and the hospitalisation-to-death distribution in [6].

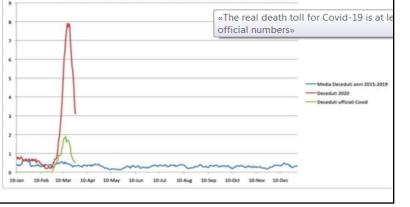
Media reports: Corriere Della Sera

«The real death toll for Covid-19 is at least 4 times the official numbers»

Nembro, one of the municipalities most affected by Covid-19, should have had - under normal conditions - about 35 deaths. 158 people were registered dead this year by the municipal offices. But the number of deaths officially attributed to Covid-19 is 31

di Claudio Cancelli Luca Foresti

Media Deceduti giornalieri a Nembro Media 2015-2019 Vs 2020 (Dati Comune di Nembro, Elaborazione Luca Foresti)



Weekly deaths: ONS and EuroMOMO

