

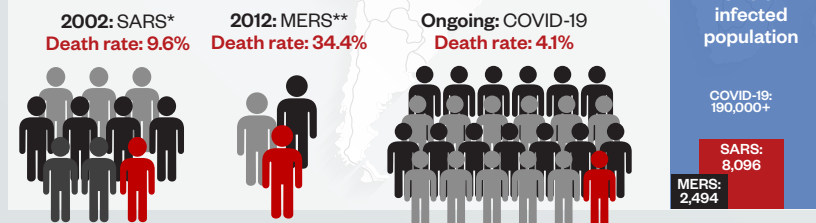
THE RACE TO STOP COVID-19

As a new strain of coronavirus threatens public health systems across the globe, researchers are racing to develop a vaccine.

DAWN CONNELLY & JULIA ROBINSON

In context

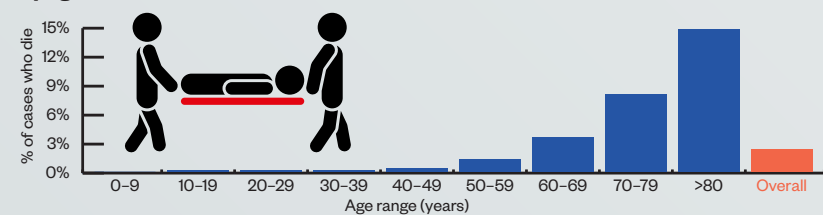
Compared with some outbreaks of novel respiratory disease, the death rate for coronavirus disease 2019 (COVID-19) is currently low. This could reduce further as the global outbreak progresses and milder cases are detected. Pandemic (H1N1) 2009 influenza was estimated to infect 1.6 billion people, with a death rate of 1 in 5,000 (0.02%); seasonal flu has a similar death rate, but infects up to 4 billion people each flu season.



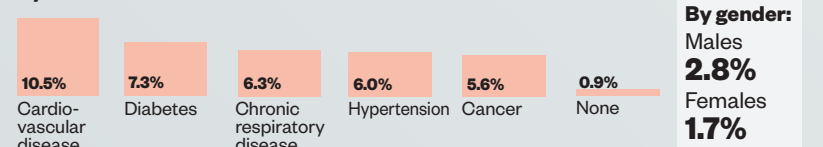
*Severe acute respiratory syndrome ** Middle East respiratory syndrome

Case fatality rates for COVID-19

By age:



By comorbid condition:



By gender:
Males 2.8%
Females 1.7%

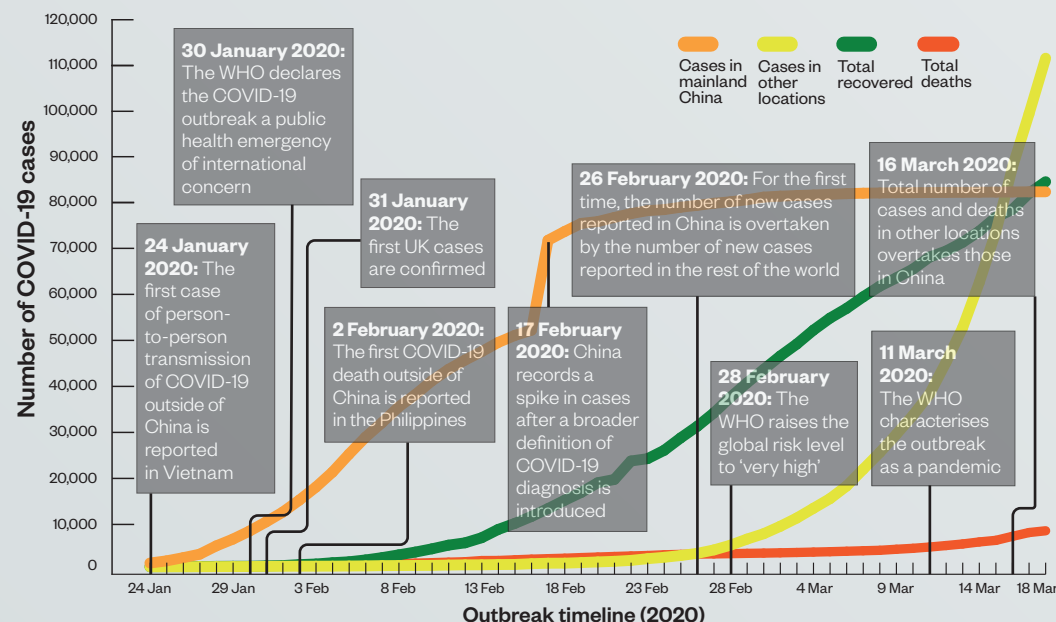
Symptoms and transmission

- Common symptoms*: **fever** (98.6%); **weakness** (69.6%); **cough** (59.4%); **muscle pains** (34.8%); **difficulties breathing** (31.2%);
- Less common symptoms*: **diarrhoea** (10.1%); **nausea and vomiting** (10.1%); **dizziness** (9.4%); **headaches** (6.5%); **stomach pain** (2.2%);
- Incubation period: 0-14 days;
- Severity: data from 44,000 cases of COVID-19 in China suggest that 80.9% of cases are mild, 13.8% are severe and 4.7% are critical. Severe cases are more prevalent in older people and those with an existing long-term condition, such as cardiovascular disease, diabetes, respiratory disease or hypertension;
- Transmission: person-to-person spread between close contacts (up to 1.8 metres) through respiratory droplets;
- Diagnosis: COVID-19 is suspected based on clinical symptoms. Cases are only confirmed when there is a positive laboratory test;
- Treatment: treatment of COVID-19 is symptomatic, such as providing oxygen. Medicines specifically licensed for COVID-19 are not currently available, but clinical trials of several antiviral drugs are being conducted in China.

*Percentages are taken from a case series of 138 consecutive hospitalised patients in China.

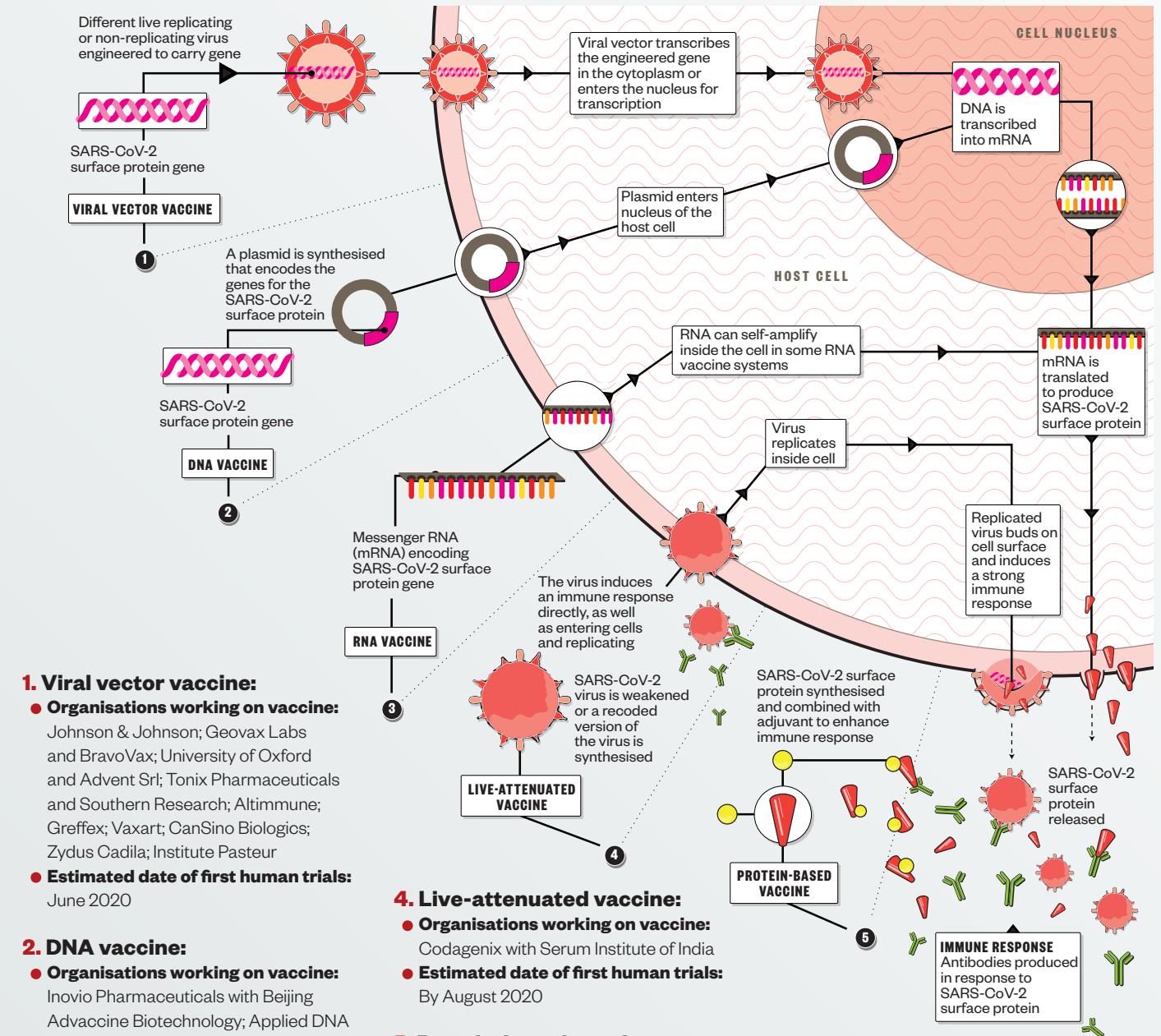
Global takeover

- 31 December 2019:** The first cases of pneumonia with unknown cause in Wuhan, China, are reported to the World Health Organization (WHO)
- 7 January 2020:** The Chinese authorities identify the novel coronavirus and temporarily name it '2019-nCoV', which is later changed to severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2)
- 10 January 2020:** The Chinese Center for Disease Control and Prevention makes the genome of SARS-CoV-2 available
- 11 January 2020:** The first COVID-19 death is reported in China
- 24 January 2020:** The first case of person-to-person transmission of COVID-19 outside of China is reported in Vietnam
- 2 February 2020:** The first COVID-19 death outside of China is reported in the Philippines
- 17 February 2020:** China records a spike in cases after a broader definition of COVID-19 diagnosis is introduced
- 28 February 2020:** The WHO raises the global risk level to 'very high'
- 11 March 2020:** The WHO characterises the outbreak as a pandemic
- 16 March 2020:** Total number of cases and deaths in other locations overtakes those in China
- 26 February 2020:** For the first time, the number of new cases reported in China is overtaken by the number of new cases reported in the rest of the world
- 31 January 2020:** The first UK cases are confirmed
- 30 January 2020:** The WHO declares the COVID-19 outbreak a public health emergency of international concern



Vaccine development

There are five main approaches being taken to develop a vaccine against SARS-CoV-2, with several biotechnology companies, academic organisations and pharmaceutical companies employing different technologies in the race to bring their vaccine candidate to clinical trials. The World Health Organization (WHO) hopes that a vaccine will be available by October 2021.



1. Viral vector vaccine:

- Organisations working on vaccine:** Johnson & Johnson; Geovax Labs and BravoVax; University of Oxford and Advent Srl; Tonix Pharmaceuticals and Southern Research; Altimune; Greffex; Vaxart; CanSino Biologics; Zydus Cadila; Institute Pasteur
- Estimated date of first human trials:** June 2020

2. DNA vaccine:

- Organisations working on vaccine:** Inovio Pharmaceuticals with Beijing Advaccine Biotechnology; Applied DNA Sciences, Takis Biotech and Ewvix; Zydus Cadila
- Estimated date of first human trials:** April 2020

3. RNA vaccine:

- Organisations working on vaccine:** CureVac; Moderna and US National Institute of Allergy and Infectious Diseases; Stermira Therapeutics, Tongji University and Chinese Center for Disease Control and Prevention; Imperial College London; Arcturus and Duke-NUS Medical School, Singapore; BioNTech and Pfizer
- Date of first human trials:** March 2020

4. Live-attenuated vaccine:

- Organisations working on vaccine:** Codagenix with Serum Institute of India
- Estimated date of first human trials:** By August 2020

5. Protein-based vaccine:

- Organisations working on vaccine:** Novavax; Clover Biopharmaceuticals with GSK; Baylor College of Medicine, University of Texas Medical Branch, New York Blood Center and Fundan University, China; University of Saskatchewan, Canada; University of Queensland, Australia, GSK and Dynavax; Vaxart; Genex; Expres2ion; Vaxil Bio; Sanofi Pasteur; iBio/CC-Pharming; Genex and EpiVax; Walter Reed Army Institute of Research and United States Army Medical Research Institute of Infectious Diseases; EpiVax and University of Georgia
- Estimated date of first human trials:** By June 2020

Immune response:

- It is not known how strong the immune response needs to be to protect against SARS-CoV-2; therefore, some of the vaccines being developed may not work.
- Before candidates reach clinical trials, investigators must also ensure they induce protective immunity, not immunopathology, as was seen in early attempts to develop a SARS-CoV vaccine after it emerged in 2002.

SOURCES: China CDC Weekly 2020;2(8):113-122; Department of Health and Social Care; JAMA 2020, doi:10.1001/jama.2020.1585; JAMA 2020, doi:10.1001/jama.2020.2648; John Hopkins Center for Systems Science and Engineering; Lancet 2012;379:687-696; World Health Organization
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All data correct as of 19 March 2020