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Infection fatality rate of COVID-19

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Infection fatality rate of COVID-19 inferred from seroprevalence data

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Abstract

Objective To estimate the infection fatality rate of coronavirus disease 2019 (COVID-19) from seroprevalence data.

Methods I searched PubMed and preprint servers for COVID-19 seroprevalence studies with a sample size ≥ 500 as of 9 September, 2020. I also retrieved additional results of national studies from preliminary press releases and reports. I assessed the studies for design features and seroprevalence estimates. I estimated the infection fatality rate for each study by dividing the number of COVID-19 deaths by the number of people estimated to be infected in each region. I corrected for the number of antibody types tested (immunoglobin, IgG, IgM, IgA).

Results I included 61 studies (74 estimates) and eight preliminary national estimates. Seroprevalence estimates ranged from 0.02% to 53.40%. Infection fatality rates ranged from 0.00% to 1.63%, corrected values from 0.00% to 1.54%. Across 51 locations, the median COVID-19 infection fatality rate was 0.27% (corrected 0.23%): the rate was 0.09% in locations with COVID-19 population mortality rates less than the global average (< 118 deaths/million), 0.20% in locations with 118–500 COVID-19 deaths/million people and 0.57% in locations with > 500 COVID-19 deaths/million people. In people < 70 years, infection fatality rates ranged from 0.00% to 0.31% with crude and corrected medians of 0.05%.

Conclusion The infection fatality rate of COVID-19 can vary substantially across different locations and this may reflect differences in population age structure and casemix of infected and deceased patients and other factors. The inferred infection fatality rates tended to be much lower than estimates made earlier in the pandemic.

Introduction

The infection fatality rate, the probability of dying for a person who is infected, is one of the most important features of the coronavirus disease 2019 (COVID-19) pandemic. The expected total mortality burden of COVID-19 is directly related to the infection fatality rate. Moreover,

justification for various non-pharmacological public health interventions depends on the infection fatality rate. Some stringent interventions that potentially also result in more noticeable collateral harms¹ may be considered appropriate, if the infection fatality rate is high. Conversely, the same measures may fall short of acceptable risk-benefit thresholds, if the infection fatality rate is low.

Early data from China suggested a 3.4% case fatality rate² and that asymptomatic infections were uncommon,³ thus the case fatality rate and infection fatality rate would be about the same. Mathematical models have suggested that 40–81% of the world population could be infected,^{4,5} and have lowered the infection fatality rate to 1.0% or 0.9%.^{5,6} Since March 2020, many studies have estimated the spread of the virus causing COVID-19 – severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) – in various locations by evaluating seroprevalence. I used the prevalence data from these studies to infer estimates of the COVID-19 infection fatality rate.

Methods

Seroprevalence studies

The input data for calculations of infection fatality rate were studies on the seroprevalence of COVID-19 done in the general population, or in samples that might approximately represent the general population (e.g. with proper reweighting), that had been published in peer-reviewed journals or as preprints (irrespective of language) as of 9 September 2020. I considered only studies with at least 500 assessed samples because smaller data sets would result in large uncertainty for any calculations based on these data. I included studies that made seroprevalence assessments at different time intervals if at least one time interval assessment had a sample size of at least 500 participants. If there were different eligible time intervals, I selected the one with the highest seroprevalence, since seroprevalence may decrease over time as antibody titres decrease. I excluded studies with data collected for more than a month that could not be broken into at least one eligible time interval less than one month duration because it would not be possible to estimate a point seroprevalence reliably. Studies were eligible regardless of the exact age range of participants included, but I excluded studies with only children.

I also examined results from national studies from preliminary press releases and reports whenever a country had no other data presented in published papers of preprints. This inclusion allowed these countries to be represented, but information was less complete than information in published papers or preprints and thus requires caution.

I included studies on blood donors, although they may underestimate seroprevalence and overestimate infection fatality rate because of the healthy volunteer effect. I excluded studies on health-care workers, since this group is at a potentially high exposure risk, which may result in seroprevalence estimates much higher than the general population and thus an improbably low infection fatality rate. Similarly, I also excluded studies on communities (e.g. shelters or religious or other shared-living communities). Studies were eligible regardless of whether they aimed to evaluate seroprevalence in large or small regions, provided that the population of reference in the region was at least 5000 people.

I searched PubMed® (LitCOVID), and medRxiv, bioRxiv and Research Square using the terms "seroprevalence" OR "antibodies" with continuous updates. I made the first search in early May and did monthly updates, with the last update on 9 September, 2020. I contacted field experts to retrieve any important studies that may have been missed.

From each study, I extracted information on location, recruitment and sampling strategy, dates of sample collection, sample size, types of antibody measured (immunoglobulin G (IgG), IgM and IgA), the estimated crude seroprevalence (positive samples divided by all samples tested), adjusted seroprevalence and the factors that the authors considered for adjustment.

Inferred infection fatality rate

If a study did not cover an entire country, I collected information on the population of the relevant location from the paper or recent census data so as to approximate as much as possible the relevant catchment area (e.g. region(s) or county(ies)). Some studies targeted specific age groups (e.g. excluding elderly people and/or excluding children) and some estimated numbers of people infected in the population based on specific age groups. For consistency, I used the entire population (all ages) and, separately, the population 0–70 years to estimate numbers of infected people. I assumed that the seroprevalence would be similar in different age groups, but I also recorded any significant differences in seroprevalence across age strata so as to examine the validity of this assumption.

I calculated the number of infected people by multiplying the relevant population size and the adjusted estimate of seroprevalence. If a study did not give an adjusted seroprevalence estimate, I used the unadjusted seroprevalence instead. When seroprevalence estimates with different adjustments were available, I selected the analysis with largest adjustment. The factors adjusted for included COVID-19 test performance, sampling design, and other factors such as age,

sex, clustering effects or socioeconomic factors. I did not adjust for specificity in test performance when positive antibody results were already validated by a different method.

For the number of COVID-19 deaths, I chose the number of deaths accumulated until the date 1 week after the midpoint of the study period (or the date closest to this that had available data) – unless the authors of the study had strong arguments to choose some other time point or approach. The 1-week lag accounts for different delays in developing antibodies versus dying from infection. The number of deaths is an approximation because it is not known when exactly each patient who died was infected. The 1-week cut-off after the study midpoint may underestimate deaths in places where patients are in hospital for a long time before death, and may overestimate deaths in places where patients die soon because of poor or even inappropriate care. Whether or not the health system became overloaded may also affect the number of deaths. Moreover, because of imperfect diagnostic documentation, COVID-19 deaths may have been both overcounted and undercounted in different locations and at different time points.

I calculated the inferred infection fatality rate by dividing the number of deaths by the number of infected people for the entire population, and separately for people < 70 years. I took the proportion of COVID-19 deaths that occurred in people < 70 years old from situational reports for the respective locations that I retrieved at the time I identified the seroprevalence studies. I also calculated a corrected infection fatality rate to try and account for the fact that only one or two types of antibodies (among IgG, IgM, IgA) might have been used. I corrected seroprevalence upwards (and inferred infection fatality rate downwards) by one tenth of its value if a study did not measure IgM and similarly if IgA was not measured. This correction is reasonable based on some early evidence, ⁷ although there is uncertainty about the exact correction factor.

Data synthesis

The estimates of the infection fatality rate across all locations showed great heterogeneity with I^2 exceeding 99.9%; thus, a meta-analysis would be inappropriate to report across all locations. Quantitative synthesis with meta-analysis across all locations would also be misleading since locations with high COVID-19 seroprevalence would tend to carry more weight than locations with low seroprevalence. Furthermore, locations with more studies (typically those that have attracted more attention because of high death tolls and thus high infection fatality rates) would be represented multiple times in the calculations. In addition, poorly conducted studies with fewer adjustments would get more weight because of spuriously narrower confidence intervals than

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more rigorous studies with more careful adjustments which allow for more uncertainty. Finally, with a highly skewed distribution of the infection fatality rate and with large between-study heterogeneity, typical random effects models would produce an incorrectly high summary infection fatality rate that approximates the mean of the study-specific estimates (also strongly influenced by high-mortality locations where more studies have been done); for such a skewed distribution, the median is more appropriate.

Therefore, in a first step, I grouped estimates of the infection fatality rate from studies in the same country (or for the United States of America, the same state) together and calculated a single infection fatality rate for that location, weighting the study-specific infection fatality rates by the sample size of each study. This approach avoided inappropriately giving more weight to studies with higher seroprevalence estimates and those with seemingly narrower confidence intervals because of poor or no adjustments, while still giving more weight to larger studies. Then, I used the single summary estimate for each location to calculate the median of the distribution of location-specific infection fatality rate estimates. Finally, I explored whether the location-specific infection fatality rates were associated with the COVID-19 mortality rate in the population (COVID-19 deaths per million people) in each location as of 12 September 2020; this analysis allowed me to assess whether estimates of the infection fatality rate tended to be higher in locations with a higher burden of death from COVID-19.

Results

Seroprevalence studies

I retrieved 61 studies with 74 eligible estimates published either in the peer-reviewed literature or as preprints as of 9 September 2020.8-68 Furthermore, I also considered another eight preliminary national estimates. ^{69–76} This search yielded a total of 82 eligible estimates (Fig. 1).

The studies varied substantially in sampling and recruitment designs (Table 1; available at: http://www.who.int/bulletin/volumes/##/##-#####). Of the 61 studies, 24 studies^{8,10,16,17,20,22,25,33,34,36,37,42,46–49,52–54,61,63,65,68} explicitly aimed for random sampling from the general population. In principle, random sampling is a stronger design. However, even then, people who cannot be reached (e.g. by email or telephone or even by visiting them at a house location) will not be recruited, and these vulnerable populations are likely to be missed. Moreover, several such studies^{8,10,16,37,42} focused on geographical locations with high numbers of deaths,

Article ID: BLT.20.265892

higher than other locations in the same city or country, and this emphasis would tend to select eventually for a higher infection fatality rate on average.

Eleven studies assessed blood donors, ^{12,15,18,24,28,31,41,44,45,55,60} which might underestimate COVID-19 seroprevalence in the general population. For example, 200 blood donors in Oise, France showed 3.00% seroprevalence, while the seroprevalence was 25.87% (171/661) in pupils, siblings, parents, teachers and staff at a high school with a cluster of cases in the same area; the true population seroprevalence may be between these two values. 13

For other studies, healthy volunteer bias 19 may underestimate seroprevalence, attracting people with symptoms²⁶ may overestimate seroprevalence, and studies of employees, ^{14,21,25,32,66} grocery store clients²³ or patient cohorts^{11,14,27–30,36,38,40,50,51,56,59,62,64,67} risk sampling bias in an unpredictable direction.

All the studies tested for IgG antibodies but only about half also assessed IgM and few assessed IgA. Only seven studies assessed all three types of antibodies and/or used pan-Ig antibodies. The ratio of people sampled versus the total population of the region was more than 1:1000 in 20 studies (Table 2; available at: http://www.who.int/bulletin/volumes/##/##/##- ######).

Seroprevalence estimates

Seroprevalence for the infection ranged from 0.02% to 53.40% (58.40% in the slum subpopulation in Mumbai; Table 3). Studies varied considerably depending on whether or not they tried to adjust their estimates for test performance, sampling (to get closer to a more representative sample), clustering (e.g. when including household members) and other factors. The adjusted seroprevalence occasionally differed substantially from the unadjusted value. In studies that used samples from multiple locations, between-location heterogeneity was seen (e.g. 0.00–25.00%) across 133 Brazilian cities).²⁵

Inferred infection fatality rate

Inferred infection fatality rate estimates varied from 0.00% to 1.63% (Table 4). Corrected values also varied considerably (0.00–1.54%).

For 15 locations, more than one estimate of the infection fatality rate was available and thus I could compare the infection fatality rate from different studies evaluating the same location. The estimates of infection fatality rate tended to be more homogeneous within each location, while

Article ID: BLT.20.265892

they differed markedly across locations (Fig. 2). Within the same location, infection fatality rate estimates tend to have only small differences, even though it is possible that different areas within the same location may also have real differences in infection fatality rate. France is one exception where differences are large, but both estimates come from population studies of outbreaks from schools and thus may not provide good estimates of population seroprevalence and may lead to an underestimated infection fatality rate.

I used summary estimates weighted for sample size to generate a single estimate for each location. Data were available for 51 different locations (including the inferred infection fatality rates from the eight preliminary additional national estimates in Table 5).

The median infection fatality rate across all 51 locations was 0.27% (corrected 0.23%). Most data came from locations with high death tolls from COVID-19 and 32 of the locations had a population mortality rate (COVID-19 deaths per million population) higher than the global average (118 deaths from COVID-19 per million as of 12 September 2020;⁷⁹ Fig. 3). Uncorrected estimates of the infection fatality rate of COVID-19 ranged from 0.01% to 0.67% (median 0.10%) across the 19 locations with a population mortality rate for COVID-19 lower than the global average, from 0.07% to 0.73% (median 0.20%) across 17 locations with population mortality rate higher than the global average but lower than 500 COVID-19 deaths per million, and from 0.20% to 1.63% (median 0.71%) across 15 locations with more than 500 COVID-19 deaths per million. The corrected estimates of the median infection fatality rate were 0.09%, 0.20% and 0.57%, respectively, for the three location groups.

For people < 70 years old, the infection fatality rate of COVId-19 across 40 locations with available data ranged from 0.00% to 0.31% (median 0.05%); the corrected values were similar.

Discussion

The infection fatality rate is not a fixed physical constant and it can vary substantially across locations, depending on the population structure, the case-mix of infected and deceased individuals and other, local factors. The studies analysed here represent 82 different estimates of the infection fatality rate of COVID-19, but they are not fully representative of all countries and locations around the world. Most of the studies are from locations with overall COVID-19 mortality rates that are higher than the global average. The inferred median infection fatality rate in locations with a COVID-19 mortality rate lower than the global average is low (0.09%). If one

could sample equally from all locations globally, the median infection fatality rate might be even substantially lower than the 0.23% observed in my analysis.

COVID-19 has a very steep age gradient for risk of death. ⁸⁰ Moreover, many, and in some cases most, deaths in European countries that have had large numbers of cases and deaths ⁸¹ and in the USA ⁸² occurred in nursing homes. Locations with many nursing home deaths may have high estimates of the infection fatality rate, but the infection fatality rate would still be low among non-elderly, non-debilitated people.

Within China, the much higher infection fatality rate estimates in Wuhan compared with other areas of the country may reflect widespread nosocomial infections, ⁸³ as well as unfamiliarity with how to manage the infection as the first location that had to deal with COVID-19. The very many deaths in nursing homes, nosocomial infections and overwhelmed hospitals may also explain the high number of fatalities in specific locations in Italy⁸⁴ and New York and neighbouring states. ^{23,27,35,56} Poor decisions (e.g. sending COVID-19 patients to nursing homes), poor management (e.g. unnecessary mechanical ventilation) and hydroxychloroquine may also have contributed to worse outcomes. High levels of congestion (e.g. in busy public transport systems) may also have exposed many people to high infectious loads and, thus, perhaps more severe disease. A more aggressive viral clade has also been speculated. ⁸⁵ The infection fatality rate may be very high among disadvantaged populations and settings with a combination of factors predisposing to higher fatalities. ³⁷

Very low infection fatality rates seem common in Asian countries. 8,11,29,48,49,51,59,61,67 A younger population in these countries (excluding Japan), previous immunity from exposure to other coronaviruses, genetic differences, hygiene etiquette, lower infectious load and other unknown factors may explain these low rates. The infection fatality rate is low also in low-income countries in both Asia and Africa, 44,49,66,67 perhaps reflecting the young age-structure. However, comorbidities, poverty, frailty (e.g. malnutrition) and congested urban living circumstances may have an adverse effect on risk and thus increase infection fatality rate.

Antibody titres may decline with time ^{10,28,32,86,87} and this would give falsely low prevalence estimates. I considered the maximum seroprevalence estimate when multiple repeated measurements at different time points were available, but even then some of this decline cannot be fully accounted for. With four exceptions, ^{10,28,32,51} the maximum seroprevalence value was at the latest time point.

Positive controls for the antibody assays used were typically symptomatic patients with positive polymerase chain reaction tests. Symptomatic patients may be more likely to develop antibodies. 87–91 Since seroprevalence studies specifically try to reveal undiagnosed asymptomatic and mildly symptomatic infections, a lower sensitivity for these mild infections could lead to substantial underestimates of the number of infected people and overestimate of the inferred infection fatality rate.

A main issue with seroprevalence studies is whether they offer a representative picture of the population in the assessed region. A generic problem is that vulnerable people at high risk of infection and/or death may be more difficult to recruit in survey-type studies. COVID-19 infection is particularly widespread and/or lethal in nursing homes, in homeless people, in prisons and in disadvantaged minorities. 92 Most of these populations are very difficult, or even impossible, to reach and sample and they are probably under-represented to various degrees (or even entirely missed) in surveys. This sampling obstacle would result in underestimating the seroprevalence and overestimating infection fatality rate.

In principle, adjusted seroprevalence values may be closer to the true estimate, but the adjustments show that each study alone may have unavoidable uncertainty and fluctuation, depending on the type of analysis chosen. Furthermore, my corrected infection fatality rate estimates try to account for undercounting of infected people when not all three antibodies (IgG, IgM and IgA) were assessed. However, the magnitude of the correction is uncertain and may vary in different circumstances. An unknown proportion of people may have responded to the virus using immune mechanisms (mucosal, innate, cellular) without generating any serum antibodies. 93–97

A limitation of this analysis is that several studies included have not yet been fully peer-reviewed and some are still ongoing. Moreover, despite efforts made by seroprevalence studies to generate estimates applicable to the general population, representativeness is difficult to ensure, even for the most rigorous studies and despite adjustments made. Estimating a single infection fatality rate value for a whole country or state can be misleading, when there is often huge variation in the population mixing patterns and pockets of high or low mortality. Furthermore, many studies have evaluated people within restricted age ranges, and the age groups that are not included may differ in seroprevalence. Statistically significant, modest differences in seroprevalence across some age groups have been observed in several studies. 10,13,15,23,27,36,38

Lower values have been seen in young children and higher values in adolescents and young adults, but these patterns are inconsistent and not strong enough to suggest major differences extrapolating across age groups.

Acknowledging these limitations, based on the currently available data, one may project that over half a billion people have been infected as of 12 September, 2020, far more than the approximately 29 million documented laboratory-confirmed cases. Most locations probably have an infection fatality rate less than 0.20% and with appropriate, precise non-pharmacological measures that selectively try to protect high-risk vulnerable populations and settings, the infection fatality rate may be brought even lower.

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Competing interests:

I am a co-author (not principal investigator) of one of the seroprevalence studies.

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Table 1. Eligible seroprevalence studies on COVID-19 published or deposited as preprints as of 9 September 2020: dates, sampling and recruitment

| Author | Country (location) | Dates | Sampling and recruitment |
|--------------------------------------|------------------------------------|--|---|
| Figar et al. ⁴⁷ | Argentina (Barrio Padre Mugica) | 10–26 June | Probabilistic sampling of a slum neighbourhood, sampling from people 14 years or older across households |
| Herzog et al. ³⁸ | Belgium | 30 March–5 April and 20–26 April | Residual sera from 10 private diagnostic laboratories in Belgium, with fixed numbers per age group, region and periodical sampling, and stratified by sex |
| Hallal et al. ²⁵ | Brazil | 15–22 May | Sampling from 133 cities (the main city in each region), selecting 25 census tracts with probability proportionate to size in each sentinel city, and 10 households at random in each tract. Aiming for 250 participants per city |
| Gomes et al.34 | Brazil (Espirito Santo) | 13–15 May | Cross-section of major municipalities with houses as the sampling units |
| Da Silva et al. ⁶⁸ | Brazil (Maranhao) | 27 July–8 August | Three-stage cluster sampling stratified by four state regions in the state of Maranhao; the estimates took clustering, stratification and non-response into account |
| Amorim Filho et al. ⁴¹ | Brazil (Rio de Janeiro) | 14–27 April (eligible: 24–27 April) | Blood donors without flulike symptoms within 30 days of donation; had close contact with suspected or confirmed COVID-19 cases in the 30 days before donation; or had travelled abroad in the past 30 days |
| Silveira et al. ¹⁷ | Brazil (Rio Grande do Sul) | 9–11 May (third round, after 11–13 April, and 25–27 April) | Multistage probability sampling in each of nine cities to select 500 households, from which one member was randomly chosen for testing |
| Tess et al. ⁴² | Brazil (Sao Paulo) | 4–12 May | Randomly selected adults and their cohabitants sampled from six districts of Sao Paulo City with high numbers of cases |
| Skowronski et al. ⁵⁰ | Canada (British Columbia) | 15–27 May (after baseline in 5–13 March) | Specimens from patients attending one of about 80 diagnostic service centres of the only outpatient laboratory network in the Lower Mainland |
| Torres et al.43 | Chile (Vitacura) | 4–19 May | Classroom stratified sample of children and all staff in a community placed on quarantine after school outbreak |
| Chang et al. ⁵⁵ | China | January–April weekly: 3–23 February (Wuhan); 24 February–15 March (Shenzhen); 10 February–1 March (Shijiazhuang) | 38 144 healthy blood donors in Wuhan, Shenzhen and Shijiazhuang who met the criteria for blood donation during the COVID-19 pandemic in China |
| Wu et al. ¹⁴ | China (Wuhan) | 3–15 April | People applying for a permission to resume work ($n = 1021$) and hospitalized patients ($n = 381$) |
| Ling et al. ³² | China (Wuhan) | 26 March–28 April | Age 16–64 years, going back to work, with no fever, headache or other symptoms of COVID-19 |
| Xu et al. ⁶⁰ | China (Guangzhou) | 23 March-2 April | Healthy blood donors in Guangzhou |

| | | Arue | cie iD: BL1.20.203892 |
|--------------------------------|------------------------------|--------------------------|---|
| Xu et al. ⁴⁰ | China (several regions) | 30 March–10 April | Voluntary participation by public call for haemodialysis patients ($n = 979$ in Zingzhou, Ubei and $n = 563$ in Guangzhou/Foshun, Guangdong) and outpatients in Chingqing ($n = 993$), and community residents in Chengdu, Sichuan ($n = 9442$), and |
| 1 | | 00.00.4 | required testing for factory workers in Guangzhou, Guandong (<i>n</i> = 442) |
| Jerkovic et al. ²⁶ | Croatia | 23–28 April | DIV Group factory workers in Split and Sibenik-Knin invited for voluntary testing |
| Erikstrup et al. ¹² | Denmark | 6 April–3 May | All Danish blood donors aged 17–69 years giving blood. Blood donors are healthy and must comply with strict eligibility criteria; they must self-defer for two weeks if they develop fever with upper respiratory symptoms |
| Determine at al 52 | Donmark (Fares | 27 April 1 May | |
| Petersen et al. ⁵² | Denmark (Faroe Islands) | 27 April–1 May | 1500 randomly selected residents invited to participate, samples collected from 1 075 |
| Fontanet et al. ³⁹ | France (Crepy-en- Valois) | 28–30 April | Pupils, their parents and relatives, and staff of primary schools exposed to SARS-CoV-2 in February and March 2020 in a city north of Paris |
| Fontanet et al. ¹³ | France (Oise) | 30 March–4 April | Pupils, their parents and siblings, as well as teachers and non-teaching staff of a high-school |
| Streeck et al. ¹⁶ | Germany (Gangelt) | 30 March-6 April | 600 adults with different surnames in Gangelt were randomly selected; all |
| | , , | · | household members were asked to participate in the study |
| Kraehling et al. ²¹ | Germany (Frankfurt) | 6–14 April | Employees of Infraserv Höchst, a large industrial site operator in Frankfurt am Main. No exclusion criteria |
| Bogogiannidou et | Greece | March and April (April | Leftover blood samples collected from a nationwide laboratory network, including |
| al. ⁶² | 0.000 | data used) | both private and public hospital laboratories (27 laboratories in total) |
| Merkely et al. ⁵⁷ | Hungary | 1–16 May | Representative sample ($n = 17.787$) of the Hungarian population ≥ 14 years living in |
| Montory of all | riangary | . To May | private households (8 283 810) |
| Gudbjatsson et | Iceland | Several cohorts | 30 576 people in Iceland, including those documented to be infected, those |
| al. ⁵⁸ | lociand | between April and | quarantined and people not known to have been exposed. |
| aı. | | June ^a | quarantined and people not known to have been exposed. |
| Malani et al. ⁶¹ | India (Mumbai) | 29 June–19 July | Geographically-spaced community sampling of households, one individual per |
| Maiaili Ct al. | maia (Mambai) | 25 Julie 15 July | household was tested in slum and non-slum communities in three wards, one each |
| | | | from the three main zones of Mumbai |
| Khan et al. ⁶⁷ | India (Srinagar) | 1–15 July | Adults (> 18 years) who visited selected hospitals across the Srinagar District |
| Shakiba et al.8 | Islamic Republic of | April (until 21 April) | Population-based cluster random sampling design through telephone call invitation, |
| Orianiba et al. | Iran (Guilan) | April (dritti 2 i April) | household-based |
| Fiore et al. ³¹ | Italy (Apulia) | 1–31 May | Blood donors 18–65 years old free of recent symptoms possibly related to COVID- |
| i lore et al. | italy (Apulla) | 1-51 Way | 19, no close contact with confirmed cases, symptom-free in the preceding 14 days, |
| | | | no contact with suspected cases |
| Doi et al. ¹¹ | Japan (Kobe) | 31 March–7 April | Randomly selected patients who visited outpatient clinics and received blood testing |
| Doi et al. | Japan (Robe) | 31 March April | for any reason. Patients who visited the emergency department or the designated |
| | | | fever consultation service were excluded |
| Takita et al. ²⁹ | Japan (Takya) | 21 April 20 May | |
| i akila el al.ºº | Japan (Tokyo) | 21 April–20 May | Two community clinics in the main railway stations in Tokyo (Navitas Clinic Shinjuku |
| | | | and Tachikawa) |

| | | Artic | ele ID: BLT.20.265892 |
|--|--|---|---|
| Nawa et al. ⁴⁸ | Japan (Utsunomiya City) | 14 June–5 July | Invitations enclosed with a questionnaire were sent to 2 290 people in 1 000 households randomly selected from Utsunomiya City's basic resident registry; 742 completed the study |
| Uyoga et al. ⁴⁴ | Kenya | 30 April–16 June (~90% of samples in last 30 days) | Residual blood donor serum samples from donors 16–65 years in four sites (Mombasa, Nairobi, Eldoret and Kisumu) |
| Snoeck et al. ²⁰ | Luxembourg | 16 April–5 May | Representative sample (no details how ensured), 1 807 of 2000 contacted provided data, were < 79 years and had serology results |
| Slot et al. ¹⁵ | Netherlands | 1–15 April | Blood donors. Donors must be completely healthy, but they may have been ill in the past, provided that they recovered at least 2 weeks before |
| Westerhuis et al. ⁶⁴ Nisar et al. ⁴⁹ | Netherlands (Rotterdam) Pakistan (Karachi) | Early March and early April 25 June–11 July (after baseline on 15–25 April) | Left-over plasma samples from patients of nine age categories in Erasmus Medical Center in Rotterdam: 879 samples in early March and 729 in early April) Cross-sectional household surveys in a low- (district Malir) and high-transmission (district East) area of Karachi with households selected using simple random sampling (Malir) and systematic random sampling (East) |
| Javed et al. ⁶⁶ | Pakistan (urban Karachi, Lahore, Multan, Peshawar and Quetta) | 06-Jul | Adult, working population aged 18–65 years, recruited from dense, urban workplaces including factories, businesses, restaurants, media houses, schools, banks, hospitals (health-care providers), and from families of positive cases in cities in Pakistan |
| Abu Raddad et al. ⁵¹ | Qatar | 12 May–12 July (highest seroprevalence on 12– 31 May) | Convenience sample of residual blood specimens collected for routine clinical screening or clinical management from 32 970 outpatient and inpatient departments for a variety of health conditions ($n = 937$ in 12–31 May) |
| Noh et al. ⁵⁹ | Republic of Korea | 25–29 May | Outpatients who visited two hospitals in south-west Seoul which serve six administrative areas |
| Pollan et al. ³⁶ | Spain | 27 April–11 May | 35 883 households selected from municipal rolls using two-stage random sampling stratified by province and municipality size, with all residents invited to participate (75.1% of all contacted individuals participated) |
| Crovetto et al. ³⁰ | Spain (Barcelona) | 14 April–5 May | Consecutive pregnant women for first trimester screening or delivery in two hospitals |
| Stringhini et al. ¹⁰ | Switzerland (Geneva) | 6 April–9 May (5 consecutive weeks) | Randomly selected previous participants of the Bus Santé study with an email (or telephone contact, if email unavailable); participants were invited to bring all members of their household aged 5 years and older |
| Emmenegger et al. ²⁸ | Switzerland (Zurich) | Prepandemic until June (patients) and May (blood donors) | Patients at the University Hospital of Zurich and blood donors in Zurich and Lucerne |
| Ward et al. ⁶⁵ | United Kingdom (England) | 20 June-13 July | Random population sample of 100 000 adults over 18 years |
| Thompson et al. ¹⁸ | United Kingdom (Scotland) | 21–23 March | Blood donors. Donors should not have felt unwell in the past 14 days; some other deferrals also applied regarding travel and COVID-19 symptoms |

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|------------------------------------|---|--|---|
| Havers et al. ³⁵ | USA (10 states) | 23 March–1 April (Washington, Puget Sound and New York, New York City), 1–8 April (Louisiana), 5–10 April (Florida, south), 13–25 April (Pennsylvania, Philadelphia, metropolitan area), 20–26 April (Missouri), 23–27 April (California, San Francisco Bay Area), 20 April–3 May (Utah), 26 April–3 May (Connecticut), 30 April–12 May (Minnesota, Minneapolis) | Convenience samples using residual sera obtained for routine clinical testing (screening or management) by two commercial laboratory companies |
| Ng et al. ²⁴ | USA (California, Bay | March | 1000 blood donors in diverse Bay Area locations (excluding those with self-reported |
| Sood ²² | Area) USA (California, Los Angeles) | 10–14 April | symptoms or abnormal vital signs) Proprietary database representative of the county. A random sample of these residents was invited, with quotas for enrolment for subgroups based on age, sex, race and ethnicity distribution |
| Chamie et al. ³³ | USA (California, San Francisco) | 25–28 April | United States census tract 022 901 population-dense area (58% Latin American) in San Francisco Mission district, expanded to neighbouring blocks on 28 April |
| Bendavid et al. ¹⁹ | USA (California, Santa Clara) | 2–3 April | Facebook advertisement with additional targeting by zip code |
| Biggs et al. ⁵³ | USA (Georgia, DeKalb and Fulton) | 28 April–3 May | Two-stage cluster sampling design used to randomly select 30 census blocks in DeKalb county and 30 census blocks in Fulton county, with a target of seven participating households per census block |
| McLaughlin et al. ⁴⁶ | USA (Idaho, Blaine county) | 4–19 May | Volunteers who registered via a secure web link, using prestratification weighting to the population distribution by age and sex within each zip code |
| Bryan et al. ⁹ | USA (Ídaho, Boise) | Late April | People from the Boise, Idaho metropolitan area, part of the Crush the Curve initiative |
| Menachemi et al. ⁵⁴ | USA (Indiana) | 25–29 April | Stratified random sampling among all persons aged ≥ 12 years using Indiana's 10 public health preparedness districts as sampling strata |
| Feehan et al. ⁶³ | USA (Louisiana, Baton Rouge) | 15–31 July | Representative sample in a method developed by Public Democracy |

Article ID: BLT.20.265892

| Feehan et al. ³⁷ | USA (Louisiana, Orleans and Jefferson Parish) | 9–15 May | Pool of potential participants reflecting the demographics of the parishes was based on 50 characteristics, then a randomized subset of 150 000 people was selected, and 25 000 were approached with digital apps, and 2 640 were recruited |
|--------------------------------|---|---|--|
| Rosenberg et al. ²³ | USA (New York) | 19–28 April | Convenience sample of people ≥ 18 years living in New York State, recruited consecutively on entering 99 grocery stores and through an in-store flyer |
| Meyers et al. ⁵⁶ | USA (New York) | 2–30 March (Columbia University Medical Center, New York City); 13–28 March (CareMount central laboratory) | Discarded clinical samples in Columbia Medical Center, New York City (<i>n</i> = 814 in 24 February–30 March, 742 of those in the period 2–30 March) and samples from CareMount central laboratory (960 samples on 13 and 14 March, 505 samples on 20/21 March, and 376 samples on 27/28 March) from its network of clinics in five counties north of New York City |
| Reifer et al. ²⁷ | USA (New York, Brooklyn) | Early May | Patients seen in an urgent care facility in Brooklyn |
| Nesbitt et al.45 | USA (Rhode Island) | 27 April–11 May | Consecutive blood donors |

COVID-19: coronavirus disease-19; SARS-CoV-2: severe acute respiratory syndrome coronavirus 2.

Note: Some studies included additional data sets that did not fulfil the eligibility criteria (e.g. had sample size < 500 or were health-care workers) and they are not presented here.

^a Sample collection time for some sub-cohorts may have exceeded 1 month, but more than half of the cases were already documented by polymerase chain reaction testing before any antibody testing and the last death occurred on 20 April.

Table 2. Sample size, types of antibodies assessed and population size in the studies included to assess COVID-19 infection fatality rate, 2020

| studies included | to assess COVID | 19 intection tat | ality rate, 2020 | |
|---|----------------------------------|--------------------------|---------------------------------------|---|
| Country (location) | Sample size ^a , no. | Antibody | Population, ^b no. | % of population < 70 years ^c |
| Argentina (Barrio Padre | 873 | IgG | 49 983 | 99 |
| Mugica) ⁴⁷ | | | | |
| Belgium ³⁸ | 3 391 (20–26 April) | IgG | 11 589 623 | 86 |
| Brazil (133 cities) ²⁵ | 24 995 | IgG and IgM | 74 656 499 | 94 (Brazil) |
| Brazil (Espirito Santo) ³⁴ | 4 608 | lgG and lgM | 4 018 650 | 94 (Brazil) |
| Brazil (Maranhao) ⁶⁸ | 3 156 | lgG and lgM | 7 114 598 | 92 |
| Brazil (Rio de Janeiro), blood | 669 (24–27 April) | IgG and IgM | 17 264 943 | 94 (Brazil) |
| donors ⁴¹ | 4.500 | | 44.077.000 | 0.4 |
| Brazil (Rio Grande do Sul) ¹⁷ | 4 500 | lgG | 11 377 239 | 91 |
| Brazil (Sao Paulo) ⁴² | 517 | IgG and IgM | 298 240 (6 districts) | 94 (Brazil) |
| Canada (British Columbia) ⁵⁰ | 885 | IgG, IgM and IgA | 5 071 000 | 94 |
| Chile (Vitacura) ⁴³ | 1 244 | IgG and IgM | 85 000 | 92 (Chile) |
| China, blood donors ⁵⁵ | 020 (2. 22 Fabruary) | laC and laM | 11 210 000 | 02 (China) |
| Wuhan | 930 (3–23 February) | IgG and IgM | 11 210 000 | 93 (China) |
| Shenzhen | 3 507 (24 February– 15 March) | lgG and lgM | 13 030 000 | 93 (China) |
| Shijiazhuang | 6 455 (10 February–1 March) | lgG and lgM | 11 030 000 | 93 (China) |
| China (Wuhan) ¹⁴ | 1 401 | IgG and IgM | 11 080 000 | 93 (China) |
| China (Wuhan) ³² | 1 196 (4–8 April) | IgG and IgM | 11 080 000 | 93 (China) |
| China (Guangzhou), blood | 2 199 | lgG, lgM and lgA | 115 210 000 (Guangdong) | 93 (China) |
| donors ⁶⁰ | | | | |
| China (several regions) ⁴⁰ | | | | |
| Hubei (not Wuhan) | 979 | lgG and lgM | 48 058 000 | 93 (China) |
| Chongqing | 993 | lgG and lgM | 31 243 200 | 93 (China) |
| Sichuan | 9 442 | lgG and lgM | 83 750 000 | 93 (China) |
| Guangdong | 1 005 | IgG and IgM | 115 210 000 | 93 (China) |
| Croatia ²⁶ | 1 494 | IgG and IgM | 4 076 000 | 86 |
| Denmark blood donors ¹² | 20 640 1 075 | lgG and lgM | 5771876 | 86 |
| Denmark (Faroe Islands) ⁵² | 1 340 | lgG and lgM | 52 428 5 978 000 (Hauts-de-France) | 88 89 |
| France (Crepy-en-Valois) ³⁹ France (Oise) ¹³ | 661 | lgG lgG | 5 978 000 (Hauts-de-France) | 89 |
| Germany (Gangelt) ¹⁶ | 919 | lgG and lgA | 12 597 | 86 |
| Germany (Frankfurt) ²¹ | 1 000 | IgG and IgA | 2 681 000 ^d | 84 (Germany) |
| Greece ⁶² | 6 586 (4 511 in April) | lgG | 10 412 967 | 84 |
| Hungary ⁵⁷ | 10 504 | lgG (also had PCR) | 9657451 | 88 |
| Iceland ⁵⁸ | 30 576 | Pan-lg | 366 854 | 90 |
| India (Mumbai) ⁶¹ | 6 904 (4 202 in | IgG | 1 414 917 (705 523 in slums, | 98 |
| | slums, 2 702 not in | .90 | 709 394 in non-slums) in the 3 | |
| | slums) | | ward areas | |
| India (Srinagar) ⁶⁷ | 2 906 | IgG | 1 500 000 | 97 |
| Islamic Republic of Iran (Guilan) ⁸ | 551 | lgG and lgM | 2354848 | 95 |
| Italy (Apulia), blood donors ³¹ | 909 | IgG and /IgM | 4 029 000 | 84 |
| Japan (Kobe) ¹¹ | 1 000 | IgG | 1 518 870 | 79 (Japan) |
| Japan (Tokyo) ²⁹ | 1 071 | lgG | 13 902 077 | 79 (Japan) |
| Japan (Utsunomiya City) ⁴⁸ | 742 | lgG | 518610 | 79 (Japan) |
| Kenya, blood donors ⁴⁴ | 3 098 | lgG | 47 564 296 | 99 |
| Luxembourg ²⁰ | 1 807 | lgG and lgA ^e | 615 729 | 90 |
| Netherlands blood donors ¹⁵ | 7 361 | lgG, lgM and lgA | 17 097 123 | 86 |
| Netherlands (Rotterdam) ⁶⁴ | 729 (early April) | lgG | 17 097 123 (Netherlands) | 86 |
| Pakistan (Karachi)49 | 1 004 | lgG and lgM | 16 7Ò0 000 ´ | 98 (Pakistan) |
| Pakistan (urban)66 | 24 210 | lgG and lgM | 79 000 000 (urban) | ` 98 |
| | | = | • | |

| | Article ID: BLT.20.265892 | | | | |
|--|---------------------------|------------------|--------------------------------|--------------------|--|
| Qatar ⁵¹ | 937 | lgG | 2800000 | 99 | |
| Republic of Korea ⁵⁹ | 1 500 | lğG | 2 667 341 | 90 (southern | |
| | | .50 | | Republic of Korea) | |
| Spain ³⁶ | 61 075 | lgG | 46 940 000 | 85 | |
| Spain (Barcelona) ³⁰ | 874 | lgG, lgM and lgA | 7 566 000 (Catalonia) | 86 | |
| Switzerland (Geneva) ¹⁰ | 577 (20–27 April) | 0 0 | 500 000 (Catalonia) | 88 | |
| | | lgG | | 88 | |
| Switzerland (Zurich) ²⁸ | 1 644 patients (1–15 | lgG | 1 520 968 (Zurich canton) | 88 | |
| 0 11 117 11129 | April) | | 1000 505 /7 : 1 11 | 00 | |
| Switzerland (Zurich) ²⁸ | 1 640 blood donors | lgG | 1 930 525 (Zurich and Lucerne) | 88 | |
| | (May) | | | | |
| United Kingdom (England) ⁶⁵ | 109 076 | lgG | 56 287 000 | 86 | |
| United Kingdom (Scotland), | 500 | IgG | 5 400 000 | 88 | |
| blood donors ¹⁸ | | | | | |
| USA (10 states) ³⁵ | | | | | |
| Washington, Puget Sound | 3 264 | Pan-lg | 4 273 548 | 90 (Washington) | |
| Utah | 1 132 | Pan-Ig | 3 282 120 | 92 | |
| New York, New York City | 2 482 | Pan-Ig | 9 260 870 | 89 | |
| Missouri | 1 882 | Pan-lg | 6 110 800 | 88 | |
| Florida, south | 1 742 | Pan-Ig | 6345345 | 86 (Florida) | |
| Connecticut | 1 431 | Pan-Ig | 3 562 989 | 88 | |
| Louisiana | 1 184 | Pan-Ig | 4 644 049 | 92= | |
| California, San Francisco Bay | 1 224 | Pan-Ig | 2 173 082 | 92 = 90 | |
| | | • | | | |
| Pennsylvania, Philadelphia | 824 | Pan-Ig | 4 910 139 | 90 | |
| Minnesota, Minneapolis | 860 | Pan-lg | 3857479 | 90 | |
| USA (California, Bay Area) ²⁴ | 1 000 | lgG | 7 753 000 | 90 | |
| USA (California, Los | 863 | IgG and IgM | 7 892 000 | 92 | |
| Angeles) ²² | | | | | |
| USA (California, San | 3 953 | IgG (also PCR | 5174 (in census tract 022 901) | 95 | |
| Francisco) ³³ | | testing) | | | |
| USA (California, Santa | 3 300 | IgG and IgM | 1 928 000 | 90 | |
| Clara) ¹⁹ | | | | | |
| USA (Idaho, Boise) ⁹ | 4 856 | lgG | 481 587 (Ada county) | 92 | |
| USA (Georgia, DeKalb and | 696 | Total Ig | 1806672 | 88 (Georgia) | |
| Fulton counties)53 | | 3 | | () | |
| USA (Idaho, Blaine county) ⁴⁶ | 917 | lgG | 23 089 | 92 | |
| USA (Indiana) ⁵⁴ | 3 629 | IgG (also RT–PCR | 6 730 000 | 89 | |
| | 0 020 | done) | 0.00000 | | |
| USA (Louisiana, Baton | 138 | lgG | 699 200 (East Baton Rouge, | 92 (Louisiana) | |
| Rouge) ⁶³ | 100 | igo | West Baton Rouge, Ascension, | oz (Lodisiaria) | |
| Nouge)** | | | | | |
| LICA /Lauriciana Orleana and | 2.640 | la C | Livingston) | 00 (Louisiana) | |
| USA (Louisiana, Orleans and | 2 640 | lgG | 825 057 | 92 (Louisiana) | |
| Jefferson Parish) ³⁷ | 45.404 | | 40.450.000 | 00 | |
| USA (New York) ²³ | 15 101 | lgG | 19 450 000 | 90 | |
| USA, New York ⁵⁶ | | | | | |
| Columbia University Medical | 742 (2–30 March) | IgG and IgM | 9 260 870 | 89 | |
| Center, New York City | | | | | |
| CareMount central laboratory, | 1 841 | lgG and lgM | 10 189 130 (New York state | 89 | |
| five New York state counties | | | excluding New York City) | | |
| USA (New York, Brooklyn) ²⁷ | 11 092 | lgG | 2 559 903 | 91 | |
| USA (Rhode Island), blood | 1 996 | lgG and lgM | 1 059 000 | 88 | |
| donors ⁴⁵ | | 5 5 | | - | |
| | | | | | |

COVID-19: coronavirus disease-19; Ig: immunoglobin; RT-PCR: real-time polymerase chain reaction.

^a Dates in brackets are the specific dates used when seroprevalence was evaluated at multiple consecutive time points or setting.

^b Some studies focused on age-restricted populations of the specific location under study, for example: people 17–70 years in the Denmark blood donor study (n = 3800000); people 18–79-years in the Luxembourg study (n = 483000);

people < 70 years in the Netherlands blood donor study (n = 13745768); people ≥ 18 years in the New York state study (n = 15280000); people > 19 years in the Utah population of the 10-state United States study (n = 2173082); people ≥ 18 years in Blaine county, Idaho (n = 17611); people 15–64 years in the Kenya blood donor study (n = 27150165); people > 14 years living in private premises in Hungary; people > 18 years (n = 551185) in Baton Rouge, Louisiana; people 18–65 years working in urban locations in Pakistan (n = 22100000); and people > 18 years in Srinagar District, India (n = 1020000). In this table and subsequent analyses, the entire population in the location is considered for consistency across studies.

- ^c Information in parenthesis specify the population.
- ^d Participants were recruited from a large number of districts, but most districts had very few participants; here I included the population of the nine districts with > 1:10 000 sampling ratio (846/1000 participants came from these nine districts).
- ^e Considered positive if both IgG and IgA were positive; in the other studies, detection of any antibody was considered positive.

Table 3. Prevalence of COVID-19 and estimated number of people infected, 2020

| | JI COVID-19 al | Second control of people infected | · |
|--|--------------------------------|--|------------------|
| Country (location) | | Seroprevalence (%) | Estimated no. of |
| | Crude | Adjusted (adjustments) | people infected |
| Argentina (Barrio Padre Mugica) ⁴⁷ | ND | 53.4 (age, sex, household, non-response) | 26 691 |
| Belgium ³⁸ | 5.7 | 6.0 (sampling, age, sex, province) | 695 377 |
| Brazil (133 cities) ²⁵ | 1.39 | 1.62 overall, varying from 0 to 25.0 across 133 cities (test, design) | 1 209 435ª |
| Brazil (Espirito Santo) ³⁴ | 2.1 | ND | 84 391 |
| Brazil (Maranhao) ⁶⁸ | 37 | 40.4 (clustering, stratification, non-response) | 2 877 454 |
| Brazil (Rio de Janeiro), blood | 6 | 4.7 (age, sex, test) | 811 452 |
| donors ⁴¹ | | | |
| Brazil (Rio Grande do Sul) ¹⁷ | 0.222 | 0.222 (sampling) ^b | 25 283 |
| Brazil (Sao Paulo) ⁴² | 5.2 | 4.7 (sampling design) | 14 017 |
| Canada (British Columbia) ⁵⁰ | 0.45 | 0.55 (age) | 27 890 |
| Chile (Vitacura) ⁴³ | 11.2 | ND | 9 500 |
| China, blood donors ⁵⁵ | | | |
| Wuhan | 3.87 | ND | 433 827 |
| Shenzhen | 0.06 | ND | 7 818 |
| Shijiazhuang | 0.02 | ND | 2 206 |
| China (Wuhan) ¹⁴ | 10 | ND | 1 108 000 |
| China (Wuhan) ³² | 8.36 (3.53 for entire period) | ND (2.80 (age, sex, test) for entire period) | 926 288 |
| China (Guangzhou), blood | 0.09 | ND | 104 783 |
| donors ⁶⁰ | | | |
| China (several regions) ⁴⁰ | | | |
| Hubei (not Wuhan) | 3.6 | ND | 1 718 110 |
| Chongqing | 3.8 | ND | 11 956 109 |
| Sichuan | 0.6 | ND | 487 847 |
| Guangdong | 2.2 | ND | 2 522 010 |
| Croatia ²⁶ | 1.27 ^c | ND | 51 765 |
| Denmark, blood donors ¹² | 2 | 1.9 (test) | 109 665 |
| Denmark (Faroe Islands) ⁵² | 0.6 | 0.7 (test) | 365 |
| France (Crepy-en-Valois) ³⁹ | 10.4 | ND | 620 105 |
| France (Oise) ¹³ | 25.9 | ND | 1 548 000 |
| Germany (Gangelt) ¹⁶ | 15 | 20.0 (test, cluster, symptoms) | 2 5 1 9 |
| Germany (Frankfurt) ²¹ | 0.6 | ND | 16 086 |
| Greece ⁶² | 0.42 (April) | 0.49 (age, sex, region)d | 51 023 |
| Hungary ⁵⁷ | 0.67 | 0.68 (design, age, sex, district) | 65 671 |
| lceland ⁵⁸ | 2.3 | 0.9 (including those positive by PCR) | 3 177 |
| | (quarantined), 0.3 (unknown | | |
| | exposure) | | |
| India (Mumbai) ⁶¹ | 54.1 in slum | 58.4 in slum areas, 17.3 in non-slum areas (test, | 534 750 |
| | areas, 16.1 in | age, sex) | |
| | non-slum areas | | |
| India (Srinagar) ⁶⁷ | 3.8 | 3.6 (age, sex) | 54 000 |
| Islamic Republic of Iran (Guilan) ⁸ | 22 | 33.0 (test, sampling) | 770 000 |
| Italy (Apulia), blood donors ³¹ | 0.99 | ND | 39 887 |
| Japan (Kobe) ¹¹ | 3.3 | 2.7 (age, sex) | 40 999 |
| Japan (Tokyo) ²⁹ | 3.83 | ND | 532 450 |
| Japan (Utsunomiya City) ⁴⁸ | 0.4 | 1.23 (age, sex, distance to clinic, district, cohabitants) | 6 378 |
| Kenya, blood donors ⁴⁴ | 5.6 | 5.2 (age, sex, region, test) | 2 783 453 |
| Luxembourg ²⁰ | 1.9 | 2.1 (age, sex, district) | 12 684 |
| Netherlands, blood donors ¹⁵ | 2.7 | ND | 461 622 |
| Netherlands (Rotterdam) ⁶⁴ | 3 | ND | 512 910 |

Article ID: BLT.20.265892 Pakistan (Karachi)⁴⁹ 16.3 (20.0 in 11.9 (age, sex; 15.1 in East, 8.7 in Malir) 1 987 300 East, 12.7 in Malir) Pakistan (urban)⁶⁶ 17.5 ND 13 825 000 Qatar⁵¹ 30.4 (24.0 for 851 200 ND entire period) Republic of Korea⁵⁹ 0.07 ND 1867 Spain³⁶ ND 5.0e (sampling, age, sex, income) 2 347 000 Spain (Barcelona)30 14.3 ND 1081938 Switzerland (Geneva)¹⁰ 10.6 10.9 (test, age, sex) 54 500 Switzerland (Zurich)²⁸ 1.3 in patients during 1–15 April and 1.6 in blood 19 773 (Zurich); Unclear donors in May (multivariate Gaussian 30 888 (Zurich conditioning) and Lucerne) United Kingdom (England)⁶⁵ 5.6 6.0 (test, sampling) 3 360 000 United Kingdom (Scotland) blood 1.2 ND 64 800 donors¹⁸ USA (six states)35 (age, sex, test) Washington, Puget Sound 48 291 1.3 1.1 2.4 2.2 71 550 New York, New York City 5.7 6.9 641778 Missouri 2.9 2.7 161 936 Florida, south 2.2 1.9 117 389 Connecticut 4.9 4.9 176 012 ND 5.8 267 033 Louisiana California, San Francisco Bay ND 1 64 626 Pennsylvania, Philadelphia ND 3.2 156 633 Minnesota, Minneapolis ND 2.4 90 651 USA (California, Bay Area)²⁴ 0.4 (blood 0.1 (test and confirmation) 7 753 donors) 4.06 USA (California, Los Angeles)²² 4.65 (test, sex, race and ethnicity, income) 367 000 USA (California, San Francisco)33 6.1 (age, sex, race and ethnicity, test) 4.3 in the 316 census track USA (California, Santa Clara)¹⁹ 2.6 (test, sampling, cluster) 51 000 1.5 USA (Idaho, Boise)9 1.79 ND 8620 USA (Georgia, DeKalb and Fulton 2.7 2.5 (age, sex, race and ethnicity) 45 167 counties)53 USA (Idaho, Blaine county)46 22.4 23.4 (test, age, sex, household) 5 403 2.8 (age, race, Hispanic ethnicity) USA (Indiana)54 2.3 (IgG or PCR) 187 802 USA (Louisiana, Baton Rouge)⁶³ 6.6 (census, race, parish) including PCR 46 147 6 positives USA (Louisiana, Orleans and 6.9 (IgG or PCR) 6.9 for IgG (census weighting, demographics) 56 578 Jefferson Parish)37 USA (New York)²³ 12.5 14.0 (test, sex, age race and ethnicity, region) 2723000 USA, New York⁵⁶ Columbia University Medical Center, 5 ND 463 044 New York City CareMount central laboratory, five 1.8 ND 183 404 New York state counties USA (New York, Brooklyn)²⁷ 47 ND 1 203 154 USA (Rhode Island), blood 3.9 41 384 ND

COVID-19: coronavirus disease 2019; ND: no data available; PCR: polymerase chain reaction; test: test performance.

donors45

^a The authors calculated 760 000 to be infected in the 90 cities that had 200–250 samples tested, but many of the other 43 cities with < 200 samples may be equally or ever better represented since they tended to be smaller than the 90 cities (mean population 356 213 versus 659 326).

- ^b An estimate is also provided adjusting for test performance, but the assumed specificity of 99.0% seems inappropriately low, since as part of the validation process the authors found that several of the test-positive individuals had household members who were also infected, thus the estimated specificity was deemed by the authors to be at least 99.95%.
- ° 1.20% in workers in Split without mobility restrictions, 3.37% in workers in Knin without mobility restrictions, 1.57% for all workers without mobility restrictions; Split and Knin tended to have somewhat higher death rates than nationwide Croatia, but residence of workers is not given, so the entire population of the country is used in the calculations.
- ^d An estimate is also provided adjusting for test performance resulting in adjusted seroprevalence of 0.23%, but this seems inappropriately low, since the authors report that all positive results were further validated by ELISA.
- e 5.0% with point of care test, 4.6% with immunoassay, 3.7% with both tests positive, 6.2% with at least one test positive.

Notes: Of the studies where seroprevalence was evaluated at multiple consecutive time points, the seroprevalence estimate was the highest in the most recent time interval with few exceptions, for example: in the Switzerland (Geneva) study, ¹⁰ the highest value was seen 2 weeks before the last time interval; in the Switzerland (Zurich) study, ²⁸ the highest value was seen in the period 1–15 April for patients at the university hospital and in May for blood donors; and in the China (Wuhan) study, ³² the highest value was seen about 3 weeks before the last time interval.

Table 4. Deaths from COVID-19 and inferred infection fatality rates, overall and in people vounger than 70 years, by location, 2020

| younger than 70 years, by location, 2020 | | | | | | |
|---|--------------------------------------|---|--|---|--|--|
| Location | Deaths from COVID- 19, no. (date) | Inferred infection fatality rate (corrected), % | % of deaths from COVID-19 in people <70 years ^a | Infection fatality rate in people < 70 years (corrected), | | |
| Argentina (Barrio Padre | 44 (1 July) | 0.16 (0.13) | ~70 | 0.11 (0.09) | | |
| Mugica) ⁴⁷ | | | | | | |
| Belgium ³⁸ | 7594 (30 April) | 1.09 (0.87) | 10 | 0.13 (0.10) | | |
| Brazil (133 cities) ²⁵ | _b | Median 0.30 (0.27) | 31 (< 60 years) | 0.10 (0.9) | | |
| Brazil (Espirito Santo) ³⁴ | 363 (21 May) | 0.43 (0.39) | 31 (Brazil, < 60 years) 23 | 0.14 (0.13) | | |
| Brazil (Maranhao) ⁶⁸ Brazil (Rio de Janeiro), blood | 4272 (8 August) 1019 (3 May) | 0.15 (0.14) 0.12 (0.11) | 31 (Brazil, < 60 years) | 0.04 (0.03) 0.04 (0.04) | | |
| donors ⁴¹ | 1019 (3 May) | 0.12 (0.11) | 31 (Diazii, ~ 00 years) | 0.04 (0.04) | | |
| Brazil (Rio Grande do Sul) ¹⁷ | 124 (14 May) | 0.49 (0.39) | 31 (Brazil, < 60 years) | 0.19 (0.15) | | |
| Brazil (Sao Paulo) ^{c,42} | Unknown (15 May) | Unknown, but likely > 0.4 | | | | |
| Canada (British Columbia) ⁵⁰ | 164 (28 May) | 0.59 (0.59) | 13 | 0.08 (0.08) | | |
| Chile (Vitacura) c,43 | Unknown (18 May) | Unknown, but likely | 36 (Chile) | Unknown, but likely | | |
| , | - (-)) | < 0.2 | , | < 0.1 | | |
| China, blood donors ⁵⁵ | | | | | | |
| Wuhan | 1935 (20 February) | 0.45 (0.41) | 50 | 0.24 (0.22) | | |
| Shenzhen | 1 (5 March) | 0.01 (0.01) | About 50 (if similar to Wuhan) | 0.01 (0.01) | | |
| Shijiazhuang | 1 (27 February) | 0.05 (0.04) | About 50 (if similar to Wuhan) | 0.03 (0.02) | | |
| China (Wuhan) ¹⁴ | 3869 (2 May) | 0.35 (0.31) | 50 | 0.19 (0.15) | | |
| China (Wuhan) ³² | 3869 (13 April) | 0.42 (0.38) | 50 | 0.23 (0.21) | | |
| China (Guangzhou), blood donors ⁶⁰ | 8 (5 April) | 0.00 (0.00) | About 50 (if similar to Wuhan) | 0.00 (0.00) | | |
| China (several regions) ⁴⁰ | C40 (40 A mil) | 0.04 (0.02) | About 50 /if similar to | 0.00 (0.00) | | |
| Hubei (not Wuhan) | 643 (12 April) | 0.04 (0.03) | About 50 (if similar to Wuhan) | 0.02 (0.02) | | |
| Chongqing | 6 (12 April) | 0.00 (0.00) | About 50 (if similar to Wuhan) | 0.00 (0.00) | | |
| Guangdong | 8 (12 April) | 0.00 (0.00) | About 50 (if similar to Wuhan) | 0.00 (0.00) | | |
| Sichuan | 3 (12 April) | 0.00 (0.00) | About 50 (if similar to Wuhan) | 0.00 (0.00) | | |
| Croatia ²⁶ | 79 (3 May) | 0.15 (0.14) | 13 ´ | 0.02 (0.02) | | |
| Denmark, blood donors ¹² | 370 (21 April) | 0.34 (0.27) | 12 | 0.05 (0.04) | | |
| Faroe Islands ⁵² | 0 (5 May) | 0.00 (0.00) | 0 | 0.00 (0.00) | | |
| France (Crepy-en-Valois) ³⁹ | 2325 (5 May) ^d | 0.37 (0.30) | 7 (France, < 65 years) | 0.04 (0.03) | | |
| France (Oise) ¹³ | 932 (7 April) ^d | 0.06 (0.05) | 7 (France, < 65 years) | 0.01 (0.01) | | |
| Germany (Gangelt) ¹⁶ | 7 (15 April) | 0.28 (0.25) | 0 14 (Cormony) | 0.00 (0.00) | | |
| Germany (Frankfurt) ²¹ Greece ⁶² | 42º (17 April) 121 (22 April) | 0.26 (0.21) 0.24 (0.19) | 14 (Germany) 30 | 0.04 (0.03) 0.09 (0.07) | | |
| Hungary ⁵⁷ | 442 (15 May) | 0.67 (0.54) | No data | No data | | |
| Iceland ⁵⁸ | 10 (1 June) | 0.30 (0.30) | 30 | 0.10 (0.10) | | |
| India (Mumbai) ⁶¹ | 495 (13–20 July) | 0.09 (0.07) | 50 (< 60 years, India) | 0.04 (0.03) | | |
| India (Srinagar) ⁶⁷ | 35 (15 July) ^f | 0.06 (0.05) | 50 (< 60 years, India) | 0.03 (0.03) | | |
| Islamic Republic of Iran (Guilan) ⁸ | 617 (23 April) | 0.08 (0.07) | No data | No data [′] | | |
| Italy (Apulia), blood donors ³¹ | 530 (22 May) | 1.33 (1.20) | 15 (Italy) | 0.24 (0.22) | | |
| Japan (Kobe) ¹¹ | 10 (mid-April) | 0.02 (0.02) | 21 (Ĵapan) | 0.01 (0.01) | | |
| Japan (Tokyo) ²⁹ | 189 (11 May) | 0.04 (0.03) | 21 (Japan) | 0.01 (0.01) | | |
| Japan (Utsunomiya City) ⁴⁸ | 0 (14 June) | 0.00 (0.00) | 0 | 0.00 (0.00) | | |

| | Article ID: BLT.20.265892 | | | | | |
|---|---|--------------------------|------------------------------------|--------------------------|--|--|
| Kenya, blood donors ⁴⁴ | 64 (31 May) | 0.00 (0.00) | 58 (< 60 years) | 0.00 (0.00) | | |
| Luxembourg ²⁰ | 92 (2 May) | 0.73 (0.58) | 9 | 0.07 (0.06) | | |
| Netherlands, blood donors ¹⁵ | 3134 (15 April) | 0.68 (0.68) | 11 | 0.09 (0.09) | | |
| Netherlands (Rotterdam) ⁶⁴ | 3134 (15 April) | 0.65 (0.52) | 11 | 0.08 (0.06) | | |
| Pakistan (Karachi) ⁴⁹ | ~1500 (9 July) ⁹ | 0.08 (0.07) | ~70 | 0.06 (0.05) | | |
| Pakistan (urban) ⁶⁶ | 5266 (13 July) ^h | 0.04 (0.04) | ~70 | 0.03 (0.03) | | |
| Qatar ⁵¹ | 93 (19 June) | 0.01 (0.01) | 74 | 0.01 (0.01) | | |
| Republic of Korea ⁵⁹ | 2 (3 June) ⁱ | 0.10 (0.09) | 0 | 0.00 (0.00) | | |
| Spain ³⁶ | 26 920 (11 May) | 1.15 (0.92) | 13 | 0.18 (0.14) | | |
| Spain (Barcelona) ³⁰ | 5137 (2 May) | 0.48 (0.48) | 13 (Spain) | 0.07 (0.07) | | |
| Switzerland (Geneva) ¹⁰ | 243 (30 April) | 0.45 (0.36) | 8 | 0.04 (0.03) | | |
| Switzerland (Zurich) ²⁸ | 107 (15 April, Zurich), | 0.51 (0.41) | 8 (Switzerland) | 0.05 (0.04) | | |
| (| 147 (22 May, Zurich | 0.01 (0111) | o (0.111 <u>2</u> 011 <u>2</u> 11) | (3.33) | | |
| | and Lucerne) | | | | | |
| England ⁶⁵ | 38 854 (9 July) | 1.16 (0.93) | 20 | 0.27 (0.22) | | |
| Scotland, blood donors ¹⁸ | 47 (1 April) | 0.07 (0.06) | 9 (< 65 years) | 0.01 (0.01) | | |
| USA (10 states) ³⁵ | (,) | 0.07 (0.00) | o (oo you.e) | 0.01 (0.01) | | |
| Washington, Puget Sound | 207 (4 April) | 0.43 (0.43) | 10 (state, < 60 years) | 0.05 (0.05) | | |
| Utah | 58 (4 May) | 0.08 (0.08) | 28 (< 65 years) | 0.03 (0.03) | | |
| New York | 4146 (4 April) | 0.65 (0.65) | 34 (state) | 0.25 (0.25) | | |
| Missouri | 329 (30 April) | 0.20 (0.20) | 23 | 0.05 (0.05) | | |
| Florida, south | 295 (15 April) | 0.25 (0.25) | 28 (state) | 0.08 (0.08) | | |
| Connecticut | 2718 (6 May) | 1.54 (1.54) | 18 | 0.31 (0.31) | | |
| Louisiana | 806 (11 April) | 0.30 (0.30) | 32 | 0.10 (0.10) | | |
| California, San Francisco Bay | 321 (1 May) | 0.50 (0.50) | 25 | 0.14 (0.14) | | |
| Pennsylvania, Philadelphia | 697 (26 April) | 0.45 (0.45) | 21 (state) | 0.10 (0.14) | | |
| Minnesota, Minneapolis | 436 (13 May) | 0.48 (0.48) | 20 (state) | 0.10 (0.10) | | |
| USA (California, Bay Area) ²⁴ | 12 (22 March) | 0.15 (0.12) | 25 | 0.04 (0.03) | | |
| USA (California, Los | 724 (19 April) | 0.20 (0.18) | 24 (< 65 years) | 0.06 (0.05) | | |
| Angeles) ²² | 721 (1074) | 0.20 (0.10) | 21 (100 yours) | 0.00 (0.00) | | |
| USA (California, San | 0 (4 May) | 0.00 (0.00) | 0 | 0.00 (0.00) | | |
| Francisco) ³³ | o (Timay) | 0.00 (0.00) | ŭ | 0.00 (0.00) | | |
| USA (California; Santa Clara) ¹⁹ | 94 (22 April) | 0.18 (0.17) | 35 | 0.07 (0.06) | | |
| USA (Idaho, Boise) ⁹ | 14 (24 April) | 0.16 (0.13) | 14 (Idaho) | 0.02 (0.02) | | |
| USA (Georgia) ⁵³ | 198 (7 May) | 0.44 (0.44) | 30 | 0.15 (0.15) | | |
| USA (Idaho, Blaine county) ⁴⁶ | 5 (19 May) | 0.10 (0.08) | 14 (Idaho) | 0.02 (0.01) | | |
| USA (Indiana) ⁵⁴ | 1099 (30 April) | 0.58 (0.46) | 24 | 0.16 (0.13) | | |
| USA (Louisiana, Baton | 420 (30 July) | 0.91 (0.73) | 32 (Louisiana) | 0.32 (0.25) | | |
| Rouge) ⁶³ | 0 (00 00) | 0.0 . (0 0) | 02 (200.000.00) | 0.02 (0.20) | | |
| USA (Louisiana, Orleans and | 925 (16 May) | 1.63 (1.31) | 32 | 0.57 (0.46) | | |
| Jefferson Parish) ³⁷ | (, , | (1101) | - | (3113) | | |
| USA (New York) ²³ | 18 610 (30 April) ^j | 0.68 (0.54) ^j | 34 | 0.26 (0.23)d | | |
| USA (New York Columbia | 965 (28 March, New | 0.15 (0.14) | 34 | 0.06 (0.05) | | |
| University Medical Center, | York state) | ((()) | - | (3.22) | | |
| New York City and CareMount | , | | | | | |
| central laboratory, five New | | | | | | |
| York state counties) ⁵⁶ | | | | | | |
| USA (New York, Brooklyn) ²⁷ | 4894 (19 May) ^j | 0.41 (0.33) ^j | 34 (New York state) | 0.15 (0.14) ^d | | |
| USA (Rhode Island), blood | 430 (11 May) | 1.04 (0.83) | 17 | 0.20 (0.16) | | |
| donors ⁴⁵ | ` ,, | (/ | | · -/ | | |
| | | | | | | |

COVID-19: coronavirus disease 2019.

^a Whenever the number or proportion of COVID-19 deaths at age < 70 years was not provided in the paper, I retrieved the proportion of these deaths from situation reports of the relevant location. If I could not find this information for the specific location, I used a larger geographic area. For Brazil, the closest information that I found was from a news report.⁷⁷ For Croatia, I retrieved data on age for 45/103 deaths through Wikipedia.⁷⁸

- ^b Data are provided by the authors for deaths per 100 000 population in each city along with inferred infection fatality rate in each city, with wide differences across cities; the infection fatality rate shown here is the median across the 36 cities with 200–250 samples and at least one positive sample (the interquartile range for the uncorrected infection fatality rate is 0.20–0.60% and across all cities is 0–2.4%, but with very wide uncertainty in each city). A higher infection fatality rate is alluded to in the preprint, but the preprint also shows a scatter diagram for survey-based seroprevalence versus reported deaths per population with a regression slope that agrees with an infection fatality rate of 0.3%.
- ^c Information on deaths was not available for the specific locations. In the Sao Paulo study, the authors selected six districts of Sao Paulo most affected by COVID-19, they do not name the districts and the number of deaths as of mid-May is not available, but using data for death rates across all Sao Paulo would give an infection fatality rate of > 0.4% overall. In the Vitacura study, similarly one can infer from the wider Santiago metropolitan area that the infection fatality rate in the Vitacura area would probably be < 0.2% overall.
- ^d For France, government situation reports provide the number of deaths per region only for in-hospital deaths; therefore, I multiplied the number of in-hospital deaths by a factor equal to: total number of deaths/in-hospital deaths for all of France.
- ^e Estimated from no. of deaths in Hesse province on 17 April × proportion of deaths in the nine districts with key enrolment (enrolment ratio > 1:10 000) in the study among all deaths in Hesse province.
- ^f I calculated the approximate number of deaths assuming the same case fatality ratio in the Srinagar district as in the Jammu and Kashmir state where it is located.
- ⁹ For Karachi, it is assumed that about 30% of COVID-19 deaths in Pakistan are in Karachi (since about 30% of the cases are there).
- ^h The number of deaths across all Pakistan; I assumed that this number is a good approximation of deaths in urban areas (most deaths occur in urban areas and there is some potential underreporting).
- ⁱ I calculated the approximate number of deaths from the number of cases in the study areas in south-western Seoul, assuming a similar case fatality as in Seoul overall.
- ^j Confirmed COVID-19 deaths; inclusion of probable COVID-19 deaths would increase the infection fatality rate estimates by about a quarter.

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Table 5. Infection fatality rates for coronavirus disease-19 inferred from preliminary nationwide seroprevalence data, 2020

| Country | Sample size (antibody) | Date | Reported seroprevalence (%) | Population, no. | Deaths, no. (date) | Inferred infection fatality rate (corrected), % |
|---------------------------|---------------------------|----------------|-----------------------------------|--------------------|-----------------------|---|
| Afghanistan ⁷⁵ | 9 500 (IgG?) | August? | 31.5 | 39 021 453 | 1300 (8 May) | 0.01 (0.01) |
| Czechia ⁷¹ | 26 549 (IgG) | 23 April-1 May | 0.4 | 10 710 000 | 252 (4 May) | 0.59 (0.47) |
| Finland ⁶⁹ | 674 (IgG) | 20–26 Aprila | 2.52 | 5 541 000 | 211 (30 April) | 0.15 (0.12) |
| Georgia ⁷⁶ | 1 068 (IgG?) | 18–27 May | 1 | 3 988 264 | 12 (30 May) | 0.03 (0.03)b |
| Israel ⁷² | 1 709 (IgG?) | May | 2–3 | 9 198 000 | 299 (10 June) | 0.13 (0.10)° |
| Russian | 650 000 (lgG?) | June? | 14 | 145 941 776 | 5859 (7 June) | 0.03 (0.03) |
| Federation ⁷⁴ | , | | | | , , | , , |
| Slovenia ⁷³ | 1368 (IgG?) | April | 3.1 | 2 079 000 | 92 (1 May) | 0.14 (0.11) |
| Sweden ⁷⁰ | 1 200 (lgG) | 18–24 May | 6.3 | 10 101 000 | 4501 (28 May) | 0.71 (0.57) |

COVID-19: coronavirus disease 2019; Ig: immunoglobin.

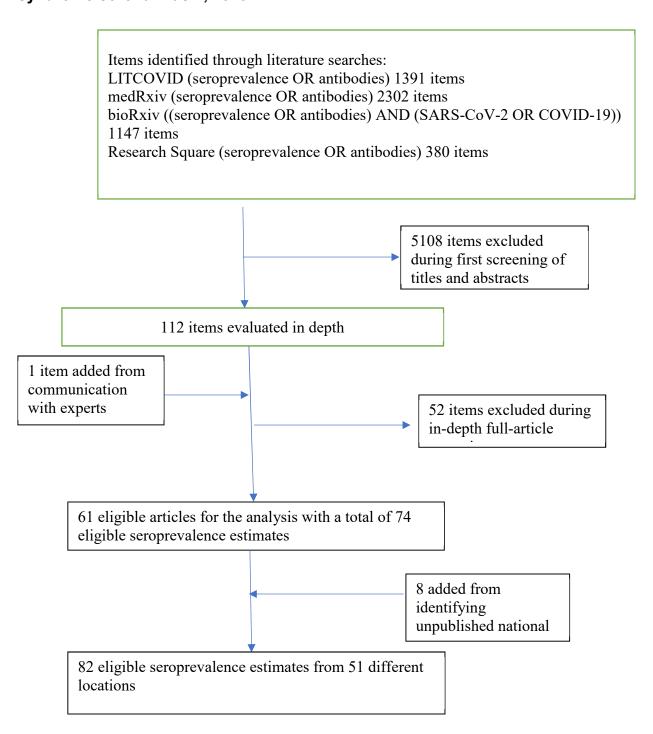
Notes: These are countries for which no eligible studies were retrieved in the literature search. The results of these studies have been announced to the press and/or in preliminary reports, but are not yet peer reviewed and published. The question marks indicate that the antibody type or date were not clear.

^a The seroprevalence was slightly lower in subsequent weeks.

^b The survey was done in Tbilisi, the capital city with a population 1.1 million. I could not retrieve the count of deaths in Tbilisi, but if more deaths happened in Tbilisi, then the infection fatality rate may be higher, but still < 0.1%.

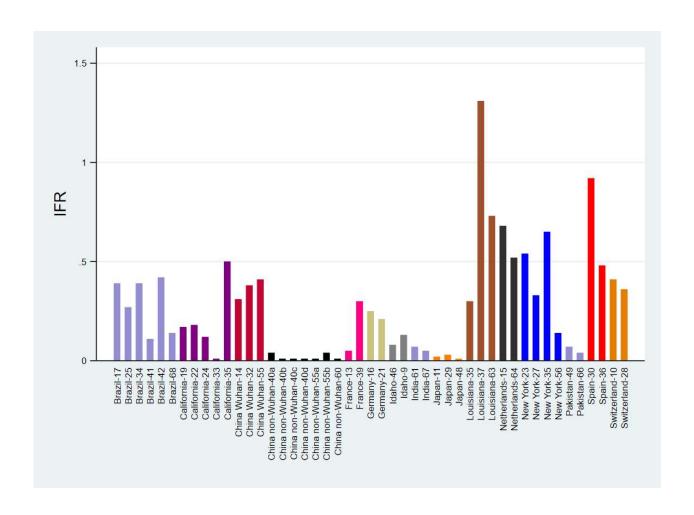
^c Assuming a seroprevalence of 2.5%.

Fig. 1. Flowchart for selection of seroprevalence studies on severe acute respiratory syndrome coronavirus 2, 2020



COVID-19: coronavirus disease 2019; SARS-CoV-2: severe acute respiratory syndrome coronavirus 2.

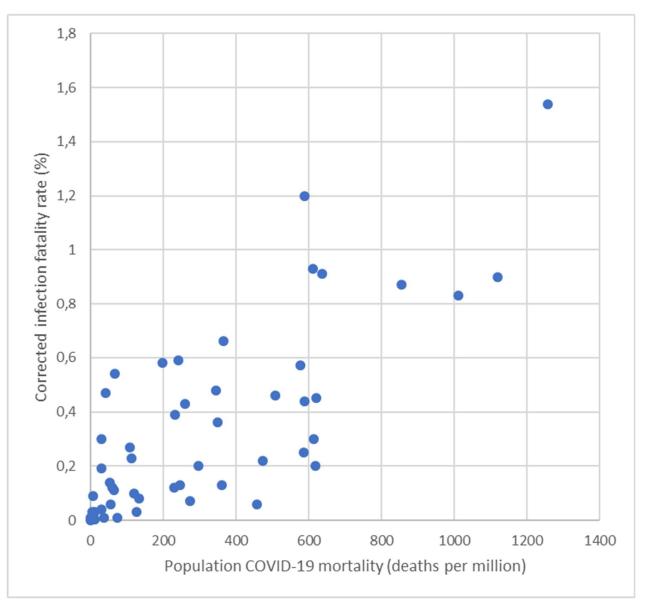
Fig. 2. Estimates of infection fatality rates for COVID-19 in locations that had two or more estimates, 2020



COVID-19: coronavirus disease 2019.

Notes: Locations are defined at the level of countries, except for the USA where they are defined at the level of states and China is separated into Wuhan and non-Wuhan areas. Corrected infection fatality rate estimates are shown (correcting for what types of antibodies were assayed).

Fig. 3. Corrected estimates of COVID-19 infection fatality rate in each location plotted against COVID-19 mortality rate as of September 12, 2020 in that location



COVID-19: coronavirus disease 2019

Notes: Locations are defined at the level of countries, except for the United Kingdom of Great Britain and Northern Ireland where they are defined by jurisdiction, USA are defined at the level of states and China is separated into Wuhan and non-Wuhan areas. Included locations are: Afghanistan; Argentina, Belgium Brazil; Canada; Chile; China (non-Wuhan and Wuhan); Croatia; Czechia; Denmark; Faroe Islands; Finland; France; Georgia; Germany; Greece; Hungary; Iceland; India; Islamic Republic of Iran (Islamic Republic of); Israel; Italy; Japan; Kenya; Luxembourg; Netherlands; Pakistan; Qatar; Russian Federation; Slovenia; Republic of Korea; Spain; Sweden; Switzerland; United Kingdom (England, Scotland); and USA (California, Connecticut, Florida, Georgia, Idaho, Indiana, Louisiana, Minnesota, Missouri, New York, Pennsylvania, Rhode Island, Utah, Washington). When several infection fatality rate estimates were available from multiple studies for a location, the sample size-weighted mean is used. One outlier location with very high deaths per million population (1702 for New York) is not shown.