

Age–sex differences in the global burden of lower respiratory infections and risk factors, 1990–2019: results from the Global Burden of Disease Study 2019



GBD 2019 LRI Collaborators*



Summary

Background The global burden of lower respiratory infections (LRIs) and corresponding risk factors in children older than 5 years and adults has not been studied as comprehensively as it has been in children younger than 5 years. We assessed the burden and trends of LRIs and risk factors across all age groups by sex, for 204 countries and territories.

Methods In this analysis of data for the Global Burden of Diseases, Injuries, and Risk Factors Study (GBD) 2019, we used clinician-diagnosed pneumonia or bronchiolitis as our case definition for LRIs. We included International Classification of Diseases 9th edition codes 079.6, 466–469, 470.0, 480–482.8, 483.0–483.9, 484.1–484.2, 484.6–484.7, and 487–489 and International Classification of Diseases 10th edition codes A48.1, A70, B97.4–B97.6, J09–J15.8, J16–J16.9, J20–J21.9, J91.0, P23.0–P23.4, and U04–U04.9. We used the Cause of Death Ensemble modelling strategy to analyse 23 109 site-years of vital registration data, 825 site-years of sample vital registration data, 1766 site-years of verbal autopsy data, and 681 site-years of mortality surveillance data. We used DisMod-MR 2.1, a Bayesian meta-regression tool, to analyse age–sex-specific incidence and prevalence data identified via systematic reviews of the literature, population-based survey data, and claims and inpatient data. Additionally, we estimated age–sex-specific LRI mortality that is attributable to the independent effects of 14 risk factors.

Findings Globally, in 2019, we estimated that there were 257 million (95% uncertainty interval [UI] 240–275) LRI incident episodes in males and 232 million (217–248) in females. In the same year, LRIs accounted for 1.30 million (95% UI 1.18–1.42) male deaths and 1.20 million (1.07–1.33) female deaths. Age-standardised incidence and mortality rates were 1.17 times (95% UI 1.16–1.18) and 1.31 times (95% UI 1.23–1.41) greater in males than in females in 2019. Between 1990 and 2019, LRI incidence and mortality rates declined at different rates across age groups and an increase in LRI episodes and deaths was estimated among all adult age groups, with males aged 70 years and older having the highest increase in LRI episodes (126.0% [95% UI 121.4–131.1]) and deaths (100.0% [83.4–115.9]). During the same period, LRI episodes and deaths in children younger than 15 years were estimated to have decreased, and the greatest decline was observed for LRI deaths in males younger than 5 years (–70.7% [–77.2 to –61.8]). The leading risk factors for LRI mortality varied across age groups and sex. More than half of global LRI deaths in children younger than 5 years were attributable to child wasting (population attributable fraction [PAF] 53.0% [95% UI 37.7–61.8] in males and 56.4% [40.7–65.1] in females), and more than a quarter of LRI deaths among those aged 5–14 years were attributable to household air pollution (PAF 26.0% [95% UI 16.6–35.5] for males and PAF 25.8% [16.3–35.4] for females). PAFs of male LRI deaths attributed to smoking were 20.4% (95% UI 15.4–25.2) in those aged 15–49 years, 30.5% (24.1–36.9) in those aged 50–69 years, and 21.9% (16.8–27.3) in those aged 70 years and older. PAFs of female LRI deaths attributed to household air pollution were 21.1% (95% UI 14.5–27.9) in those aged 15–49 years and 18.2% (12.5–24.5) in those aged 50–69 years. For females aged 70 years and older, the leading risk factor, ambient particulate matter, was responsible for 11.7% (95% UI 8.2–15.8) of LRI deaths.

Interpretation The patterns and progress in reducing the burden of LRIs and key risk factors for mortality varied across age groups and sexes. The progress seen in children younger than 5 years was clearly a result of targeted interventions, such as vaccination and reduction of exposure to risk factors. Similar interventions for other age groups could contribute to the achievement of multiple Sustainable Development Goals targets, including promoting wellbeing at all ages and reducing health inequalities. Interventions, including addressing risk factors such as child wasting, smoking, ambient particulate matter pollution, and household air pollution, would prevent deaths and reduce health disparities.

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*Collaborators listed at the end of the paper

Correspondence to:
Dr Hmwe Hmwe Kyu, Institute for Health Metrics and Evaluation, University of Washington, Seattle, WA 98195, USA
hmwekyu@uw.edu

Research in context

Evidence before this study

The burden of lower respiratory infections (LRI) among children younger than 5 years has been studied extensively by several groups, including the WHO Maternal and Child Epidemiology Estimation group and the Global Burden of Diseases, Injuries, and Risk Factors Study (GBD). We searched PubMed for the terms (“lower respiratory infection*”OR “LRI”)AND (“burden” OR “estimates”) AND (“age” OR “sex” OR “gender”) AND (“differenc*” OR “discrepan*” OR “disparit*”), with no language restrictions, for publications from Jan 1, 1980, to July 22, 2022. Our search identified 21 studies that reported population-based LRI morbidity and mortality estimates. Of these studies, 15 focused on either a single location or a subset of countries or regions, and six studies reported the LRI estimates at the global level. None of those studies reported the burden of LRIs attributable to risk factors for people older than 5 years by age and sex. We also did not find any studies reporting risk-deleted LRI mortality estimates. GBD 2017 estimated 2.56 million (95% uncertainty interval [UI] 2.44–2.66) LRI deaths among all ages and 0.80 million (0.75–0.87) LRI deaths in children younger than 5 years in 2017. The GBD 2017 LRI paper evaluated the risk factors and interventions that have affected the burden of LRIs among children younger than 5 years in 195 countries and territories.

Added value of this study

GBD 2019 included new data sources on LRI mortality and morbidity and used an enhanced standardised approach to adjust data from different sources (using different case definitions or measurement methods) to improve the comparability of data. We assessed the LRI burden for all age groups by sex for 204 countries and territories. We also assessed, for the first time, the burden of LRIs attributable to risk factors for children aged 5–14 years, as well as different adult age groups. Lastly, for the first time, we provided the risk-deleted mortality estimates that represent the LRI mortality rates that would have been observed if the combined effects of all evaluated risk factors were removed.

Implications of all the available evidence

Our study provides a comprehensive assessment of the LRI burden and risk factors across different age groups by sex. We identify the regions, countries, and age–sex groups with the highest LRI incidence and mortality to inform targeted interventions. By analysing the LRI burden by time, and identifying the leading risk factors by age groups separately for males and females, we provide insight into policy planning and resource prioritisation for addressing the uneven progress in reducing the LRI burden.

Introduction

Lower respiratory infections (LRIs), mainly caused by bacteria such as *Streptococcus pneumoniae* and *Haemophilus influenzae* type b and viruses such as influenza and respiratory syncytial virus, are a leading cause of death globally, killing more than 2 million people every year.¹ LRIs are also the leading underlying cause of sepsis, which is a major cause of health loss and death worldwide.² Global initiatives to tackle LRIs, such as the Global Action Plan for the Prevention and Control of Pneumonia and Diarrhoea,³ the Stop Pneumonia Initiative,⁴ and the Integrated Management of Childhood Illness initiative,⁵ are targeted at children younger than 5 years. Current literature on the burden of LRIs also focuses primarily on children younger than 5 years; less attention is paid to the LRI burden among children older than 5 years and adults. Evidence indicates that males are more susceptible to LRIs than females, possibly due to factors such as differences in immune response to infection and behavioural factors such as smoking.⁶ Understanding the current burden and trends of LRIs across all age groups by sex is essential for identifying areas of intervention.

Although measuring the burden of LRIs is a crucial input in policy decision making, the assessment of modifiable risk factors for LRIs can inform preventive interventions. With the ageing of populations, it is increasingly important to assess LRI risk factors, especially those for which exposure is not declining, such as ambient

particulate matter air pollution, and compare them to risk factors for which exposure is decreasing, such as household air pollution.⁷ Understanding the changing LRI burden attributable to various risk factors across the entire age spectrum can assist in identifying priorities for targeted interventions. To our knowledge, the global burden of LRIs attributable to risk factors for age groups other than those younger than 5 years has not been comprehensively studied. The objective of this study is to assess the burden and trends of LRIs and risk factors across all age groups by sex for 204 countries and territories. This manuscript was produced as part of the Global Burden of Diseases, Injuries, and Risk Factors Study (GBD) Collaborator Network and in accordance with the GBD Protocol.

Methods

Overview

Detailed methods for GBD 2019 have been published elsewhere.¹⁷ Here, we describe the methods and estimation strategies for LRIs and risk factors. In compliance with the Guidelines for Accurate and Transparent Health Estimates Reporting (GATHER), input data sources and code for each step of the estimation process are available on the Global Health Data Exchange.

Case definition

We used clinician-diagnosed pneumonia or bronchiolitis as our case definition for LRIs. We included

For input data sources and code see <http://ghdx.healthdata.org/gbd-2019/data-input-sources>

International Classification of Diseases 9th edition codes 079.6, 466–469, 470.0, 480–482.8, 483.0–483.9, 484.1–484.2, 484.6–484.7, and 487–489, and International Classification of Diseases 10th edition codes A48.1, A70, B97.4–B97.6, J09–J15.8, J16–J16.9, J20–J21.9, J91.0, P23.0–P23.4, and U04–U04.9 (appendix 1 pp 81–84).

LRI mortality

The GBD Cause of Death database collates all available data from vital registration systems, surveillance systems, and verbal autopsy studies. Input data for LRI mortality estimation included 23 109 site-years (the number of years for which data are available for a particular location) of vital registration data, 825 site-years of sample vital registration data (ie, data covering a sample of the population), 1766 site-years of verbal autopsy data, and 681 site-years of mortality surveillance data. Country-specific data sources and citations are available on the Global Health Data Exchange. Vital registration data were adjusted for completeness and garbage coding.¹⁸ Data before and after garbage code redistribution are available in the online data visualisation tool.

We used the Cause of Death Ensemble modelling (CODEm) strategy¹⁹ to generate LRI mortality estimates by location, year, age, and sex. CODEm assesses a vast array of sub-models with varying combinations of predictive covariates (eg, undernutrition and air pollution) that are run through four model categories (ie, mixed-effects regression models and spatiotemporal Gaussian process regression models for cause fractions and mortality rates; appendix 1 pp 13–14). Sub-models are evaluated using out-of-sample predictive validity and combined into an ensemble with the best predictive performance.

LRI morbidity

To estimate age–sex-specific incidence and prevalence of LRIs, we used data identified via systematic reviews of the literature. Additionally, we used population-based survey data, claims data, and inpatient data to estimate incidence and prevalence (appendix 1 pp 6–10). For GBD 2019, we used an enhanced standardised approach, compared with previous GBD iterations, to adjust definitions in data sources that did not use our reference case definition to be comparable with our reference case definition (ie, clinician-diagnosed pneumonia or bronchiolitis). To do so, we first computed the ratio of the data based on alternative case definitions to the data based on the reference case definition, on the basis of all available data matched by location, year, age, and sex. We then ran a meta-regression to pool the ratios and used the pooled ratio to adjust the data based on alternative case definitions to the level of the data based on the reference case definition (appendix 1 pp 8–10).

Our inclusion criteria for scientific literature included a study duration of at least 1 year to avoid bias in the seasonal timing of LRIs and a sample size of at least 100 people (the sample size threshold was chosen arbitrarily). Survey data

were adjusted for seasonality by fitting a generalised additive mixed-effects model with a forced periodicity for each GBD region, accounting for the year of the survey and the case definition used. The percentage difference between the monthly model-fit LRI prevalence and the corresponding regional mean LRI prevalence was computed to adjust survey data by month and geography. The mean duration of LRIs was 7.79 days (uncertainty interval [UI] 6.20–9.64); this was determined on the basis of a systematic review and meta-analysis,¹⁰ and was used to convert incidence data to prevalence. We modelled these data together with LRI mortality estimates using DisMod-MR 2.1,¹¹ a Bayesian meta-regression tool that imposes coherence between data for different parameters, to produce final incidence and prevalence estimates. Details on the preparation of data sources and the modelling in DisMod can be found in appendix 1 (pp 6–11).

Risk factors

Detailed methods for GBD risk factor estimation have been published elsewhere.⁷ In summary, we first selected risk–outcome pairs (eg, LRIs attributable to smoking) on the basis of evidence of a convincing or probable causal relationship between the risk and the outcome. A full list of LRI risk factors and the mechanism through which each risk factor could cause LRIs can be found in appendix 1 (pp 15–17). The population attributable fractions (PAFs) of risk factors were quantified by estimating the risk factor exposure distributions and the relative risk of the association between each risk factor and the outcome, and determining the theoretical minimum-risk exposure level. The PAF is the fraction of LRI mortality that would have been reduced if the exposure to the risk factor had been at the theoretical minimum-risk exposure level. The attributable burden was computed by multiplying the location–year–age–sex-specific PAFs of risk factors by corresponding LRI deaths. We also calculated risk-deleted mortality rates to represent the LRI mortality rate that would have been observed had the risk factors been set to their corresponding theoretical minimum-risk exposure levels. Full details of the methods used for estimating each of the 14 LRI risk factors are provided in appendix 1 (pp 18–79).

Uncertainty intervals and age-standardisation

We computed 95% UIs based on 1000 draws from the posterior distribution of each stage in the estimation process using the 2.5th and 97.5th percentiles of the 1000 ordered values.

We used the GBD world population age standard²² to calculate age-standardised LRI incidence and mortality rates.

Role of the funding source

The funder of the study had no role in study design, data collection, data analysis, data interpretation, or writing of the report.

See Online for appendix 1

For the online data visualisation tool see <https://vizhub.healthdata.org/cod/>

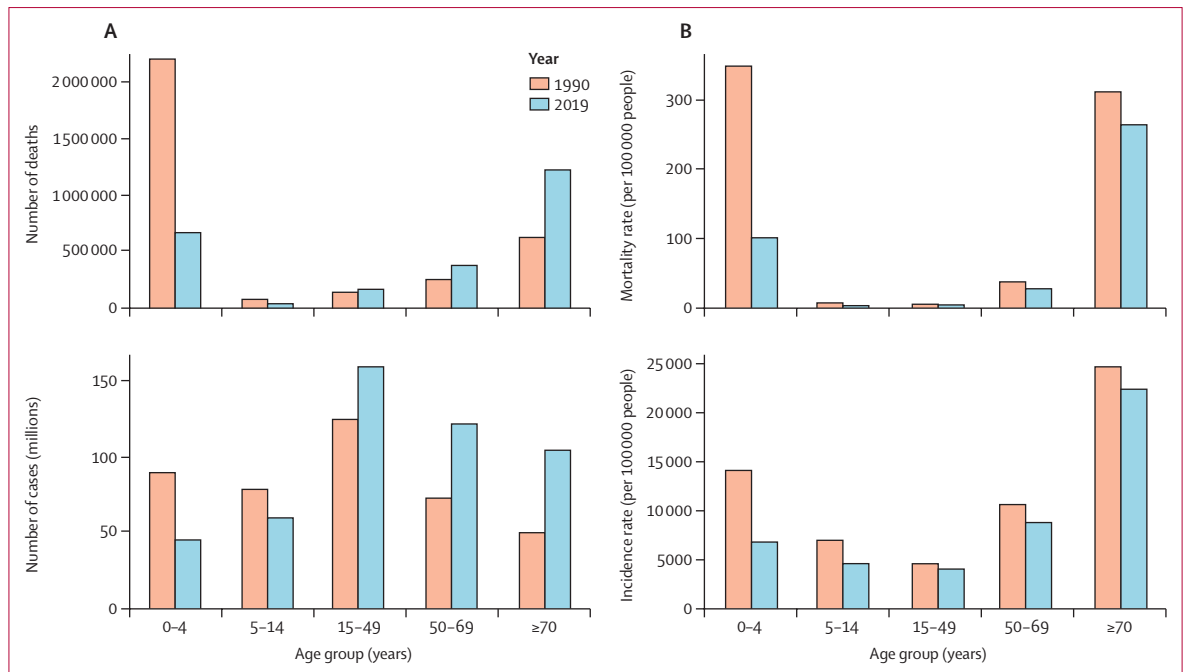


Figure 1: Incidence and mortality counts (A) and rates (B) due to lower respiratory infections for both sexes combined in 1990 and 2019, by age group

Results

On a global scale, in 2019, the total number of LRI incident episodes was 257 million (95% UI 240–275) for males and 232 million (217–248) for females, reflecting an increase of 20.0% (95% UI 15.8–24.5) for males and 15.8% (11.9–19.7) for females since 1990 (appendix 2 pp 3, 5). The age-standardised incidence rate was 1.17 (95% UI 1.16–1.18) times greater in males than in females in 2019. When looking at specific age–sex groups, we estimated that there was a decrease in LRI episodes between 1990 and 2019 in children younger than 15 years and an increase in this period in all adult age groups (figure 1; appendix 2 pp 3, 34). Among children, the decrease varied from 20.8% (95% UI 16.2–25.6) among males aged 5–14 years to 49.9% (48.5–51.6) among females younger than 5 years. Among adult age groups, the increase varied from 26.1% (23.2–29.1) for females aged 15–49 years to 126.0% (121.4–131.1) for males aged 70 years and older during the same period (appendix 2 p 34).

Between 1990 and 2019, children younger than 5 years saw the greatest improvement, with a decrease in incidence rate per 100 000 population of 51.7% (95% UI 50.0–53.5) for males and 52.1% (50.7–53.7) for females. Other age groups did not show similar improvements (figure 2; appendix 2 pp 34–37). We estimated only an 8.6% (6.6–10.5) decrease in incidence rate for males aged 70 years and older and only an 11.2% (9.2–13.0) decrease in incidence rate for females aged 70 years and older between 1990 and 2019.

In 2019, we estimated that individuals aged 15–49 years had the lowest global incidence rate of LRI episodes per 100 000 population among all age groups: 4128.1

(95% UI 3726.8–4583.5) for males and 3944.6 episodes (3541.5–4421.4) for females (figures 1, 2; appendix pp 34–37). Individuals aged 70 years and older, on the other hand, had the highest incidence rate per 100 000 population of all age groups: 25786.6 (23 182.5–28 975.4) for males and 19819.9 (17 921.3–22 072.6) for females. Of all super-regions, South Asia had the highest incidence rate per 100 000 population among both males aged 70 years and older (48 185.3 [95% UI 42 327.6–56 191.8]) and females aged 70 years and older (38 852.6 [34 264.3–44 606.1]; appendix 2 pp 34–37).

Globally, in terms of absolute numbers, LRIs accounted for 1.30 million (95% UI 1.18–1.42) deaths in 2019 among males and 1.20 million (1.07–1.33) deaths among females (appendix 2 p 38). The age-standardised mortality rate was 1.31 (95% UI 1.23–1.41) times greater in males than in females in 2019. We estimated an increase in LRI deaths among all adult age groups between 1990 and 2019 (figure 1; appendix 2 p 4), with males aged 70 years and older having the highest increase in deaths (100.0% [95% UI 83.4–115.9]; table). In high-income countries, we estimated a 70.7% (95% UI 58.3–77.9) increase in death counts for males aged 70 years and older and a 54.3% (39.7–63.0) increase for females aged 70 years and older (table). This increase in the number of deaths between 1990 and 2019 is visible regardless of age, with 20 countries showing an increase of more than 100% in death counts attributable to LRIs for males and 19 countries for females (appendix 2 pp 38–67).

Between 1990 and 2019, children younger than 5 years showed the greatest improvement in death rates for LRIs

See Online for appendix 2

(figures 1, 2), with a decrease in mortality rate per 100 000 people of 72.1% (95% UI 63.7–78.4) for males and a decrease of 69.7% (61.7–76.9) for females (table). Despite this finding, in 2019, there were still 672 000 LRI deaths (95% UI 551 000–826 000) in children younger than 5 years for both sexes combined (table; figure 1). Mortality rates for individuals aged 70 years and older decreased at a much slower pace globally over the same period (figure 2B); we estimated a 19.1% (95% UI 12.7–25.8) decrease in mortality rate for males and a 12.7% (5.6–20.6) decrease in mortality rate for females (table).

In 2019, global mortality rates due to LRIs were highest in individuals aged 70 years and older; the mortality rate per 100 000 was 294.8 (95% UI 262.3–317.8) for males and 241.2 (202.6–266.2) for females (table; figures 1, 2B). In contrast, children aged 5–14 years had the lowest mortality rates, with a mortality rate per 100 000 people of only 3.3 (2.8–3.9) for males and 3.3 (2.7–3.9) for females. We estimated that sub-Saharan Africa was the super-region with the highest mortality rate in individuals aged 70 years and older, with a mortality rate per 100 000 people of 850.8 (758.1–941.4) for males and 672.1 (553.9–766.2) for females (table).

Globally, in 2019, we estimated that 876 000 LRI deaths (95% UI 770 000–987 000) among males (PAF 67.6% [95% UI 62.9–72.1]) and 725 000 deaths (95% UI 629 000–826 000) among females (PAF 60.6% [95% UI 55.6–65.6]) were attributable to all evaluated LRI risk factors (appendix 2 p 68). Globally, the number of LRI deaths attributable to all risk factors decreased by 41.4% (95% UI 32.0–49.3) for males and 44.5% (34.9–53.9) for females between 1990 and 2019. Children younger than 5 years had the greatest percentage decrease in number of deaths and mortality rate attributable to all risk factors between 1990 and 2019 (appendix 2 p 97). The greatest percentage increase in attributable deaths between 1990 and 2019 was estimated to be in males aged 70 years and older (66.6% [95% UI 50.4–82.8]). Global age-standardised attributable mortality rate per 100 000 population due to all risk factors in 2019 was 26.2 (95% UI 23.1–29.5) for males and 19.4 (16.8–22.2) for females. Between 1990 and 2019, this rate decreased by 56.1% (95% UI 50.0–60.9) for males and 59.1% (52.5–65.6) for females (appendix 2 p 68).

In 2019, the leading risk factor for LRI mortality in children younger than 5 years was child wasting, in both males (PAF 53.0% [95% UI 37.7–61.8]) and females (56.4% [40.7–65.1]; figure 3; appendix p 117). Child wasting was also the largest risk factor in children younger than 5 years in 1990 and had decreased only slightly by 2019 (figure 3). For children younger than 5 years in 2019, the second largest PAF was for household air pollution; male and female children in this age group had near identical PAFs (31.4% [95% UI 21.5–41.5] vs 31.2% [21.3–41.5]; appendix p 117). Household air pollution was the second largest PAF in 1990 and had decreased substantially by 2019 (figure 3).

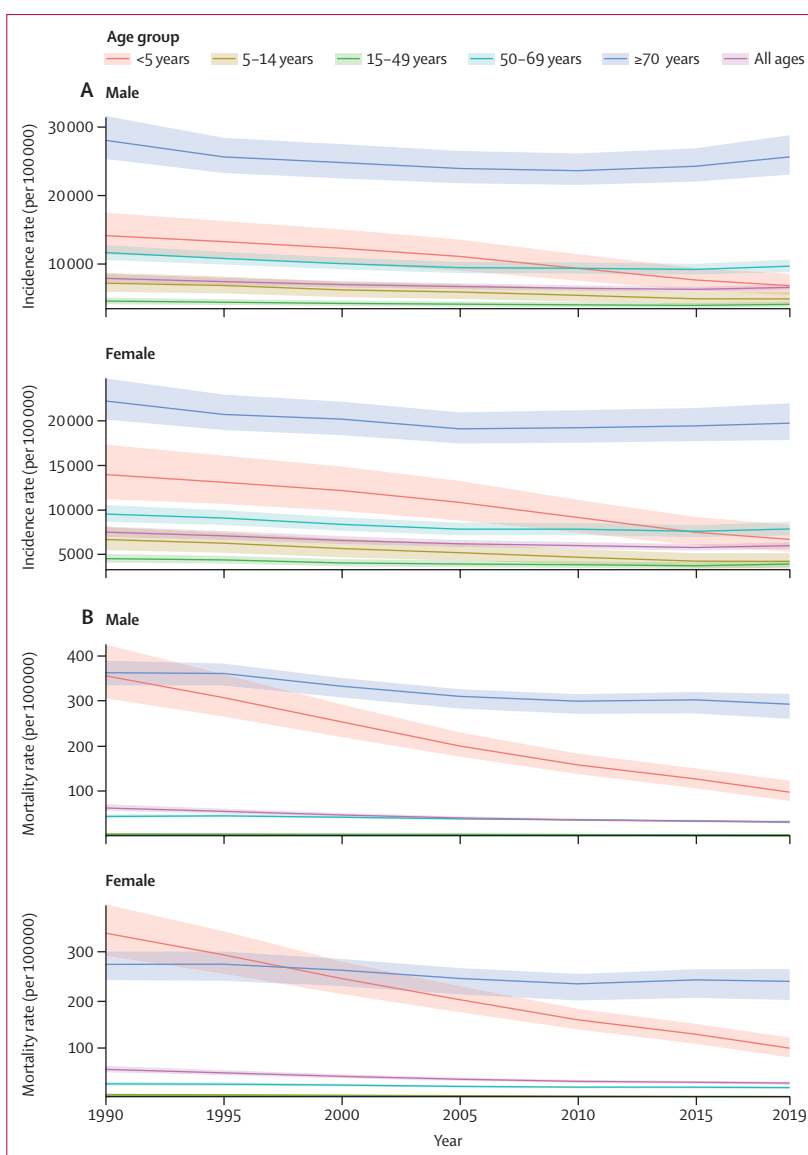


Figure 2: Global time trend of lower respiratory infection incidence rates (A) and mortality rates (B) by age and sex, 1990–2019

In 2019, the largest risk factor for children aged 5–14 years was household air pollution (PAF 26.0% [95% UI 16.6–35.5] for males and 25.8% [16.3–35.4] for females; figure 4; appendix p 134). For males aged 15–49 years, the risk factor with the highest PAF in 2019 was smoking (20.4% [15.4–25.2]); the risk factor with the lowest effect on the same group was high temperature (3.0% [1.3–6.9]; appendix p 147). These findings differ from findings in males aged 15–49 years in 1990, for whom the largest risk factor was household air pollution (29.0% [20.0–37.9]; figure 4). In 2019, females aged 15–49 years had the highest PAF from household air pollution (21.1% [14.5–27.9]), and the lowest PAF from alcohol use (2.0% [0.6–3.4]; figure 4; appendix p 147).

	Male				Female			
	2019		Percentage change from 1990 to 2019		2019		Percentage change from 1990 to 2019	
	Number of deaths*	Mortality rate (per 100 000 people)	Number of deaths	Mortality rate (per 100 000 people)	Number of deaths*	Mortality rate (per 100 000 people)	Number of deaths	Mortality rate (per 100 000 people)
Global								
0-4 years	341 000 (275 000 to 428 000)	99.7 (80.2 to 124.9)	-70.7% (-77.2 to -61.8)	-72.1% (-78.4 to -63.7)	331 000 (270 000 to 401 000)	103.2 (84.2 to 125.0)	-68.4% (-75.9 to -60.0)	-69.7% (-76.9 to -61.7)
5-14 years	21 800 (18 500 to 25 900)	3.3 (2.8 to 3.9)	-45.0% (-54.2 to -30.9)	-52.8% (-60.6 to -40.6)	20 500 (16 700 to 24 700)	3.3 (2.7 to 3.9)	-50.2% (-59.8 to -39.7)	-56.6% (-65.0 to -47.5)
15-49 years	105 000 (95 000 to 116 000)	5.3 (4.8 to 5.8)	27.5% (14.1 to 43.7)	-11.9% (-21.2 to -0.7)	65 200 (56 500 to 74 400)	3.4 (2.9 to 3.8)	5.7% (-7.3 to 21.1)	-27.3% (-36.3 to -16.7)
50-69 years	228 000 (210 000 to 248 000)	33.8 (31.1 to 36.7)	47.7% (32.7 to 63.9)	-26.5% (-34.0 to -18.4)	153 000 (132 000 to 171 000)	21.8 (18.8 to 24.3)	50.1% (31.7 to 71.4)	-26.2% (-35.2 to -15.7)
≥70 years	600 000 (534 000 to 646 000)	294.8 (262.3 to 317.8)	100.0% (83.4 to 115.9)	-19.1% (-25.8 to -12.7)	628 000 (527 000 to 693 000)	241.2 (202.6 to 266.2)	90.5% (73.3 to 105.8)	-12.7% (-20.6 to -5.6)
Central Europe, eastern Europe, and central Asia								
0-4 years	7800 (6300 to 9700)	54.9 (44.3 to 68.1)	-78.1% (-82.8 to -72.4)	-72.0% (-78.0 to -64.8)	6200 (5100 to 7600)	46.5 (38.0 to 57.2)	-78.2% (-82.6 to -72.7)	-71.5% (-77.2 to -64.3)
5-14 years	800 (700 to 900)	3.0 (2.6 to 3.3)	-42.2% (-49.6 to -33.9)	-26.3% (-35.8 to -15.8)	700 (600 to 800)	2.6 (2.3 to 3.0)	-46.5% (-52.9 to -37.4)	-30.1% (-38.4 to -18.2)
15-49 years	13 100 (11 600 to 14 700)	13.1 (11.6 to 14.7)	83.4% (62.8 to 105.7)	87.4% (66.4 to 110.3)	4400 (3900 to 5100)	4.4 (3.9 to 5.1)	49.3% (30.0 to 71.9)	53.6% (33.8 to 76.8)
50-69 years	20 000 (17 700 to 22 400)	44.8 (39.6 to 50.2)	72.2% (52.9 to 92.2)	40.3% (24.5 to 56.6)	6600 (5900 to 7300)	12.0 (10.9 to 13.3)	38.3% (24.4 to 53.3)	19.3% (7.3 to 32.2)
≥70 years	18 700 (16 600 to 20 400)	147.3 (130.4 to 160.7)	69.8% (54.8 to 84.7)	0.8% (-8.1 to 9.7)	21 500 (18 700 to 23 600)	87.8 (76.3 to 96.2)	35.4% (24.6 to 45.9)	-2.5% (-10.2 to 5.1)
High-income								
0-4 years	900 (700 to 1000)	3.1 (2.6 to 3.6)	-74.5% (-78.8 to -69.6)	-72.5% (-77.1 to -67.2)	700 (600 to 800)	2.5 (2.1 to 2.8)	-75.3% (-79.2 to -71.4)	-73.4% (-77.6 to -69.2)
5-14 years	200 (200 to 200)	0.3 (0.3 to 0.3)	-64.4% (-67.6 to -58.9)	-63.3% (-66.5 to -57.5)	200 (200 to 200)	0.3 (0.3 to 0.3)	-62.2% (-66.1 to -53.7)	-61.0% (-65.0 to -52.2)
15-49 years	4500 (4300 to 4600)	1.8 (1.7 to 1.9)	-29.6% (-32.6 to -26.5)	-32.1% (-35.0 to -29.1)	2900 (2800 to 3000)	1.2 (1.2 to 1.3)	-15.1% (-19.0 to -11.2)	-17.7% (-21.5 to -13.9)
50-69 years	23 800 (22 900 to 24 700)	17.8 (17.2 to 18.5)	11.0% (6.8 to 15.5)	-30.8% (-33.4 to -27.9)	13 500 (12 900 to 13 900)	9.6 (9.3 to 10.0)	17.7% (13.2 to 22.2)	-22.4% (-25.4 to -19.4)
≥70 years	189 000 (164 000 to 202 000)	302.2 (263.3 to 323.4)	70.7% (58.3 to 77.9)	-21.9% (-27.5 to -18.5)	201 000 (160 000 to 223 000)	240.5 (191.9 to 266.7)	54.3% (39.7 to 63.0)	-12.2% (-20.5 to -7.2)
Latin America and Caribbean								
0-4 years	11 000 (8400 to 14 200)	44.9 (34.0 to 57.9)	-79.9% (-85.4 to -73.1)	-79.2% (-84.9 to -72.2)	8700 (6800 to 10900)	37.0 (28.8 to 46.4)	-81.4% (-86.4 to -75.6)	-80.5% (-85.7 to -74.4)
5-14 years	1000 (800 to 1200)	2.1 (1.7 to 2.4)	-60.3% (-67.4 to -53.1)	-61.7% (-68.5 to -54.7)	900 (800 to 1100)	2.0 (1.7 to 2.2)	-62.1% (-67.5 to -56.1)	-62.6% (-67.9 to -56.7)
15-49 years	10 200 (9200 to 11 300)	6.8 (6.1 to 7.5)	19.0% (7.6 to 32.1)	-24.3% (-31.6 to -15.9)	5800 (5100 to 6500)	3.7 (3.3 to 4.2)	5.1% (-7.9 to 19.0)	-32.5% (-40.8 to -23.5)
50-69 years	20 000 (18 100 to 22 100)	42.8 (38.8 to 47.2)	89.3% (72.4 to 110.8)	-25.1% (-31.8 to -16.6)	13 600 (12 400 to 14 900)	26.1 (23.7 to 28.6)	94.5% (75.6 to 114.8)	-27.0% (-34.1 to -19.3)
≥70 years	55 400 (47 700 to 60 800)	385.2 (331.5 to 422.7)	160.9% (139.7 to 183.4)	-8.6% (-16.1 to -0.8)	64 600 (53 500 to 71 600)	354.0 (293.4 to 392.7)	187.6% (161.9 to 210.2)	-7.2% (-15.5 to 0.1)

(Table continues on next page)

Males aged 50–69 years in 2019 had the highest PAF from smoking (30.5% [95% UI 24.1–36.9]) and the lowest PAF from high temperature (3.0% [1.3–6.4]; figure 4; appendix p 160). Females in the same age group had the highest PAF from household air pollution (18.2% [12.5–24.5]) and the lowest from alcohol use (1.8% [0.5–3.1]; figure 4; appendix p 160). In individuals aged 70 and older, males in 2019 had the highest PAF from smoking (21.9% [16.8–27.3]) and the lowest PAF from high temperature (2.3% [1.1–4.8]; figure 4; appendix

p 173). Females in this age group had the highest PAF from ambient particulate matter (11.7% [8.2–15.8]), and the lowest PAF from alcohol use (1.8% [0.5–3.2]; figure 4; appendix p 173). Differing from females in 2019, females in 1990 had the highest PAF from household air pollution (21.2% [15.4–27.6]), and the lowest PAF from high temperature (1.8% [0.7–6.2]; figure 4).

Ambient particulate matter tended to affect males and females similarly across all age ranges (figure 4). PAFs for low temperatures were higher for people aged

	Male				Female			
	2019		Percentage change from 1990 to 2019		2019		Percentage change from 1990 to 2019	
	Number of deaths*	Mortality rate (per 100 000 people)	Number of deaths	Mortality rate (per 100 000 people)	Number of deaths*	Mortality rate (per 100 000 people)	Number of deaths	Mortality rate (per 100 000 people)
(Continued from previous page)								
North Africa and Middle East								
0–4 years	15300 (11 800 to 19 500)	50.0 (38.6 to 63.6)	–80.3% (–86.6 to –72.9)	–82.5% (–88.1 to –75.9)	15 400 (12 000 to 19 100)	52.9 (41.3 to 65.9)	–80.1% (–85.6 to –73.6)	–82.2% (–87.1 to –76.3)
5–14 years	1800 (1400 to 2300)	3.0 (2.3 to 3.8)	–48.3% (–64.1 to –32.8)	–60.0% (–72.2 to –47.9)	1600 (1100 to 2100)	2.9 (2.0 to 3.7)	–52.0% (–63.9 to –37.4)	–62.5% (–71.7 to –51.0)
15–49 years	6500 (5400 to 7800)	3.7 (3.1 to 4.5)	68.0% (40.5 to 99.6)	–19.9% (–33.1 to –4.9)	4900 (3800 to 6000)	3.1 (2.4 to 3.8)	37.4% (11.3 to 65.5)	–31.5% (–44.6 to –17.6)
50–69 years	11 600 (9700 to 13 700)	28.2 (23.6 to 33.3)	89.7% (56.4 to 134.0)	–25.6% (–38.6 to –8.2)	7700 (5900 to 9400)	19.9 (15.3 to 24.4)	80.7% (46.1 to 144.3)	–28.4% (–42.1 to –3.2)
≥70 years	23 200 (20 100 to 26 700)	238.7 (206.3 to 274.0)	143.9% (111.4 to 195.5)	–8.2% (–20.4 to 11.2)	19 700 (16 500 to 23 100)	201.5 (168.1 to 236.0)	131.4% (92.0 to 204.4)	–9.7% (–25.1 to 18.8)
South Asia								
0–4 years	87 800 (69 700 to 111 000)	102.4 (81.3 to 129.1)	–76.1% (–82.3 to –67.2)	–76.6% (–82.7 to –67.9)	103 000 (82 700 to 126 000)	131.1 (105.1 to 160.7)	–69.9% (–78.2 to –60.3)	–70.2% (–78.4 to –60.6)
5–14 years	6600 (5000 to 8500)	3.6 (2.7 to 4.6)	–46.8% (–58.6 to –28.5)	–58.2% (–67.4 to –43.8)	7400 (5500 to 9400)	4.4 (3.3 to 5.5)	–58.4% (–68.1 to –46.6)	–67.3% (–74.9 to –58.0)
15–49 years	16 900 (13 800 to 20 700)	3.4 (2.8 to 4.2)	21.1% (–4.8 to 51.1)	–33.1% (–47.4 to –16.5)	15 000 (11 600 to 18 900)	3.1 (2.4 to 4.0)	7.1% (–18.5 to 37.4)	–42.8% (–56.5 to –26.7)
50–69 years	48 200 (38 600 to 58 300)	39.2 (31.4 to 47.5)	42.3% (10.4 to 79.0)	–33.9% (–48.7 to –16.8)	49 600 (38 000 to 62 000)	40.5 (31.1 to 50.6)	70.3% (26.4 to 126.0)	–29.9% (–47.9 to –6.9)
≥70 years	102 000 (83 800 to 122 000)	309.2 (253.6 to 368.1)	133.3% (84.1 to 187.3)	–20.9% (–37.6 to –2.6)	111 000 (86 700 to 138 000)	308.4 (240.1 to 382.4)	178.6% (107.1 to 271.0)	–17.4% (–38.6 to 10.0)
Southeast Asia, east Asia, and Oceania								
0–4 years	30 600 (25 800 to 35 800)	41.2 (34.7 to 48.2)	–90.2% (–92.5 to –87.6)	–87.6% (–90.4 to –84.3)	23 400 (19 900 to 27 200)	35.4 (30.0 to 41.1)	–91.6% (–93.4 to –89.3)	–89.1% (–91.4 to –86.1)
5–14 years	2200 (1900 to 2600)	1.6 (1.4 to 1.8)	–78.8% (–82.1 to –64.7)	–74.4% (–78.4 to –57.4)	1700 (1500 to 2000)	1.4 (1.2 to 1.6)	–79.1% (–82.6 to –70.0)	–73.4% (–77.8 to –61.7)
15–49 years	16 700 (14 800 to 19 000)	2.9 (2.6 to 3.3)	–20.2% (–33.4 to 4.5)	–33.5% (–44.4 to –12.9)	8400 (7200 to 10000)	1.5 (1.3 to 1.8)	–38.1% (–49.1 to –24.2)	–48.4% (–57.6 to –36.8)
50–69 years	47 900 (42 100 to 54 500)	19.4 (17.1 to 22.1)	42.2% (19.0 to 71.5)	–39.1% (–49.0 to –26.5)	25 000 (20 300 to 28 700)	10.0 (8.1 to 11.5)	10.7% (–6.7 to 32.0)	–54.9% (–62.0 to –46.2)
≥70 years	138 000 (122 000 to 154 000)	221.4 (195.1 to 246.0)	119.6% (89.6 to 162.2)	–23.1% (–33.6 to –8.2)	138 000 (112 000 to 158 000)	178.4 (144.8 to 203.4)	79.2% (55.9 to 110.8)	–33.5% (–42.2 to –21.8)
Sub-Saharan Africa								
0–4 years	188 000 (143 000 to 243 000)	224.1 (170.1 to 289.2)	–40.1% (–55.1 to –17.6)	–67.8% (–75.9 to –55.7)	173 000 (135 000 to 220 000)	211.5 (165.3 to 268.8)	–35.3% (–52.9 to –14.3)	–64.8% (–74.3 to –53.3)
5–14 years	9200 (7300 to 11 400)	6.3 (5.0 to 7.8)	2.4% (–21.6 to 42.8)	–52.7% (–63.8 to –34.1)	8000 (6100 to 10 100)	5.6 (4.2 to 7.0)	4.7% (–22.9 to 39.7)	–51.0% (–63.9 to –34.5)
15–49 years	36 800 (31 100 to 43 600)	14.6 (12.3 to 17.3)	73.2% (43.0 to 106.9)	–26.8% (–39.5 to –12.5)	23 700 (18 700 to 29 400)	8.9 (7.0 to 11.1)	27.8% (3.2 to 57.3)	–46.2% (–56.6 to –33.8)
50–69 years	56 700 (48 600 to 65 900)	139.6 (119.8 to 162.4)	52.3% (28.3 to 80.8)	–26.8% (–38.4 to –13.2)	37 500 (29 900 to 45 200)	84.2 (67.3 to 101.5)	62.6% (35.8 to 99.2)	–30.8% (–42.2 to –15.2)
≥70 years	73 100 (65 200 to 80 900)	850.8 (758.1 to 941.4)	79.7% (54.2 to 106.1)	–10.0% (–22.8 to 3.2)	71 500 (58 900 to 81 500)	672.1 (553.9 to 766.2)	101.5% (76.0 to 130.9)	–5.4% (–17.4 to 8.4)

95% uncertainty intervals are shown in parentheses. GBD=Global Burden of Diseases, Injuries, and Risk Factors Study. *Six-digit numbers are reported to the nearest 1000 deaths and all other numbers are reported to the nearest 100 deaths.

Table: Lower respiratory infection deaths and mortality rates in 2019 and the percentage change in deaths and mortality rates between 1990 and 2019 by age, sex, and GBD super-region

70 years and older than for the younger age ranges. In 2019, second-hand smoke produced differing patterns of effect between males and females: males aged 5–14 years had a higher PAF (7.9% [95% UI 4.5–11.5]) than other age categories, whereas females aged 15–49 years were estimated to have a slightly higher PAF

from second-hand smoke (10.1% [5.8–14.4]) than other age categories. Alcohol-use PAFs were much higher in males across all age categories than in females. Lastly, high temperature PAFs were consistent across both sexes and tended to show a greater effect on individuals in younger age categories.

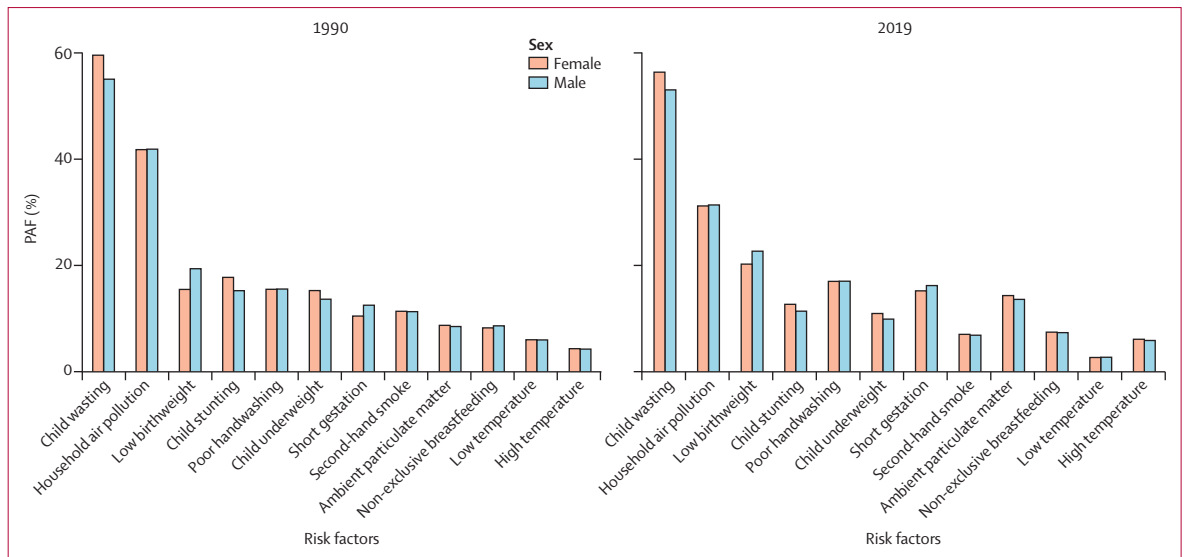


Figure 3: PAFs of lower respiratory infection deaths due to evaluated risk factors among males and females younger than 5 years in 1990 and 2019. Poor handwashing is defined as no access to a handwashing station with available soap and water. Non-exclusive breastfeeding is defined as the proportion of children under 6 months of age who are not exclusively breastfed. PAF=population attributable fraction.

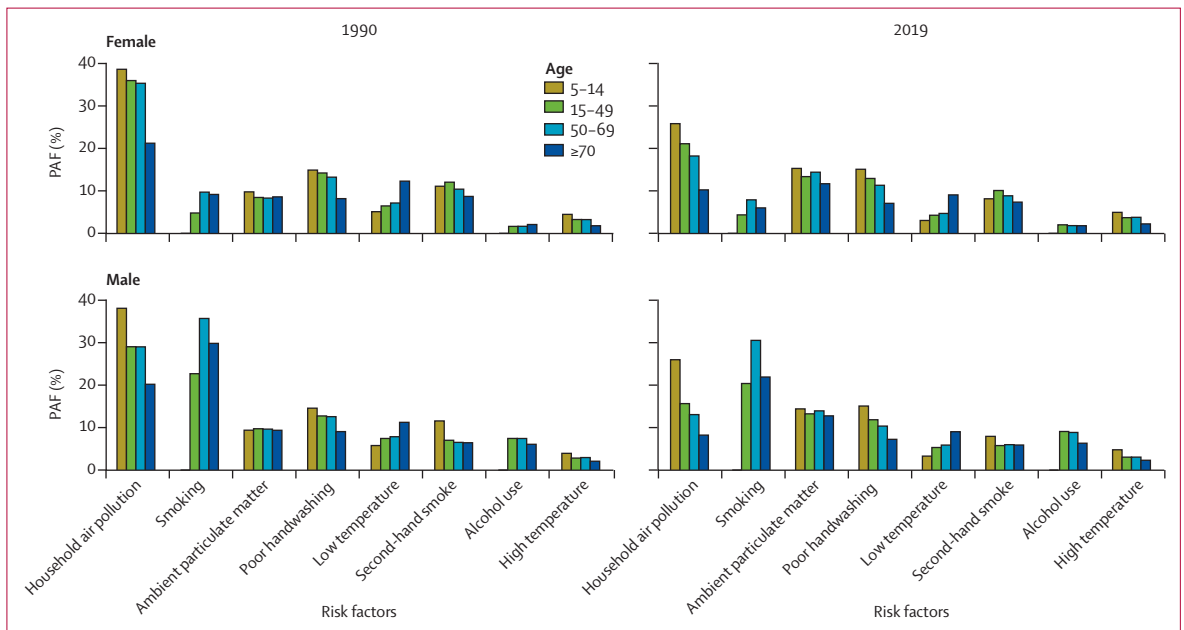


Figure 4: PAFs of lower respiratory infection deaths due to evaluated risk factors among males and females aged 5–14 years, 15–49 years, 50–69 years, and 70 years and older in 1990 and 2019. Poor handwashing is defined as no access to a handwashing station with available soap and water. Non-exclusive breastfeeding is defined as the proportion of children under 6 months of age who are not exclusively breastfed. PAF=population attributable fraction.

Mortality and incidence ratios were calculated between males and females across countries for the year 2019 (figure 5). Male-to-female ratios of age-standardised incidence rates were the highest in Ukraine and Moldova (figure 5A). Male-to-female ratios of age-standardised mortality rates were the highest in Russia, Belarus, Ukraine, Estonia, Lithuania, Japan, Ghana, and Moldova (figure 5B).

In 2019, the male-to-female ratio in global age-standardised mortality rates was 1.31 (95% UI 1.23–1.41). When all risk factors for LRIs were removed, the global age-standardised mortality rate per 100 000 population in 2019 in males was 13.5 (95% UI 11.4–15.7), and in females was 11.1 (9.2–12.7; appendix 2 p 102). Therefore, the ratio of risk-deleted mortality rates between males and females in 2019 was 1.22 (95% UI 1.12–1.36).

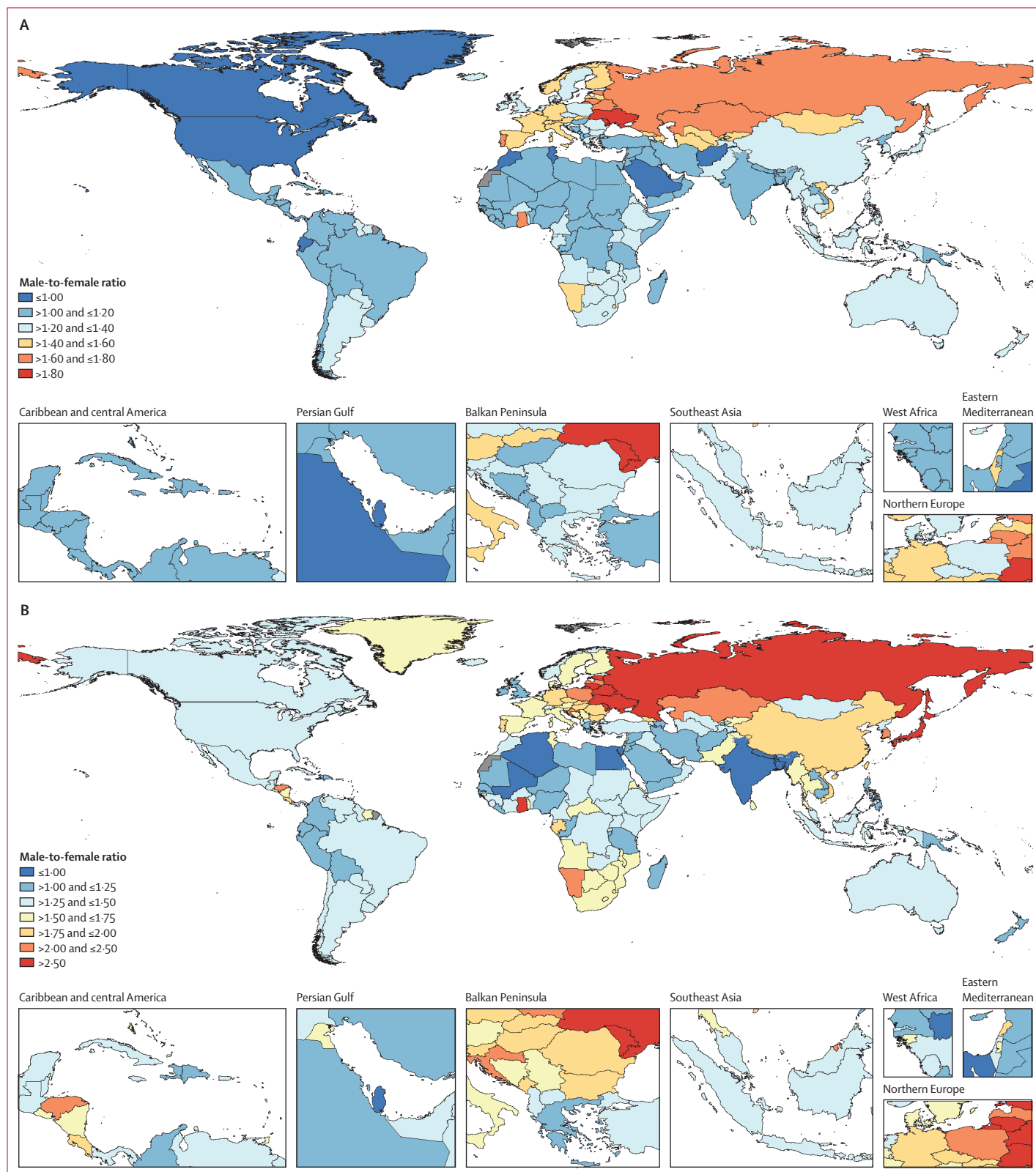


Figure 5: Male-to-female ratio of age-standardised lower respiratory infection incidence rates (A) and mortality rates (B), 2019

Comparatively, in 1990, the male-to-female ratio in global age-standardised mortality rates was 1.24 (95% UI 1.15–1.33). The global risk-deleted mortality rates per 100 000 population were 15.6 (95% UI 13.0–18.3) for males and 13.4 (11.2–15.7) for females (appendix p 102). The risk-deleted mortality rate ratio between males and females in 1990 was 1.17 (95% UI 1.07–1.28). In 2019, the central Europe, eastern Europe, and central Asia region had the largest risk-deleted male-to-female ratio of all regions at 1.62 (95% UI 1.43–1.83). South Asia had the lowest male-to-female ratio at 0.88 (95% UI 0.67–1.14), where females had a higher risk-deleted mortality rate than males (appendix p 110).

Discussion

Between 1990 and 2019, the greatest progress in reducing LRI incidence and mortality rates was observed in children younger than 5 years, indicating the success of initiatives targeting children in this age bracket. Much less progress has been made to reduce LRI incidence and mortality rates in adults, particularly in older age groups. Over the same period, in terms of absolute numbers, an increase in LRI incident episodes and deaths was estimated among all adult age groups due to population growth and aging. Globally, in 2019, age-standardised incidence rates were 1.2 times greater in males than in females and mortality rates were 1.3 times greater. Smoking was the leading risk factor for LRI mortality in adult males, responsible for about one-third of LRI deaths in those aged 50–69 years and one-fifth of LRI deaths in other age groups in 2019. PAFs of LRI deaths attributable to ambient particulate matter pollution have gone up for both males and females of all ages since 1990. On the other hand, PAFs of LRI deaths due to household air pollution have decreased among all age groups since 1990. PAFs of LRI deaths due to child wasting, stunting, and being underweight have also decreased among children younger than 5 years during the same period.

Despite the substantial progress made in children younger than 5 years, there were still 672 000 LRI deaths in this age group in 2019, and 93.5% (95% UI 90.4–95.7) of those deaths were attributable to preventable risk factors. Over the past two decades, global initiatives to combat wasting, the leading LRI risk factor, have focused mainly on treating wasting children, particularly in humanitarian situations.¹³ Although treatment coverage has steadily increased over time, only one-third of severely wasted children received treatment in 2019.¹³ This situation has been compounded by COVID-19 pandemic-related disruptions to nutrition and other fundamental services;¹⁴ the global prevalence of child wasting was estimated to increase by 14.3% during the first year of the pandemic.¹⁵ Given that undernutrition is the main risk factor not only for LRIs but also for other leading causes of death in children younger than 5 years such as diarrhoea and measles, long-term adverse

implications are foreseeable unless the recommended actions (eg, protecting and facilitating access to healthy, nutritious, and affordable food and reactivation and scaling up of early detection and treatment services for child wasting¹⁴) are taken promptly.

In 2015, the era of the Millennium Development Goals (MDGs) ended and the global community unanimously adopted the Sustainable Development Goals (SDGs). Although substantial progress had been made towards the MDG goal of reducing under-5 mortality by two-thirds between 1990 and 2015,¹⁶ achieving the new SDG target of 25 or fewer deaths per 1000 livebirths by 2030 would require promoting child survival by accelerating the decline of the major causes of death in young children. LRIs, which were still the leading infectious cause of death among children younger than 5 years in 2019, are largely preventable through vaccination and addressing key risk factors.^{10,17}

In contrast to the progress seen in children younger than five years, little has been achieved in reducing the LRI burden among adults, indicating a need for initiatives that address LRI risk factors in the adult age groups. Ambient particulate matter pollution was a leading risk factor for LRI mortality in all adult age groups in 2019. Studies from 2020 have also suggested associations between elevated exposures to particulate matter with a diameter less than 2.5 µm (PM_{2.5}) and higher COVID-19 cases and deaths.^{18–20} Contributors to global ambient particulate matter pollution include wildfires, biomass burning, sandstorms, chemical plants, and vehicle combustion sources.^{21–25} A study of how countries have followed the WHO ambient air quality guidelines found there were no air quality standards in 57 (34%) of the 170 countries examined.²⁶ The same study also found that air quality standards for some pollutants, including PM_{2.5}, were non-compliant with WHO guidelines in many countries.²⁶

Our results showed that global LRI deaths attributable to household air pollution decreased among all age groups between 1990 and 2019; however, exposure to household air pollution was responsible for more than a quarter of LRI deaths among children younger than 5 years and children aged 5–14 years, and more than a fifth of LRI deaths among women aged 15–49 years, in 2019. Sub-Saharan Africa had the largest PAFs, and South Asia had the second largest PAFs, for household air pollution across all age groups in 2019. More than 890 million people do not have access to clean cooking fuels in sub-Saharan Africa.²⁷ In India, the Pradhan Mantri Ujjwala Yojana, one of India's primary policies to provide households with liquid petroleum gas, a clean cooking fuel, was scheduled to be implemented in 102 cities and towns and related villages in 2019.²⁸ An evaluation study done in a rural community in Odisha found that the majority of Pradhan Mantri Ujjwala Yojana recipients did not refill their liquid petroleum gas cylinders (ie, solid fuels were still being used for cooking),

indicating the need for interventions to address challenges faced by rural households to ensure a complete transition from polluting to clean fuels.²⁹

Consistent with previous studies,⁶ we found higher LRI incidence and mortality among males than females, especially among adults. Potential reasons for this difference include sex differences in the immune response to infection and behavioural factors such as smoking and alcohol use.^{6,30} Females generally have a stronger immune system than males.³¹ Smoking is not only immunosuppressive but also causes changes such as ciliary dysfunction in the respiratory tract, leading to decreased pathogen clearance.³² The highest LRI mortality rates attributable to smoking among men were observed in countries in east Asia, southeast Asia, and eastern Europe. Despite a gradual decline in smoking prevalence in most of these countries, the declines were not sufficient to compensate for population growth, leading to a steady or growing number of smokers with time.³³ In many countries worldwide, progress towards reducing smoking prevalence has stalled in the past decade.^{7,33} The number of countries that have implemented at least one key intervention of the WHO Framework Convention on Tobacco Control has increased over time; however, only 62 countries had a complete ban on smoking in public and workplace settings, and only 23 countries provided comprehensive support for smokers seeking assistance in quitting smoking, as of 2018.³⁴

Results showed that PAFs attributable to alcohol use were much higher in males than females across all adult age categories. Alcohol use increases the risk of microbe aspiration and weakens the host immune system.³⁵ Although alcohol use is generally higher among men than women, it is increasing among women in different parts of the world, including some countries in sub-Saharan Africa.^{36–38} Increased government support and engagement are essential for adopting and enforcing effective alcohol policies in sub-Saharan Africa, which is a target region for alcohol companies to expand their market.³⁹

We found that the male-to-female ratio in global age-standardised LRI mortality rates decreased from 1.31 to 1.22 after removing the combined effects of all evaluated risk factors. Despite the smaller ratio, males still had a higher mortality rate than females, suggesting that other factors such as genetics and hormones could have a role in differential regulation of the immune system and the greater risk of mortality among males than females.³¹

Our results suggest that reducing the LRI burden and targeting the key risk factors that are different across age–sex groups will help in achieving multiple SDG targets, including SDG 3 (ensuring healthy lives and promoting wellbeing for all ages), SDG 7 (affordable and clean energy), and SDG-10 (reducing inequalities).⁴⁰ The remarkable progress made in children younger than 5 years was a result of the scale-up of proven interventions,

including vaccination and reducing exposure to known risk factors.¹⁰ Similar interventions for other age groups could contribute to the achievement of the SDG targets. Pneumococcal conjugate vaccines have been shown to have a direct protective effect on young children and an indirect protective effect on unvaccinated adults.⁴¹ The global pneumococcal conjugate vaccine coverage (third dose) among young children was estimated to be 47.9% (95% UI 47.0–48.9) in 2019.⁴² The gap in childhood immunisation coverage has become wider as the COVID-19 pandemic disrupted routine immunisation services worldwide, indicating an urgent need for catch-up and expansion of immunisation.^{43,44} Studies published since 2019 have shown that direct immunisation of older adults with PCV13 significantly reduced the disease burden.^{45–47} Immunisation of older adults, as well as addressing key leading risk factors such as child wasting, smoking, ambient particulate matter pollution, and household air pollution, could help reduce the burden of LRIs across all age groups. Additionally, supportive care, such as oxygen therapy, is a key part of the management of severe LRIs, and interventions to strengthen oxygen systems in low-resource settings could further help reduce LRI mortality.⁴⁸

This study has several limitations. One of the key limitations is the availability of data. In the absence of data for a particular country, estimates were dependent on the regional patterns, covariates, and out-of-sample predictive validity assessment. The absence of data in a given country translated into wide intervals of uncertainty. Even in countries with data, delays in data reporting prevented their timely integration into the GBD estimation. The most recent years for which cause of death data were available were 2016 and 2017. We were able to validate our estimation method by comparing two sets of estimates produced for a particular year with and without using any data for that year. For example, GBD 2016 produced LRI mortality estimates for 2016 using data available up to 2013 and 2014; these estimates were compared with GBD 2019 estimates for the same year that were informed by empirical data for 2016. GBD 2016⁴⁹ estimated a mortality rate per 100 000 population of 37.0 (95% UI 34.1–40.0) and GBD 2019 estimated 37.9 (32.5–40.8) for all ages and both sexes combined for the high-income super-region in 2016. Although the estimates are not identical, they are sufficiently close enough to support the validity of our approach. In this study, we were unable to evaluate the contribution of individual causes to the LRI burden. We plan to do a comprehensive assessment of the burden attributable to various pathogens in our future GBD estimation. Additionally, we have not assessed the LRI burden attributable to some potentially important risk factors such as overcrowding and incomplete immunisation.⁵⁰ Current risk–outcome pairs were included on the basis of the World Cancer Research Fund criteria for convincing or probable evidence. We could evaluate

whether additional risk factors are eligible for inclusion in future GBD iterations. Lastly, our current estimates of risk-attributable burden are limited by the quality of the primary data underlying the analysis. For example, data on some risk factors such as smoking and second-hand smoke were self-reported. Studies have indicated that self-reported smoking prevalence data might be prone to underestimation depending on respondents' perception of the social acceptance of smoking.^{51,52} Second-hand smoke exposure data might also be prone to recall bias.⁵³ Biomarker-based exposure assessment such as cotinine could help improve the accuracy of smoking and second-hand smoke data.^{52,53}

Although our results represent the LRI burden before the COVID-19 pandemic, the effect of the pandemic on LRIs needs to be investigated further. The pandemic was linked to a reduction of influenza and respiratory syncytial virus infections, probably as a result of mitigation measures, including mask wearing and social distancing.^{54,55} With the relaxation of measures, some countries started to see a rebound in influenza and respiratory syncytial virus infections in late 2020.^{56,57} As the data become more widely available, in future rounds of the GBD, we can quantify the indirect effects of the COVID-19 pandemic on the burden and causes of LRIs.

In conclusion, our results showed that despite an overall global decline in LRI incidence and mortality rates between 1990 and 2019, the pace of decline has been unequal across age groups. The observed progress in children younger than 5 years was clearly a result of targeted interventions, including improving vaccination and reducing exposure to risk factors. Similar interventions for other age groups could contribute to the achievement of multiple SDG targets, including promoting well-being at all ages and reducing health inequalities.

GBD 2019 Lower Respiratory Infection Collaborators

Hmwe Hmwe Kyu, Avina Vongpradith, Sarah Brooke Sirota, Amanda Novotney, Christopher E Troeger, Matthew C Doney, Rose G Bender, Jorge R Ledesma, Molly H Biehler, Samuel B Albertson, Joseph Jon Frostad, Katrin Burkart, Fiona B Bennitt, Jeff T Zhao, William M Gardner, Hailey Hagins, Dana Bryazka, Regina-Mae Villanueva Dominguez, Semagn Mekonnen Abate, Michael Abdelmasset, Amir Abdoli, Gholamreza Abdoli, Aidin Abedi, Vida Abedi, Tadesse M Abegaz, Hassan Abidi, Richard Gyan Aboagye, Hassan Abolhassan, Yonas Derso Abteu, Hiwa Abubaker Ali, Eman Abu-Gharbieh, Ahmed Abu-Zaid, Kidist Adamu, Isaac Yeboah Addo, Oyelola A Adegboye, Mohammad Adnan, Qorinah Estiningtyas Sakilah Adnani, Muhammad Sohail Afzal, Saira Afzal, Bright Opoku Ahinkorah, Aqeel Ahmad, Araz Ramazan Ahmad, Sajjad Ahmad, Ali Ahmadi, Sepideh Ahmadi, Haroon Ahmed, Jivan Qasim Ahmed, Tarik Ahmed Rashid, Mostafa Akbarzadeh-Khiavi, Hanadi Al Hamad, Luciana Albano, Mamoon A Aldeyab, Bezatu Mengistie Alemu, Keyfalew Addis Alene, Abdelazeem M Algammal, Fadwa Alhalaqa Najji Alhalaqa, Robert Kaba Alhassan, Beriwan Abdulqadir Ali, Liaqat Ali, Musa Mohammed Ali, Syed Shujait Ali, Yousef Alimohamadi, Wahid Alipour, Adel Al-Jumaily, Syed Mohamed Aljunid, Sami Almustanyir, Rajaa M Al-Raddadi, Rami H Hani Al-Rifai, Saif Aldeen S AlRyalat, Nelson Alvis-Guzman, Nelson J Alvis-Zakzuk, Edward Kwabena Ameyaw, Javad Javad Aminian Dehkordi, John H Amuasi, Dickson A Amugsi, Etsay Woldu Anbesu, Adnan Ansar,

Anayochukwu Edward Anyasodor, Jalal Arabloo, Demelash Areda, Ayele Mamo Argaw, Zeleke Gebru Argaw, Judie Arulappan, Raphael Taiwo Aruleba, Mulusew A Asemahagn, Seyyed Shamsadin Athari, Daniel Atlaw, Engi F Attia, Sameh Attia, Avinash Aujayeb, Tewachew Awoke, Tegegn Mulatu Ayana, Martin Amogre Ayanore, Sina Azadnajafabad, Mohammadreza Azangou-Khyavy, Samad Azari, Amirhossein Azari Jafari, Muhammad Badar, Ashish D Badiye, Nayereh Baghcheghi, Sara Bagherieh, Atif Amin Baig, Maciej Banach, Indrajit Banerjee, Mainak Bardhan, Francesco Barone-Adesi, Hiba Jawdat Barqawi, Amadou Barrow, Azadeh Bashiri, Quique Bassat, Abdul-Monim Mohammad Batiha, Abate Bekele Belachew, Melaku Ashagrie Belete, Uzma Iqbal Belgaumi, Akshaya Srikanth Bhagavathula, Nikha Bhardwaj, Pankaj Bhardwaj, Parth Bhatt, Vijayalakshmi S Bhojaraja, Zulfiqar A Bhutta, Soumitra S Bhuyan, Ali Bijani, Saeid Bitaraf, Belay Boda Abule Bodicha, Nikolay Ivanovich Briko, Danilo Buonsenso, Muhammad Hammad Butt, Jiao Cai, Paulo Camargos, Luis Alberto Cámera, Promit Ananyo Chakraborty, Muluken Genetu Chanie, Jaykaran Charan, Vijay Kumar Chattu, Patrick R Ching, Sungchul Choi, Yuen Yu Chong, Sonali Gajanan Choudhari, Enayet Karim Chowdhury, Devasahayam J Christopher, Dinh-Toi Chu, Natalie L Cobb, Aaron J Cohen, Natália Cruz-Martins, Omid Dadras, Fentaw Teshome Dagnaw, Xiaochen Dai, Lalit Dandona, Rakhi Dandona, An Thi Minh Dao, Sisay Abebe Debelo, Biniyam Demisse, Fitsum Wolde Demisse, Solomon Demissie, Diriba Dereje, Hardik Dineshbhai Desai, Abebaw Alemayehu Desta, Belay Desye, Sameer Dhingra, Nancy Diao, Daniel Diaz, Lankamo Ena Digesa, Linh Phuong Doan, Milad Dodangeh, Deepa Dongarwar, Fariba Dorostkar, Wendel Mombaqué dos Santos, Haneil Larson Dsouza, Eleonora Dubljanin, Oyewole Christopher Durojaiye, Hisham Atan Edinur, Elham Ehsani-Chimeh, Ebrahim Eini, Michael Ekhloenetale, Temitope Cyrus Ekundayo, Eman D El Desouky, Iman El Sayed, Maysaa El Sayed Zaki, Muhammed Elhadi, Ahmed Mahmoud Rabie Elkhapery, Amir Emami, Luchuo Engelbert Bain, Ryenchindorj Erkhembayar, Farshid Etaee, Mohamad Ezati Asar, Adeniyi Francis Fagbamigbe, Shahab Falahi, Aida Fallahzadeh, Anwar Faraj, Emerito Jose A Faraon, Ali Fatehizadeh, Pietro Ferrara, Allegra Allegra Ferrari, Getahun Fetensa, Florian Fischer, Joanne Flavel, Masoud Foroutan, Peter Andras Gaal, Abhay Motiramji Gaidhane, Santosh Gaihre, Nasrin Galehdar, Alberto L Garcia-Basteiro, Tushar Garg, Mesfin Damtew Gebrehiwot, Mathewos Alemu Gebremichael, Yibeltal Yismaw Gela, Belete Negese Belete Gemedo, Bradford D Gessner, Melaku Getachew, Asmare Getie, Seyyed-Hadi Ghamari, Mohammad Ghasemi Nour, Ahmad Ghashghaee, Ali Gholamrezaezhad, Abdolmajid Gholizadeh, Rakesh Ghosh, Sherief Ghozy, Pouya Goleij, Mohamad Golitaleb, Giuseppe Gorini, Alessandra C Goulart, Girma Garedeu Goyomsa, Habtamu Alganah Guadie, Zewdie Gudisa, Rashid Abdi Guled, Sapna Gupta, Veer Bala Gupta, Vivek Kumar Gupta, Alemu Guta, Parham Habibzadeh, Arvin Haj-Mirzaian, Rabih Halwani, Samer Hamidi, Md Abdul Hannan, Mehdi Harorani, Ahmed I Hasaballah, Hamidreza Hasani, Abbas M Hassan, Shokoufeh Hassani, Hossein Hassanian-Moghaddam, Hadi Hassankhani, Khezar Hayat, Behzad Heibati, Mohammad Heidari, Demisu Zembaba Heyi, Kamal Hezam, Ramesh Holla, Sung Hwi Hong, Nobuyuki Horita, Mohammad-Salar Hosseini, Mehdi Hosseinzadeh, Mihaela Hostiuc, Mowafa Househ, Soodabeh Hoveidamanesh, Junjie Huang, Nawfal R Hussein, Ivo Iavicoli, Segun Emmanuel Ibitoye, Kevin S Ikuta, Olayinka Stephen Ilesanmi, Irena M Ilic, Milena D Ilic, Mustapha Immurana, Nahlah Elkudssiah Ismail, Masao Iwagami, Jalil Jaafari, Elham Jamshidi, Sung-In Jang, Amirreza Javadi Mamaghani, Tahereh Javaheri, Fatemeh Javanmardi, Javad Javidnia, Sathish Kumar Jayapal, Umesh Jayarajah, Shubha Jayaram, Aelign Tasew Jema, Wonjeong Jeong, Jost B Jonas, Nitin Joseph, Farahnaz Joukar, Jacek Jerzy Jozwiak, Vaishali K, Zubair Kabir, Salah Eddine Oussama Kacimi, Vidya Kadashetti, Laleh R Kalankesh, Rohollah Kalhor, Ashwin Kamath, Bhushan Dattatray Kamble, Himal Kandel, Tesfaye K Kanko, Ibraheem M Karaye, André Karch, Samad Karkhah, Bekalu Getnet Kassa, Patrick DMC Katoto, Harkiran Kaur, Rimple Jeet Kaur, Leila Keikavoosi-Arani,

Mohammad Keykhaei, Yousef Saleh Khader, Himanshu Khajuria, Ejaz Ahmad Khan, Gulfaraz Khan, Imteyaz A Khan, Maseer Khan, Md Nuruzzaman Khan, Moien AB Khan, Yusra H Khan, Moawiah Mohammad Khatatbeh, Mina Khosravifar, Jagdish Khubchandani, Min Seo Kim, Ruth W Kimokoti, Adnan Kisa, Sezer Kisa, Niranjan Kisson, Luke D Knibbs, Sonali Kochhar, Farzad Kompani, Hamid Reza Koohestani, Vladimir Andreevich Korshunov, Soewarta Kosen, Parvaiz A Koul, Ai Koyanagi, Kewal Krishan, Barthelemy Kuate Defo, G Anil Kumar, Om P Kurmi, Ambily Kuttikkattu, Dharmesh Kumar Lal, Judit Lám, Iván Landires, Caterina Ledda, Sang-waong Lee, Miriam Levi, Sonia Lewycka, Gang Liu, Wei Liu, Rakesh Lodha, László Lorenzovici, Mojgan Lotfi, Joana A Loureiro, Farzan Madadzadeh, Ata Mahmoodpoor, Razzagh Mahmoudi, Marzieh Mahmoudimanes, Jamal Majidpoor, Alaa Makki, Elaheh Malakan Rad, Ahmad Azam Malik, Tauqeer Hussain Mallhi, Yosef Manla, Clara N Matei, Alexander G Mathioudakis, Richard James Maude, Entezar Mehrabi Nasab, Addisu Melese, Ziad A Memish, Oliver Mendoza-Cano, Alexios-Fotios A Mentis, Tuomo J Meretoja, Mehari Woldeariam Merid, Tomislav Mestrovic, Ana Carolina Micheletti Gomide Nogueira de Sá, Gelana Fekadu Worku Mijena, Le Huu Nhat Minh, Shabir Ahmad Mir, Reza Mirfakhraie, Seyedmohammadsadeq Mirmoenei, Agha Zeeshan Mirza, Moonis Mirza, Mohammad Mirza-Aghazadeh-Attari, Abay Sisay Misganaw, Awoke Temesgen Misganaw, Esmaeil Mohammadi, Mokhtar Mohammadi, Arif Mohammed, Shafiu Mohammed, Syam Mohan, Mohammad Mohseni, Nagabhishek Moka, Ali H Mokdad, Sara Momtazmanesh, Lorenzo Monasta, Md Moniruzzaman, Fateme Montazeri, Catrin E Moore, Abdolvahab Moradi, Lidia Morawska, Jonathan F Mosser, Ebrahim Mostafavi, Majid Motaghinejad, Haleh Mousavi Isfahani, Seyed Ali Mousavi-Aghdas, Sumaira Mubarik, Efrén Murillo-Zamora, Ghulam Mustafa, Sanjeev Nair, Tapas Sadasivan Nair, Houshang Najafi, Atta Abbas Naqvi, Sreenivas Narasimha Swamy, Zuhair S Natto, Biswa Prakash Nayak, Seyed Aria Nejadghaderi, Huy Van Nguyen Nguyen, Robina Khan Niazi, Antonio Tolentino Nogueira de Sá, Hasti Nouraei, Ali Nowroozi, Virginia Nuñez-Samudio, Chimezie Igwegbe Nzopotam, Ogochukwu Janet Nzopotam, Bogdan Oancea, Chimesduren Ochir, Oluwakemi Ololade Odukoya, Hassan Okati-Aliabad, Akinkunmi Paul Okekunle, Osaretin Christabel Okonji, Andrew T Olagunju, Isaac Iyinoluwa Olufadewa, Ahmed Omar Bali, Emad Omer, Eyal Oren, Erika Ota, Nikita Ostavnov, Abderrahim Oulhaj, Mahesh P A, Jagadish Rao Padubidri, Keyvan Pakshir, Reza Pakzad, Tamás Palicz, Anamika Pandey, Suman Pant, Shahina Pardhan, Eun-Cheol Park, Eun-Kee Park, Fatemeh Pashazadeh Kan, Rajan Paudel, Shrikant Pawar, Minjin Peng, Gavin Pereira, Simone Perna, Navaraj Perumalsamy, Ionela-Roxana Petcu, David M Pigott, Zahra Zahid Piracha, Vivek Podder, Roman V Polibin, Maarten J Postma, Hamid Pourasghari, Naeimeh Pourtaheri, Mirza Muhammad Fahd Qadir, Mathieu Raad, Mohammad Rabiee, Navid Rabiee, Saber Raeghi, Alireza Rafiei, Fakher Rahim, Mehran Rahimi, Vafa Rahimi-Movaghar, Azizur Rahman, Md Obaidur Rahman, Mosiur Rahman, Muhammad Aziz Rahman, Amir Masoud Rahmani, Vahid Rahmanian, Pradhun Ram, Kiana Ramezanzadeh, Jewel Rana, Priyanga Ranasinghe, Usha Rani, Sowmya J Rao, Sina Rashedi, Mohammad-Mahdi Rashidi, Azad Rasul, Zubair Ahmed Ratan, David Laith Rawaf, Salman Rawaf, Reza Rawassizadeh, Mohammad Sadeq Razeghinia, Elrashdy Moustafa Mohamed Redwan, Marissa B Reitsma, Andre M N Renzaho, Mohsen Rezaeian, Abanoub Riad, Reza Rikhtegar, Jefferson Antonio Buendia Rodriguez, Emma L B Rogowski, Luca Ronfani, Kristina E Rudd, Basema Saddik, Erfan Sadeghi, Umar Saeed, Azam Safary, Sher Zaman Safi, Maryam Sahebazzamani, Amirhossein Sahebkar, Sateesh Sakhamuri, Sana Salehi, Muhammad Salman, Hossein Samadi Kafil, Abdallah M Samy, Milena M Santric-Milicevic, Bruno Piassi Sao Jose, Maryam Sarkhosh, Brijesh Sathian, Monika Sawhney, Ganesh Kumar Saya, Abdul-Aziz Seidu, Allen Seylani, Amira A Shaheen, Masood Ali Shaikh, Elaheh Shaker, Hina Shamshad, Mequanent Melaku Sharew,

Asaad Sharhani, Azam Sharifi, Purva Sharma, Ali Sheidaei, Suchitra M Shenoy, Jeevan K Shetty, Damtew Solomon Shiferaw, Mika Shigematsu, Jae Il Shin, Hesamaddin Shirzad-Aski, K M Shivakumar, Siddharudha Shivalli, Parnian Shobeiri, Wudneh Simegn, Colin R Simpson, Harpreet Singh, Jasvinder A Singh, Paramdeep Singh, Samarjeet Singh Siwal, Valentin Yurievich Skryabin, Anna Aleksandrovna Skryabina, Mohammad Sadeq Soltani-Zangbar, Suhang Song, Yimeng Song, Prashant Sood, Chandrashekhar T Sreeramareddy, Paschalis Steiropoulos, Muhammad Suleman, Seyed-Amir Tabatabaeizadeh, Alireza Tahamtan, Majid Taheri, Moslem Taheri Soodejani, Elahe Taki, Iman M Talaat, Mircea Tampa, Sarmila Tandukar, Nathan Y Tat, Vivian Y Tat, Yibekal Manaye Tefera, Gebremaryam Temesgen, Mohamad-Hani Temsah, Azene Tesfaye, Degefa Gomora Tesfaye, Belay Tessema, Rekha Thapar, Jansje Henny Vera Ticoalu, Amir Tiyuri, Imad I Tleyjeh, Munkhsaikhan Togtmol, Marcos Roberto Tovani-Palome, Derara Girma Tufa, Irfan Ullah, Era Upadhyay, Sahel Valadan Tahbaz, Pascual R Valdez, Rohollah Valizadeh, Constantine Vardavas, Tommi Juhani Vasankari, Bay Vo, Linh Gia Vu, Birhanu Wagaye, Yasir Waheed, Yu Wang, Abdul Waris, T Eoin West, Nuwan Darshana Wickramasinghe, Xiaoyue Xu, Sajad Yaghoubi, Gahin Abdulraheem Tayib Yahya, Seyed Hossein Yahyazadeh Jabbari, Dong Keon Yon, Naohiro Yonemoto, Burhan Abdullah Zaman, Alireza Zandifar, Moein Zangiabadian, Heather J Zar, Iman Zare, Zahra Zarehshahabadi, Armin Zarrintan, Mikhail Sergeevich Zastrozhin, Wu Zeng, Mengxi Zhang, Zhi-Jiang Zhang, Chenwen Zhong, Mohammad Zoladl, Alimuddin Zumla, Stephen S Lim, Theo Vos, Mohsen Naghavi, Michael Brauer, Simon I Hay, Christopher J L Murray.

Affiliations

Institute for Health Metrics and Evaluation (H H Kyu PhD, A Vongpradith BA, S B Sirota MA, A Novotney MPH, C E Troeger MPH, R G Bender BS, J R Ledesma BA, S B Albertson BS, J J Frostad MPH, K Burkart PhD, F B Bennitt BA, J T Zhao BA, W M Gardner AB, H Hagins MSPH, D Bryazka BA, R V Dominguez BS, A J Cohen DSc, X Dai PhD, Prof L Dandona MD, Prof R Dandona PhD, K S Ikuta MD, T Mestrovic PhD, A H Mokdad PhD, J F Mosser MD, D M Pigott PhD, M B Reitsma BS, E L B Rogowski BA, Prof S S Lim PhD, Prof T Vos PhD, Prof M Naghavi PhD, Prof M Brauer DSc, Prof S I Hay FMedSci, Prof C J L Murray DPhil), Department of Health Metrics Sciences, School of Medicine (H H Kyu PhD, K Burkart PhD, X Dai PhD, Prof R Dandona PhD, A T Misganaw PhD, A H Mokdad PhD, D M Pigott PhD, Prof S S Lim PhD, Prof T Vos PhD, Prof M Naghavi PhD, Prof S I Hay FMedSci, Prof C J L Murray DPhil), Division of Pulmonary, Critical Care, and Sleep Medicine (E F Attia MD, N L Cobb MD), Division of Allergy and Infectious Diseases (K S Ikuta MD), Department of Global Health (S Kochhar MD), Department of Medicine (T E West MD), University of Washington, Seattle, WA, USA (Prof E Oren PhD); Urban Indian Health Institute, Seattle Indian Health Board, Seattle, WA, USA (M C Dooxey MPH); Division of Data, Analytics, and Delivery for Impact (M H Biehl MPH), Health Workforce Department (T S Nair MD), World Health Organization, Geneva, Switzerland; Department of Anesthesiology, Dilla University, Addis Ababa, Ethiopia (S M Abate MSc); Department of Surgery, Marshall University, Huntington, WV, USA (M Abdelmasseh MD); Zoonoses Research Center (A Abdoli PhD), Department of Community Medicine (V Rahmanian PhD), Jahrom University of Medical Sciences, Jahrom, Iran; Department of Epidemiology (G Abdoli PhD), Students' Research Committee (M Khosravifar MD), Department of Physiology (H Najafi PhD), Kermanshah University of Medical Sciences, Kermanshah, Iran; Department of Neurosurgery (A Abedi MD), Keck School of Medicine (A Abedi MD), Department of Radiology (A Gholamrezaezhad MD), Mark and Mary Stevens Neuroimaging and Informatics Institute, University of Southern California, Los Angeles, CA, USA (S Salehi MD); Department of Molecular and Functional Genomics, Geisinger Health System, Danville, PA, USA (V Abedi PhD); Biocomplexity Institute, Virginia Tech, Blacksburg, VA, USA (V Abedi PhD); Department of Clinical Pharmacy (T M Abegaz MS), Department of Surgical Nursing (A A Desta MSc), Human Physiology (Y Gela MSc), Institute of Public Health (M M Sharew MPH), Social and Administrative Pharmacy (W Simegn MSc), Department of Medical Microbiology

(B Tessema PhD), University of Gondar, Gondar, Ethiopia; Pharmacoeconomics and Health Outcomes Research Department, Florida A&M University, Tallahassee, FL, USA (T M Abegaz MS); Laboratory Technology Sciences Department (H Abidi PhD), Department of Nursing (M Zoladl PhD), Yasuj University of Medical Sciences, Yasuj, Iran; Department of Family and Community Health, School of Public Health, University of Health and Allied Sciences, Hohoe, Ghana (R G Aboagye MPH); Research Center for Immunodeficiencies (H Abolhassani PhD), Department of Epidemiology and Biostatistics (Y Alimohamadi PhD), Non-Communicable Diseases Research Center (S Azadnajafabad MD, M Azangou-Khyavy MD, S Ghamari MD, M Keykhaei MD, S Momtazmanesh MD, M Rashidi MD), National Institute for Health Research (E Ehsani-Chimeh PhD), School of Medicine (A Fallahzadeh MD, S Momtazmanesh MD, A Nowroozi BMedSc), The Institute of Pharmaceutical Sciences (TIPS) Tehran University of Medical Sciences (S Hassani PhD), Students' Scientific Research Center (M Keykhaei MD), Children's Medical Center (F Kompani MD), Pediatrics Department-Pediatric Cardiology (Prof E Malakan Rad MD), Tehran Heart Center (E Mehrabi Nasab MD), Faculty of Medicine (E Mohammadi MD, E Shaker MD, P Shobeiri MD), Metabolomics and Genomics Research Center (F Rahim PhD), Sina Trauma and Surgery Research Center (Prof V Rahimi-Movaghar MD), Department of Cardiology (S Rashedi MD), Department of Environmental Health Engineering (M Sarkhosh PhD), School of Public Health (A Sheidaei MSc), Department of Microbiology (E Taki PhD), Tehran University of Medical Sciences, Tehran, Iran; Department of Biosciences and Nutrition, Karolinska University Hospital, Huddinge, Sweden (H Abolhassani PhD); Biomedical Sciences Department (Y D Abtey MSc, B B A Bodicha MSc, T K Kanko MSc), Department of Public Health (Z G Argaw MPH), School of Nursing (T M Ayana MSc, B Demisse MSc), School of Public Health (A B Belachew MSc), Department of Midwifery (F W Demisse MSc, G Temesgen MSc), Department of Anatomy (S Demissie MSc), Department of Comprehensive Nursing (L E Digesa MSc), Department of Epidemiology and Biostatistics (M A Gebremichael MPH), Medical Laboratory (A Tesfaye MSc), Arba Minch University, Arba Minch, Ethiopia; University of Human Development (Prof H Abubaker Ali PhD), Department of Computer Science (M Hosseinzadeh PhD), Diplomacy and Public Relations Department (A Omar Bali PhD), University of Human Development, Sulaymaniyah, Iraq; Clinical Sciences Department (E Abu-Gharbieh PhD, H J Barqawi MPhil, Prof R Halwani PhD, Prof I M Talaat PhD), College of Medicine (Prof R Halwani PhD), Mass Communication Department (A Makki PhD), Sharjah Institute for Medical Research (B Saddik PhD), University of Sharjah, Sharjah, United Arab Emirates; Department of Surgery, Department of Obstetrics & Gynecology, Department of Medicine (A Abu-Zaid MD), College of Medicine (S Almustanyir MD, Prof Z A Memish MD), Alfaisal University, Riyadh, Saudi Arabia; College of Graduate Health Sciences, University of Tennessee, Memphis, TN, USA (A Abu-Zaid MD); Department of Health System Management (K Adamu MPH), Department of Medical Laboratory Science (M A Belete MSc), School of Public Health (M G Chanie MPH), Department of Environmental Health (M D Gebrehiwot PhD), College of Medicine and Health Sciences School of Public Health Department of Public Health Nutrition (B Wagaye MPH), Wollo University, Dessie, Ethiopia; Centre for Social Research in Health (I Y Addo PhD), School of Population Health (X Xu PhD), University of New South Wales, Sydney, NSW, Australia; Quality and Systems Performance Unit, Cancer Institute NSW, Sydney, NSW, Australia (I Y Addo PhD); Public Health and Tropical Medicine, James Cook University, Townsville, QLD, Australia (O A Adegbeye PhD); Department of Neonatology, Indiana University Health Ball Memorial Hospital, Muncie, IN, USA (M Adnan MD); Faculty of Medicine, Universitas Padjadjaran (Padjadjaran University), Bandung, Indonesia (Q E S Adnani PhD); Department of Life Sciences, University of Management and Technology, Lahore, Pakistan (M S Afzal PhD, I Ullah PhD); Department of Community Medicine, King Edward Memorial Hospital, Lahore, Pakistan (Prof S Afzal PhD); Department of Public Health, Public Health Institute, Lahore, Pakistan (Prof S Afzal PhD); The Australian Centre for Public and Population Health Research, University of Technology Sydney, Sydney, NSW, Australia (B O Ahinkorah MPH, E K Ameyaw MPhil); Department of Medical Biochemistry, College of Medicine, Shaqra University, Shaqra, Saudi Arabia (A Ahmad PhD); College of Nursing, International Relations & Diplomacy, Ranya - Al Sulaimaniyah, Iraq (A R Ahmad PhD); International Relations & Diplomacy (A R Ahmad PhD), School of Pharmacy (B A Ali PhD), Tishk International University, Erbil, Iraq; Department of Health and Biological Sciences, Abasyn University, Peshawar, Pakistan (S Ahmad PhD); Department of Epidemiology and Biostatistics (A Ahmadi PhD), Community-Oriented Nursing Midwifery Research Center (M Heidari PhD), Shahrekord University of Medical Sciences, Shahrekord, Iran; Department of Epidemiology (A Ahmadi PhD), School of Advanced Technologies in Medicine (S Ahmadi PhD), Social Determinants of Health Research Center (M Azangou-Khyavy MD, S Ghamari MD, Prof H Hassanian-Moghaddam MD, M Rashidi MD), Department of Pharmacology (A Haj-Mirzaian MD, K Ramezanzadeh PharmD), Obesity Research Center (A Haj-Mirzaian MD), Functional Neurosurgery Research Center (E Jamshidi PharmD), Department of Parasitology (A Javadi Mamaghani PhD), Department of Genetics (R Mirfakhraie PhD), School of Medicine (F Montazeri MD, S Nejadghaderi MD, M Zangjabadian MD), Chronic Respiratory Disease Research Center, National Research Institute of Tuberculosis and Lung Diseases (M Motaghinejad PhD), Medical Ethics and Law Research Center (M Taheri PhD), Shahid Beheshti University of Medical Sciences, Tehran, Iran; Department of Biosciences, COMSATS Institute of Information Technology, Islamabad, Pakistan (H Ahmed PhD); Department of Pathology and Microbiology (J Q Ahmed MSc, G A Yahya MSc), Department of Pharmacology, College of pharmacy (B A Zaman MSc), University of Duhok, Duhok, Iraq; Department of Computer Science and Engineering, University of Kurdistan Hewler, Erbil, Iraq (T Ahmed Rashid PhD); Liver and Gastrointestinal Diseases Research Center (M Akbarzadeh-Khiavi PhD), School of Nursing and Midwifery (H Hassankhani PhD), Student Research Committee (M Hosseini MD), Department of Parasitology (A Javadi Mamaghani PhD), Department of Medical Surgical Nursing (M Lotfi PhD), Medical Education Research Center (M Lotfi PhD), Anesthesiology and Critical Care (Prof A Mahmoodpoor MD), Department of Radiology (M Mirza-Aghazadeh-Attari MD, A Zarrintan MD), Tuberculosis and Lung Diseases Research Center (S Mousavi-Aghdas MD), Cardiovascular Research Center (M Rahimi MD), Connective Tissue Diseases Research Center (A Safari PhD), Drug Applied Research Center (H Samadi Kafil PhD), Department of Immunology (M Soltani-Zangbar MSc), Tabriz University of Medical Sciences, Tabriz, Iran; Geriatric and Long Term Care Department (H Al Hamad MD, B Sathian PhD), Rumailah Hospital (H Al Hamad MD), Hamad Medical Corporation, Doha, Qatar; Department of Experimental Medicine, University of Campania Luigi Vanvitelli, Naples, Italy (L Albano MD); Department of Pharmacy, University of Huddersfield, Huddersfield, UK (M A Aldeyab PhD); Faculty of Health Sciences (K A Alene MPH), School of Public Health (E K Chowdhury PhD), Curtin University, Perth, WA, Australia; Wesfarmers Centre of Vaccines and Infectious Diseases, Telethon Kids Institute, Perth, WA, Australia (K A Alene MPH); Department of Bacteriology, Immunology, and Mycology, Suez Canal University, Ismailia, Egypt (Prof A M Algammal PhD); Faculty of Nursing, Philadelphia University, Amman, Jordan (F A N Alhalaiqa PhD, Prof A M Batiha PhD); Psychological Sciences Association/Jordan, Amman, Jordan (F A N Alhalaiqa PhD); Institute of Health Research (R K Alhassan PhD, M Immurana PhD), Department of Health Policy Planning and Management (M A Ayanore PhD), University of Health and Allied Sciences, Ho, Ghana; Erbil Technical Health College, Erbil Polytechnic University, Erbil, Iraq (B A Ali PhD); Department of Biological Sciences, National University of Medical Sciences, Rawalpindi, Pakistan (L Ali PhD); School of Medical Laboratory, Module of Microbiology and Parasitology, Hawassa University, Hawassa, Ethiopia (M M Ali PhD); Center for Biotechnology and Microbiology, University of Swat, Swat, KPK, Pakistan (S S Ali PhD); Pars Advanced and Minimally Invasive Medical Manners Research Center (Y Alimohamadi PhD), Health Management and Economics Research Center (V Alipour PhD, J Arabloo PhD), Department of Health

Economics (V Alipour PhD), Hospital Management Research Center (S Azari PhD, H Pourasghari PhD), School of Medicine (M Dodangeh MD), Department of Medical Laboratory Sciences (F Dorostkar PhD), Health Services Management (H Mousavi Isfahani PhD), Trauma and Injury Research Center (M Taheri PhD), Department of Epidemiology and Biostatistics (A Tiyuri MSc), Department of Epidemiology (R Valizadeh PhD), Iran University of Medical Sciences, Tehran, Iran (F Pashazadeh Kan BSN); School of Computing, Mathematics and Engineering (Prof A Al-Jumaily PhD), Data Mining Research Unit (A Rahman PhD), Charles Sturt University, Wagga Wagga, NSW, Australia; ENSTA Bretagne's Information and Communication Sciences and Technologies Pole, Mathematics, Algorithms and Decision Team, ENSTA Bretagne, Brest, France (Prof A Al-Jumaily PhD); Department of Health Policy and Management, Kuwait University, Kuwait (Prof S M Aljunid PhD); International Centre for Casemix and Clinical Coding, National University of Malaysia, Bandar Tun Razak, Malaysia (Prof S M Aljunid PhD); Research & Innovation Center (Prof Z A Memish MD), Ministry of Health, Riyadh, Saudi Arabia (S Almustanyir MD); Department of Community Medicine (R M Al-Raddadi PhD), Rabigh Faculty of Medicine (A A Malik PhD), Department of Dental Public Health (Z S Natto DrPH), King Abdulaziz University, Jeddah, Saudi Arabia; Institute of Public Health, College of Medicine and Health Sciences, United Arab Emirates University, Abu Dhabi, Al Ain, United Arab Emirates (R H Al-Rifai PhD); Department of Special Surgery, The University of Jordan, Amman, Jordan (S S AlRyalat MD); Research Group in Hospital Management and Health Policies (Prof N Alvis-Guzman PhD), Department of Economic Sciences (N J Alvis-Zakzuk MSc), Universidad de la Costa (University of the Coast), Barranquilla, Colombia; Research Group in Health Economics, University of Cartagena, Cartagena, Colombia (Prof N Alvis-Guzman PhD); National Health Observatory, National Institute of Health, Bogota, Colombia (N J Alvis-Zakzuk MSc); Applied Science and Technology, University of California Berkeley, Berkeley, CA, USA (J J Aminian Dehkordi PhD); Chemical Engineering Department-Biotechnology group, Tarbiat Modares University, Tehran, Iran (J J Aminian Dehkordi PhD); Department of Global Health, Kwame Nkrumah University of Science and Technology, Kumasi, Ghana (J H Amuasi PhD); Global Health and Infectious Diseases, Kumasi Center for Collaborative Research in Tropical Medicine, Kumasi, Ghana (J H Amuasi PhD); Maternal and Child Wellbeing Department, African Population and Health Research Center, Nairobi, Kenya (D A Amugsi PhD); Department of Public Health, Samara University, Samara, Ethiopia (E W Anbesu MPH); School of Nursing and Midwifery, La Trobe University, Melbourne, VIC, Australia (A Ansar MPH, M Rahman PhD); Special Interest Group International Health, Public Health Association of Australia, Canberra, ACT, Australia (A Ansar MPH); School of Community Health, Charles Sturt University, Orange, NSW, Australia (A E Anyasodor PhD); College of Art and Science, Ottawa University, Surprise, AZ, USA (D Areda PhD); College of Liberal Arts and Sciences, Arizona State University, Tempe, AZ, USA (D Areda PhD); Department of Clinical Pharmacy (A M Argaw MSc), Department of Biomedical Science (D Atlaw MSc), Department of Anesthesia (Z Gudisa MSc), Department of Public Health (A Jema MPH), Department of Anatomy (D S Shiferaw MSc), Biomedical Department (D S Shiferaw MSc), Madda Walabu University, Goba, Ethiopia; Department of Maternal and Child Health, Sultan Qaboos University, Muscat, Oman (J Arulappan DSc); Department of Molecular and Cell Biology (R T Aruleba MSc), Department of Paediatrics & Child Health (Prof H J Zar PhD), University of Cape Town, Cape Town, South Africa; School of Public Health (M A Asemahagn PhD), Department of Medical Laboratory Sciences, College of Medicine and Health Sciences (T Awoke MSc, A Melese MSc), Health Systems Management (M G Chanie MPH), Department of Health Informatics (H A Guadie MPH), Bahir Dar University, Bahir Dar, Ethiopia; Department of Immunology, Zanjan University of Medical Sciences, Zanjan, Iran (S Athari PhD); Oral and Maxillofacial Surgery Department, Justus Liebig University of Giessen, Giessen, Germany (S Attia MSc); Northumbria HealthCare NHS Foundation Trust (A Aujayeb MBBS), National Health Service (NHS) Scotland, Newcastle upon Tyne, UK; Department of Health Economics, Centre for Health Policy Advocacy Innovation & Research in Africa, Accra, Ghana (M A Ayanore PhD); School of Medicine, Shahrood University of Medical Sciences, Shahrood, Iran (A Azari Jafari MD, S Mirmoeeni MD); Gomal Center of Biochemistry and Biotechnology, Gomal University, Dera Ismail Khan, Pakistan (M Badar PhD); Department of Forensic Science, Government Institute of Forensic Science, Nagpur, India (A D Badiye MSc); Department of Nursing, Saveh University of Medical Sciences, Saveh, Iran (N Baghcheghi PhD); School of Medicine (S Bagherieh BSc), Department of Environmental Health Engineering (A Fatehizadeh PhD), Health Services Management (M Mohseni PhD), Department of Biostatistics and Epidemiology (E Sadeghi PhD), Isfahan University of Medical Sciences, Isfahan, Iran; Unit of Biochemistry (A A Baig PhD), Universiti Sultan Zainal Abidin (Sultan Zainal Abidin University), Kuala Terengganu, Malaysia; Department of Hypertension, Medical University of Lodz, Lodz, Poland (Prof M Banach PhD); Polish Mothers' Memorial Hospital Research Institute, Lodz, Poland (Prof M Banach PhD); Department of Pharmacology, Sir Seewoosagur Ramgoolam Medical College, Mauritius, Belle Rive, Mauritius (I Banerjee MD); Department of Molecular Microbiology and Bacteriology, National Institute of Cholera and Enteric Diseases, Kolkata, India (M Bardhan MD); Department of Molecular Microbiology (M Bardhan MD), Indian Council of Medical Research, New Delhi, India (Prof L Dandona MD); Department of Translational Medicine, University of Eastern Piedmont, Novara, Italy (F Barone-Adesi PhD); Department of Public & Environmental Health, University of The Gambia, Brikama, The Gambia (A Barrow MPH); Epidemiology and Disease Control Unit, Ministry of Health, Kotu, The Gambia (A Barrow MPH); Health Information Management (A Bashiri PhD), Microbiology Department of Burn and Wound Healing Research Center (A Emami PhD), Research Center for Health Sciences, Institute of Health (P Habibzadeh MD), Burn and Wound Healing Research Center (F Javanmardi MSc), Department of Medical Mycology and Parasitology (H Nouraei MSc, Prof K Pakshir PhD, Z Zarehshahrabadi PhD), Shiraz University of Medical Sciences, Shiraz, Iran; Barcelona Institute for Global Health, University of Barcelona, Barcelona, Spain (Prof Q Bassat MD); Catalan Institution for Research and Advanced Studies, Barcelona, Spain (Prof Q Bassat MD, A Koyanagi MD); Center for Environmental and Respiratory Health Research, University of Oulu, Oulu, Finland (A B Belachew MSc, B Heibati PhD); Department of Oral Pathology and Microbiology (U I Belgaumi MD), Department of Oral Pathology and Microbiology, Forensic Odontology (V Kadashetti MDS), Krishna Institute of Medical Sciences Deemed to be University, Karad, India; Department of Social and Clinical Pharmacy, Charles University, Hradec Kralova, Czech Republic (A S Bhagavathula PharmD); Institute of Public Health (A S Bhagavathula PharmD), Department of Medical Microbiology & Immunology (Prof G Khan PhD), Family Medicine Department (M A Khan MSc), United Arab Emirates University, Al Ain, United Arab Emirates; Department of Anatomy (Prof N Bhardwaj MD), Department of Community Medicine and Family Medicine (P Bhardwaj MD), School of Public Health (P Bhardwaj MD), Department of Pharmacology (J Charan MD, R J Kaur PhD), All India Institute of Medical Sciences, Jodhpur, India; Department of Pediatrics, United Hospital Center, WVU Medicine, Bridgeport, WV, USA (P Bhatt MD); Department of Anatomy (V S Bhojaraja MD), Department of Biochemistry (J K Shetty MD), Royal College of Surgeons in Ireland Medical University of Bahrain, Busaiteen, Bahrain; Centre for Global Child Health, Aga Khan University, Karachi, Pakistan (Prof Z A Bhutta PhD), University of Toronto, Toronto, ON, Canada; Centre of Excellence in Women & Child Health (Prof Z A Bhutta PhD); Health Administration (S S Bhuyan PhD), Department of Pediatrics (I A Khan MD), Rutgers University, New Brunswick, NJ, USA; Social Determinants of Health Research Center, Babol University of Medical Sciences, Babol, Iran (A Bijani PhD); Department of Biostatistics and Epidemiology (Prof S Bitaraf PhD), Department of Orthodontics (E Eini DDS), Department of Epidemiology (A Sharhani PhD), Ahvaz Jundishapur University of Medical Sciences, Ahvaz, Iran; Department of Epidemiology and Evidence-Based Medicine, IM Sechenov First Moscow State Medical University, Moscow, Russia (Prof N I Briko DSc, V A Korshunov PhD, R V Polibin PhD); Department of Woman and Child Health and Public Health, Agostino

Gemelli University Polyclinic IRCCS, Roma, Italy (D Buonsenso MD); Global Health Research Institute, Catholic University of Sacred Heart, Roma, Italy (D Buonsenso MD); Faculty of Pharmacy, University of Central Punjab, Lahore, Pakistan (M Butt MS); Institute for Health and Environment, Chongqing University of Science and Technology, Chongqing, China (J Cai MSc, W Liu PhD); Department of Pediatrics (Prof P Camargos PhD), Department of Maternal and Child Nursing and Public Health (Prof A C Micheletti Gomide Nogueira de Sá MSc), Departamento de Clínica Médica (A T Nogueira de Sá MSc), Department of Infectious Diseases and Tropical Medicine (B P Sao Jose PhD), Federal University of Minas Gerais, Belo Horizonte, Brazil; Internal Medicine Department, Italian Hospital of Buenos Aires, Buenos Aires, Argentina (Prof L A Cámara MD); Board of Directors (Prof L A Cámara MD), Argentine Society of Medicine, Buenos Aires, Argentina (Prof P R Valdez MD); School of Population and Public Health (P A Chakraborty MPH, Prof M Brauer DSc), Department of Pediatrics (Prof N Kisson MD), University of British Columbia, Vancouver, BC, Canada; Department of Community Medicine, Datta Meghe Institute of Medical Sciences, Sawangi, India (V Chattu MD); Saveetha Medical College, Saveetha University, Chennai, India (V Chattu MD); Division of Infectious Diseases, Department of Medicine, Washington University in St Louis, St Louis, MO, USA (P R Ching MD); College of Medicine (S Choi, Prof J Shin MD), Department of Pediatrics (S Hong MD), Department of Public Health, Graduate School, Yonsei University (W Jeong PhD), Department of Preventive Medicine (Prof E Park PhD), Institute of Health Services Research (Prof E Park PhD), Yonsei University, Seoul, South Korea; The Nethersole School of Nursing (Y Chong PhD), Jockey Club School of Public Health and Primary Care (J Huang MD, C Zhong MD), The Chinese University of Hong Kong, Hong Kong, China; Department of Community Medicine, Datta Meghe Institute of Medical Sciences, Wardha, India (Prof S G Choudhari MD, Prof A M Gaidhane MD); Department of Epidemiology and Preventative Medicine, Monash University, Melbourne, VIC, Australia (E K Chowdhury PhD); Department of Pulmonary Medicine, Christian Medical College and Hospital (CMC), Vellore, India (Prof D J Christopher MD); Center for Biomedicine and Community Health, VNU-International School, Hanoi, Vietnam (D Chu PhD); Health Effects Institute, Boston, MA, USA (A J Cohen DSc); Department of Medicine (Prof N Cruz-Martins PhD), Laboratory for Process Engineering, Environment, Biotechnology and Energy (J Loureiro PhD), University of Porto, Porto, Portugal; Department of Health Sciences, Institute of Research and Advanced Training in Health Sciences and Technologies, Famalicão, Portugal (Prof N Cruz-Martins PhD); School of Public Health, Walailak University, Thai Buri, Thailand (O Dadras DrPH); Department of Global Public Health and Primary Care, University of Bergen, Bergen, Norway (O Dadras DrPH); Department of Public Health (F T Dagnaw MPH), Department of Midwifery (B G Kassa MSc), Debre Tabor University, Debre Tabor, Ethiopia; Department of Research (H Kaur MPH, A Pandey PhD), Public Health Foundation of India, Gurugram, India (Prof L Dandona MD, Prof R Dandona PhD, G Kumar PhD, D K Lal MD); School of Public Health (A T Dao PhD), Mater Research Institute (M Moniruzzaman PhD), The University of Queensland, Brisbane, QLD, Australia; Epidemiology Department, Hanoi Medical University, Hanoi, Vietnam (A T Dao PhD); School of Public Health (S Debela MPH), Public Health, Epidemiology, Biostatistics, Research (D G Tufa MPH), Salale University, Fiche, Ethiopia; Department of Biomedical Sciences, Jimma University, Jimma, Ethiopia (D Dereje MSc); Graduate Medical Education, Gujarat Adani Institute of Medical Sciences, Bhuj, India (H D Desai MD); Department of Public Health, Adigrat University, Adigrat, Ethiopia (B Desye MSc); Department of Pharmacy Practice, National Institute of Pharmaceutical Education and Research, Hajipur, India (S Dhingra PhD); Department of Environmental Health (N Diao DSc), Department of Health Policy and Oral Epidemiology (Z S Natto DrPH), Harvard University, Boston, MA, USA; Center of Complexity Sciences, National Autonomous University of Mexico, Mexico City, Mexico (Prof D Diaz PhD); Faculty of Veterinary Medicine and Zootechnics, Autonomous University of Sinaloa, Culiacán Rosales, Mexico (Prof D Diaz PhD); Institute for Global Health Innovations (L P Doan MSc, L G Vu MSc), Faculty of Medicine (L P Doan MSc, L G Vu MSc), Institute of Research and Development (M Hosseinzadeh PhD), Duy Tan University, Da Nang, Vietnam; Center of Excellence in Health Equity, Training and Research, Baylor College of Medicine, Houston, TX, USA (D Dongarwar MS); Responsabilidade Social (W M dos Santos PhD), Hospital Alemão Oswaldo Cruz (Oswaldo Cruz German Hospital), São Paulo, Brazil; Brazilian Centre for Evidence-based Healthcare: An Affiliate Centre of the Joanna Briggs Institute, Joanna Briggs Institute, São Paulo, Brazil (W M dos Santos PhD); Forensic Medicine and Toxicology, Kasturba Medical College Mangalore (H L Dsouza MD), Department of Community Medicine (N Joseph MD, R Thapar MD), Manipal Academy of Higher Education, Mangalore, India; Forensic Medicine and Toxicology, Kasturba Medical College Mangalore, Mangalore, India (H L Dsouza MD); Institute of Microbiology and Immunology (E Dubljanin PhD), Faculty of Medicine (I M Ilic PhD, Prof M M Santric-Milicevic PhD), School of Public Health and Health Management (Prof M M Santric-Milicevic PhD), University of Belgrade, Belgrade, Serbia; Infection and Tropical Medicine, University of Sheffield, Sheffield, UK (O C Durojaiye MPH); School of Health Sciences, University of Science Malaysia, Kubang Kerian, Malaysia (H A Edinur PhD); Department of Epidemiology and Medical Statistics (M Ekholuenetale MSc, A F Fagbamigbe PhD), Faculty of Public Health (M Ekholuenetale MSc, I I Olufadewa MHS), Department of Health Promotion and Education (S E Ibitoye MPH), Department of Community Medicine (O S Ilesanmi PhD), College of Medicine (A P Okekunle PhD), University of Ibadan, Ibadan, Nigeria; Department of Biological Sciences, University of Medical Sciences, Ondo, Ondo, Nigeria (T C Ekundayo PhD); Department of Epidemiology and Biostatistics, Cairo University, Cairo, Egypt (E D El Desouky PhD); Biomedical Informatics and Medical Statistics Department (I El Sayed PhD), Pathology Department (Prof I M Talaat PhD), Alexandria University, Alexandria, Egypt; Department of Clinical Pathology, Mansoura University, Mansoura, Egypt (Prof M El Sayed Zaki PhD); Faculty of Medicine, University of Tripoli, Tripoli, Libya (M Elhadi MD); Internal Medicine Department, Rochester General Hospital, Rochester, NY, USA (A M R Elkhapery MD); Lincoln International Institute for Rural Health, University of Lincoln, Lincoln, UK (L Engelbert Bain PhD); Department of International Cyber Education, Mongolian National University of Medical Sciences, Ulaanbaatar, Mongolia (R Erkhembayar MD, Prof C Oehir PhD); Internal Medicine (F Etace MD), Department of Genetics (S Pawar PhD), Yale University, New Haven, CT, USA; Department of Health Education & Health Promotion, School of Public Health, Semnan University of Medical Sciences, Semnan, Iran (M Ezati Asar PhD); Population and Behavioural Sciences Division, University of St Andrews, St Andrews, UK (A F Fagbamigbe PhD); Zoonotic Disease Research Center (S Falahi PhD), Department of Epidemiology (R Pakzad PhD), Ilam University of Medical Sciences, Ilam, Iran; Endocrinology and Metabolism Population Sciences Institute (A Fallahzadeh MD), Department of Epidemiology (S Nejadghaderi MD, S Rashedi MD, E Shaker MD), Quantitative Department (A Sheidaei MSc), Department of International Studies (P Shobeiri MD), Non-Communicable Diseases Research Center, Tehran, Iran (M Khosravifar MD, E Mohammadi MD, F Montazeri MD); Department of Political Science, University of Human Development, Sulaimaniyah, Iraq (Prof A Faraj PhD); Department of Health Policy and Administration, University of the Philippines Manila, Manila, Philippines (E A Faraon MD); Research Center on Public Health, University of Milan Bicocca, Monza, Italy (P Ferrara MD); Department of Health Sciences, University of Genoa, University of Genoa, Genoa, Italy (A A Ferrari MD); Department of Nursing, Wollega University, Nekemte, Ethiopia (G Fetensa MSc); Institute of Public Health, Charité Medical University Berlin, Berlin, Germany (F Fischer PhD); College of Medicine and Public Health, Flinders University, Adelaide, SA, Australia (J Flavel PhD); Department of Medical Parasitology (M Foroutan PhD), Faculty of Medicine (M Foroutan PhD), Abadan University of Medical Sciences, Abadan, Iran; Health Services Management Training Centre (P A Gaal PhD), Faculty of Health and Public Administration (J Lâm PhD, T P P Palicz MD), Semmelweis University, Budapest, Hungary; Department of Applied Social Sciences, Sapientia Hungarian University of Transylvania, Târgu-Mureş, Romania (P A Gaal PhD); Nutrition Innovation Centre for Food and Health, Ulster University, Coleraine,

UK (S Gaihr PhD); Department of Surgical Technology, Lorestan University of Medical Sciences, Khorramabad, Iran (N Galehdar PhD); Department of Tuberculosis, Manhiça Health Research Center, Manhiça, Mozambique (A L Garcia-Basteiro PhD); Viral and Bacterial Infections Research Program, Barcelona Institute for Global Health, Barcelona, Spain (A L Garcia-Basteiro PhD); Department of Radiology, King Edward Memorial Hospital, Mumbai, India (T Garg MBBS); Department of Nursing, Debre Berhan University, Debre Birhan, Ethiopia (B N B Gameda MSc); Pfizer Vaccines, Collegeville, PA, USA (B D Gessner MD); Agency of Preventive Medicine, Paris, France (B D Gessner MD); Department of Emergency and Critical Care Medicine (M Getachew MD), Department of Nursing (G F W Mijena MSc), Haramaya University, Harar, Ethiopia; School of Nursing, Arba Minch University, Arbaminch, Ethiopia (A Getie MSc); E-Learning Center, Faculty of Health (M Ghasemi Nour MD), Applied Biomedical Research Center (A Sahebkar PhD), Biotechnology Research Center (A Sahebkar PhD), Social Determinants of Health Research Center (M Sarkhosh PhD), Mashhad University of Medical Sciences, Mashhad, Iran; School of Public Health (A Ghahghaee BSc), Institute for Prevention of Non-communicable Diseases (R Kalhor PhD), Health Services Management Department (R Kalhor PhD), Department of Food Hygiene and Safety and Medical Microbiology Research Center (Prof R Mahmoudi PhD), Qazvin University of Medical Sciences, Qazvin, Iran; Department of Environmental Health Engineering, North Khorasan University of Medical Sciences, Bojnurd, Iran (A Gholizadeh PhD); Institute for Global Health Sciences (R Ghosh PhD), Department of Bioengineering and Therapeutic Sciences (Prof M S Zastrozhin PhD), University of California San Francisco, San Francisco, CA, USA; Department of Radiology (S Ghozy MD), Division of Infectious Diseases (Prof I I Tleyjeh MD), Mayo Clinic, Rochester, MN, USA; Department of Genetics, Sana Institute of Higher Education, Sari, Iran (P Goleij MSc); Department of Nursing, Arak University of Medical Sciences, Arak, Iran (M Golitaleb PhD, M Harorani MSc); Oncological Network, Prevention and Research Institute, Institute for Cancer Research, Prevention and Clinical Network, Florence, Italy (G Gorini MD); Center for Clinical and Epidemiological Research (A C Goulart PhD), Department of Internal Medicine (A C Goulart PhD), University of São Paulo, São Paulo, Brazil; Public Health Department, Salale University, Fitcha, Ethiopia (G G Goyomsa MPH); College of Medicine and Health Science, Jijiga University, Jijiga, Ethiopia (R A Guled PhD); Toxicology Department, Shriram Institute for Industrial Research, Delhi, Delhi, India (S Gupta MSc); School of Medicine, Deakin University, Geelong, VIC, Australia (V Gupta PhD); Department of Clinical Medicine (Prof V K Gupta PhD), School of Engineering (N Rabiee PhD), Macquarie University, Sydney, NSW, Australia; Department of Midwifery, Dire Dawa University, Dire Dawa, Ethiopia (A Guta MSc); School of Health and Environmental Studies, Hamdan Bin Mohammed Smart University, Dubai, United Arab Emirates (Prof S Hamidi DrPH); Department of Biochemistry and Molecular Biology, Bangladesh Agricultural University, Mymensingh, Bangladesh (Prof M Hannan PhD); Department of Anatomy, Dongguk University, Gyeongju, South Korea (Prof M Hannan PhD); Department of Zoology and Entomology, Al Azhar University, Cairo, Egypt (A I Hasaballah PhD); Department of Ophthalmology, IUMS, Karaj, Iran (H Hasani MD); Department of Plastic Surgery, The University of Texas MD Anderson Cancer Center, University of Texas, Houston, TX, USA (A M Hassan MD); Chapter of Addiction Medicine (Prof H Hassanian-Moghaddam MD), Save Sight Institute (H Kandel PhD), School of Public Health (L D Knibbs PhD), University of Sydney, Sydney, NSW, Australia; Independent Consultant, Tabriz, Iran (H Hassankhani PhD); Institute of Pharmaceutical Sciences, University of Veterinary and Animal Sciences, Lahore, Pakistan (K Hayat MS); Department of Pharmacy Administration and Clinical Pharmacy, Xian Jiaotong University, Xian, China (K Hayat MS); Public Health, Environmental Health, Epidemiology and Health Education Department (D Z Heyi MPH), Department of Midwifery (D G Tesfaye MSc), Madda Walabu University, Robe, Ethiopia; Department of Applied Microbiology, Taiz University, Taiz, Yemen (K Hezam PhD); Department of Microbiology, Nankai University, Tianjin, China (K Hezam PhD); Kasturba Medical College, Mangalore (R Holla MD, A Kamath MD), Department of Physiotherapy (Prof V K PhD), Forensic Medicine and Toxicology, Kasturba Medical College Mangalore (J Padubidri MD), Manipal Academy of Higher Education, Manipal, India (A Kamath MD); Research Department, Electronic Medical Records for the Developing World, York, UK (S Hong MD); Department of Pulmonology, Yokohama City University, Yokohama, Japan (N Horita PhD); National Human Genome Research Institute, National Institutes of Health, Bethesda, MD, USA (N Horita PhD); Internal Medicine Department (M Hostiuic PhD), Department of Dermatology (C N Matei PhD, M Tampa PhD), Carol Davila University of Medicine and Pharmacy, Bucharest, Romania; College of Science and Engineering, Hamad Bin Khalifa University, Doha, Qatar (Prof M Househ PhD); Burn Research Center (S Hoveidamanesh MD), Shahid Motahari Hospital, Tehran, Iran; Department of Biomolecular Sciences, University of Zakho, Zakho, Iraq (N R Hussein PhD); Department of Public Health, University of Naples Federico II, Naples, Italy (Prof I Iavicoli PhD); Department of Community Medicine, University College Hospital, Ibadan, Ibadan, Nigeria (O S Ilesanmi PhD); Department of Epidemiology, University of Kragujevac, Kragujevac, Serbia (Prof M D Ilic PhD); Department of Clinical Pharmacy, MAHSA University, Bandar Saujana Putra, Malaysia (Prof N Ismail PhD); Department of Health Services Research, University of Tsukuba, Tsukuba, Japan (M Iwagami PhD); Department of Non-Communicable Disease Epidemiology (M Iwagami PhD), Medical Statistics Department (S Shivalli MD), London School of Hygiene & Tropical Medicine, London, UK; Department of Environmental Health Engineering (J Jaafari PhD), Gastrointestinal and Liver Diseases Research Center (F Joukar PhD), Caspian Digestive Disease Research Center (F Joukar PhD), Department of Medical-Surgical Nursing (S Karkhah MSc), Guilan University of Medical Sciences, Rasht, Iran; Division of Pulmonary Medicine, Lausanne University Hospital, Lausanne, Switzerland (E Jamshidi PharmD); Department of Preventive Medicine, Yonsei University, Seodaemun-gu, South Korea (Prof S Jang PhD); Health Informatic Lab (T Javaheri PhD), Department of Computer Science (R Rawassizadeh PhD), Boston University, Boston, MA, USA; Department of Medical Mycology (J Javidnia PhD), Department of Immunology (Prof A Rafiei PhD), Molecular and Cell Biology Research Center (Prof A Rafiei PhD), Mazandaran University of Medical Sciences, Sari, Iran; Centre of Studies and Research, Ministry of Health, Muscat, Oman (S Jayapal PhD); Postgraduate Institute of Medicine (U U Jayarajah MD), Department of Pharmacology (P Ranasinghe PhD), University of Colombo, Colombo, Sri Lanka; Department of Surgery, National Hospital, Colombo, Sri Lanka (U U Jayarajah MD); Department of Biochemistry, Government Medical College, Mysuru, India (Prof S Jayaram MD); Institute of Molecular and Clinical Ophthalmology Basel, Basel, Switzerland (Prof J B Jonas MD); Department of Ophthalmology, Heidelberg University, Mannheim, Germany (Prof J B Jonas MD); Department of Family Medicine and Public Health, University of Opole, Opole, Poland (J J Jozwiak PhD); School of Public Health, University College Cork, Cork, Ireland (Z Kabir PhD); Department of Medicine, Faculty of Medicine, University of Tlemcen, Tlemcen, Algeria (S O Kacimi MD); Social Determinants of Health Research Center (L R Kalankesh PhD), Department of Anatomy (J Majidpoor PhD), Department of Nutrition (S Tabatabaeizadeh PhD), Gonabad University of Medical Sciences, Gonabad, Iran; Department of Community Medicine and Family Medicine, All India Institute of Medical Sciences, Hyderabad, India (B D Kamble MD); Department of Community Medicine, Banaras Hindu University, Varanasi, India (B D Kamble MD); Sydney Eye Hospital, South Eastern Sydney Local Health District, Sydney, NSW, Australia (H Kandel PhD); School of Health Professions and Human Services, Hofstra University, Hempstead, NY, USA (I M Karaye MD); Institute for Epidemiology and Social Medicine, University of Münster, Münster, Germany (A Karch MD); Centre for Tropical Diseases and Global Health, Catholic University of Bukavu, Bukavu, Democratic Republic of the Congo (P D Katoto PhD); Department of Global Health, Stellenbosch University, Cape Town, South Africa (P D Katoto PhD); Department of Healthcare Services Management, School of Health, Alborz University of Medical Sciences, Karaj, Iran (L Keikavoosi-Arani PhD); Department of Public Health, Jordan University of Science and Technology, Irbid, Jordan (Prof Y S Khader PhD); Department of Epidemiology and

Biostatistics (E A Khan MPH), Department of Public Health (Z Z Piracha PhD), Health Services Academy, Islamabad, Pakistan; Epidemiology Department (M Khan MD), Substance Abuse and Toxicology Research Center (S Mohan PhD), Jazan University, Jazan, Saudi Arabia; Department of Population Science, Jatiya Kabi Kazi Nazrul Islam University, Mymensingh, Bangladesh (M Khan PhD); Primary Care Department, NHS North West London, London, UK (M A Khan MSc); Department of Clinical Pharmacy, Jouf University, Sakaka, Saudi Arabia (Y H Khan PhD, T Mallhi PhD); Basic Medical Sciences, Yarmouk University, Irbid, Jordan (M M Khatatbeh PhD); Department of Public Health, New Mexico State University, Las Cruces, NM, USA (Prof J Khubchandani PhD); Department of Genomics and Digital Health, Samsung Advanced Institute for Health Sciences & Technology, Seoul, South Korea (M Kim MD); Public Health Center, Ministry of Health and Welfare, Wando, South Korea (M Kim MD); Department of Nutrition, Simmons University, Boston, MA, USA (R W Kimokoti MD); School of Health Sciences, Kristiania University College, Oslo, Norway (Prof A Kisa PhD); Department of Global Community Health and Behavioral Sciences (Prof A Kisa PhD), Department of Medicine (M F Qadir PhD), Tulane University, New Orleans, LA, USA; Department of Nursing and Health Promotion, Oslo Metropolitan University, Oslo, Norway (S Kisa PhD); Global Healthcare Consulting, New Delhi, India (S Kochhar MD); Social Determinants of Health Research Center, Saveh University of Medical Sciences, Saveh, Iran (H Koohestani PhD); Independent Consultant, Jakarta, Indonesia (S Kosen MD); Department of Internal and Pulmonary Medicine, Sheri Kashmir Institute of Medical Sciences, Srinagar, India (Prof P A Koul MD); Biomedical Research Networking Center for Mental Health Network, San Juan de Dios Sanitary Park, Sant Boi de Llobregat, Spain (A Koyanagi MD); Department of Anthropology, Panjab University, Chandigarh, India (Prof K Krishan PhD); Department of Demography (Prof B Kuate Defo PhD), Department of Social and Preventive Medicine (Prof B Kuate Defo PhD), University of Montreal, Montreal, QC, Canada; Faculty of Health and Life Sciences, Coventry University, Coventry, UK (O P Kurmi PhD); Department of Medicine (O P Kurmi PhD), Department of Psychiatry and Behavioural Neurosciences (A T Olagunju MD), McMaster University, Hamilton, ON, Canada; Department of Nephrology, Usha Hospital, Chengannur, India (A Kuttikkattu MD); NEVES Society for Patient Safety, NEVES Society for Patient Safety, Budapest, Hungary (J Lám PhD); Unit of Genetics and Public Health (Prof I Landires MD), Unit of Microbiology and Public Health (V Nuñez-Samudio PhD), Institute of Medical Sciences, Las Tablas, Panama; Department of Public Health (V Nuñez-Samudio PhD), Ministry of Health, Herrera, Panama (Prof I Landires MD); Department of Clinical and Experimental Medicine, University of Catania, Catania, Italy (C Ledda PhD); Pattern Recognition and Machine Learning Lab, Gachon University, Seongnam, South Korea (Prof S Lee PhD); Department of Prevention, USL Tuscany Center, Firenze, Italy (M Levi PhD); Department of Health Sciences, University of Florence, Florence, Italy (M Levi PhD); Centre for Tropical Medicine and Global Health (S Lewycka PhD), Nuffield Department of Medicine (Prof R J Maude PhD), Big Data Institute (C E Moore PhD), University of Oxford, Oxford, UK; Oxford University Clinical Research Unit, Wellcome Trust Asia Programme, Hanoi, Vietnam (S Lewycka PhD); School of Life Sciences, University of Technology Sydney, Ultimo, NSW, Australia (G Liu PhD); Centre for Inflammation, Centenary Institute, Camperdown, NSW, Australia (G Liu PhD); Department of Paediatrics, All India Institute of Medical Sciences, New Delhi, India (Prof R Lodha MD); Department of Health Economics, Syreon Research Romania, Târgu Mureş, Romania (L Lorenzovici MSc); Department of Doctoral Studies, George Emil Palade University of Medicine, Pharmacy, Science, and Technology from Targu Mures, Târgu Mureş, Romania (L Lorenzovici MSc); School of Health, Polytechnic Institute of Porto, Porto, Portugal (J Loureiro PhD); Department of Biostatistics and Epidemiology, Yazd University of Medical Sciences, Yazd, Iran (F Madadzadeh PhD); Department of Biostatistics and Epidemiology (M Mahmoodimanesh PhD), Department of Immunology (M Razeghinia MSc), Kerman University of Medical Sciences, Kerman, Iran; University Institute of Public Health (A A Malik PhD), Department of Pharmacy Practice, Faculty of Pharmacy (M Salman PhD), The University of Lahore, Lahore, Pakistan; Heart and Vascular Institute, Cleveland Clinic Abu Dhabi, Abu Dhabi, United Arab Emirates (Y Manla MD); Board of Directors, Association of Resident Physicians, Bucharest, Romania (C N Matei PhD); Division of Infection, Immunity and Respiratory Medicine, University of Manchester, Manchester, UK (A G Mathioudakis MD); North West Lung Centre, Manchester University NHS Foundation Trust, Manchester, UK (A G Mathioudakis MD); Epidemiology Department, Mahidol Oxford Tropical Medicine Research Unit, Bangkok, Thailand (Prof R J Maude PhD); Civil Engineering Faculty, University of Colima, Coquimatlán City, Mexico (Prof O Mendoza-Cano PhD); University Research Institute, National and Kapodistrian University of Athens, Athens, Greece (A A Mentis MD); Breast Surgery Unit, Helsinki University Hospital, Helsinki, Finland (T J Meretoja MD); University of Helsinki, Helsinki, Finland (T J Meretoja MD); College of Medicine and Health Sciences, Institute of Public Health, Department of Epidemiology and Biostatistics, University of Gondar, Addis Ababa, Ethiopia (M W Merid MPH); University Centre Varazdin, University North, Varazdin, Croatia (T Mestrovic PhD); Department of Medicine, University of Medicine and Pharmacy at Ho Chi Minh City, Ho Chi Minh City, Vietnam (L Minh MD); College of Applied Medical Sciences, Majmaah University, Riyadh, Saudi Arabia (S A Mir PhD); Department of Chemistry, Umm Al Qura University, Makkah, Saudi Arabia (A Mirza PhD); Department of Hospital Administration (M Mirza MD), Department of Radiodiagnosis (P Singh MD), All India Institute of Medical Sciences, Bathinda, India; Social Determinants of Health Center, Urmia University of Medical Science, Urmia, Iran (M Mirza-Aghazadeh-Attari MD); Department of Medical Laboratory Sciences (A S Misganaw MSc), Department of Microbial Cellular and Molecular Biology (A S Misganaw MSc), Addis Ababa University, Addis Ababa, Ethiopia; National Data Management Center for Health (A T Misganaw PhD), Infection Prevention & Control and Water, Sanitation and Hygiene Unit (B Wagaye MPH), Ethiopian Public Health Institute, Addis Ababa, Ethiopia; Department of Information Technology, Lebanese French University, Erbil, Iraq (M Mohammadi PhD); Department of Biology, University of Jeddah, Jeddah, Saudi Arabia (A Mohammed PhD); Health Systems and Policy Research Unit, Ahmadu Bello University, Zaria, Nigeria (S Mohammed PhD); Department of Health Care Management, Technical University of Berlin, Berlin, Germany (S Mohammed PhD); School of Health Sciences, University of Petroleum and Energy Studies, Dehradun, India (S Mohan PhD); Health Services Management, Iran University of Medical Sciences, Iran, Iran (M Mohseni PhD); Oncology Department, Appalachian Regional Healthcare, Hazard, KY, USA (N Moka MD); Department of Internal Medicine, University of Kentucky, Lexington, KY, USA (N Moka MD); Clinical Epidemiology and Public Health Research Unit, Burlo Garofolo Institute for Maternal and Child Health, Trieste, Italy (L Monasta DSc, I Ronfani PhD); School of Medicine (Prof A Moradi PhD), Infectious Diseases Research Center (H Shirzad-Aski PhD), Department of Microbiology (A Tahamtan PhD), Golestan University of Medical Sciences, Gorgan, Iran; International Laboratory for Air Quality and Health, Queensland University of Technology, Brisbane, QLD, Australia (Prof L Morawska PhD); Department of Medicine (E Mostafavi PhD), Stanford Cardiovascular Institute (E Mostafavi PhD), Stanford University, Palo Alto, CA, USA; Department of Epidemiology and Biostatistics (S Mubarik MS), School of Medicine (Z Zhang PhD), Wuhan University, Wuhan, China; Epidemiology, Family Medicine Unit 19, Mexican Institute of Social Security, Colima, Mexico (E Murillo-Zamora PhD); Postgraduate in Medical Sciences, Universidad de Colima, Colima, Mexico (E Murillo-Zamora PhD); Department of Pediatric Medicine, The Children's Hospital & The Institute of Child Health, Multan, Pakistan (Prof G Mustafa MD); Department of Pediatrics & Pediatric Pulmonology, Institute of Mother & Child Care, Multan, Pakistan (Prof G Mustafa MD); Department of Pulmonary Medicine, Government Medical College Trivandrum, Trivandrum, India (S Nair MD); Health Action by People, Trivandrum, India (S Nair MD); School of Pharmacy, University of Reading, Reading, UK (A Naqvi PhD); Mysore Medical College and Research Institute, Government Medical College, Mysore, India (Prof S Narasimha Swamy MD); Amity Institute of Forensic Sciences, Amity University, Noida, India (B P Nayak PhD); Health Innovation and Transformation Centre, Federation University Australia,

Brisbane, QLD, Australia (H V N Nguyen PhD); Department of Population and Quantitative Health Sciences, University of Massachusetts Medical School, Worcester, MA, USA (H V N Nguyen PhD); International Islamic University Islamabad, Islamabad, Pakistan (R K Niazi PhD); Center of Excellence in Reproductive Health Innovation, University of Benin, Benin City, Nigeria (C I Nzoputam MPH); Department of Physiology, University of Benin, Edo, Nigeria (O J Nzoputam PhD); Department of Physiology, Benson Idahosa University, Benin City, Nigeria (O J Nzoputam PhD); Administrative and Economic Sciences Department, University of Bucharest, Bucharest, Romania (Prof B Oancea PhD); Advisory Board, Ministry of Health, Ulaanbaatar, Mongolia (Prof C Ochir PhD); Department of Community Health and Primary Care, University of Lagos, Idi Araba, Nigeria (O O Odukoya MSc); Department of Family and Preventive Medicine, University of Utah, Salt Lake City, UT, USA (O O Odukoya MSc); Health Promotion Research Center, Zahedan University of Medical Sciences, Zahedan, Iran (H Okati-Aliabad PhD); Department of Food and Nutrition, Seoul National University, Seoul, South Korea (A P Okekunle PhD); School of Pharmacy, University of the Western Cape, Cape Town, South Africa (O C Okonji MSc); Department of Psychiatry, University of Lagos, Lagos, Nigeria (A T Olagunju MD); Slum and Rural Health Initiative Research Academy, Slum and Rural Health Initiative, Ibadan, Nigeria (I I Olufadewa MHS); Mass Communication Department, Ajman University, Dubai, United Arab Emirates (E Omer PhD); Graduate School of Public Health, San Diego State University, San Diego, CA, USA (Prof E Oren PhD); Department of Global Health Nursing, St Luke's International University, Chuo-ku, Japan (Prof E Ota PhD); Laboratory of Public Health Indicators Analysis and Health Digitalization, Moscow Institute of Physics and Technology, Dolgoprudny, Russia (N Oststavnov BA); Department of Epidemiology and Population Health, Khalifa University, Abu Dhabi, United Arab Emirates (A Oulhaj PhD); Department of Respiratory Medicine, Jagadguru Sri Shivarathreeswara Academy of Health Education and Research, Mysore, India (Prof M P A DNB); Hungarian Health Management Association, Hungarian Health Management Association, Budapest, Hungary (T P P Palicz MD); Clinical Research, Nepal Health Research Council, Kathmandu, Nepal (S Pant MPH); Vision and Eye Research Institute, Anglia Ruskin University, Cambridge, UK (Prof S Pardhan PhD); Department of Medical Humanities and Social Medicine, Kosin University, Busan, South Korea (Prof E Park PhD); Central Department of Public Health, Tribhuvan University, Kathmandu, Nepal (R Paudel MPH); Department of Infection Control, Taihe Hospital, Shiyuan, China (M Peng MPH); The First Clinical College, Hubei University of Medicine, Shiyuan, China (M Peng MPH); School of Public Health, Curtin University, Bentley, WA, Australia (Prof G Pereira PhD); Centre for Fertility and Health, Norwegian Institute of Public Health, Oslo, Norway (Prof G Pereira PhD); College of Science, Department of Biology, University of Bahrain, Sakir, Bahrain (Prof S Perna PhD); Department of Zoology, Yadava College, Madurai District, India (Prof N Perumalsamy PhD); Department of Zoology, Annai Fathima College, Madurai, Madurai District, India (Prof N Perumalsamy PhD); Department of Statistics and Econometrics, Bucharest University of Economic Studies, Bucharest, Romania (I Petcu PhD); Medical College, Tairunnessa Memorial Medical College and Hospital, Gazipur, Bangladesh (V Podder HSC); School of Public Health, University of Adelaide, Adelaide, SA, Australia (V Podder HSC); University Medical Center Groningen (Prof M J Postma PhD), School of Economics and Business (Prof M J Postma PhD), University of Groningen, Groningen, Netherlands; Non-Communicable Diseases Research Center, Bam University of Medical Sciences, Bam, Iran (N Pourtaheri PhD); International Operations Department, Mérieux Foundation, Lyon, France (M Raad MD); Biomedical Engineering Department, Amirkabir University of Technology, Tehran, Iran (Prof M Rabiee PhD); Pohang University of Science and Technology, Pohang, South Korea (N Rabiee PhD); Laboratory Sciences Department, Maragheh University of Medical Sciences, Maragheh, Iran (S Raeghi PhD); Center for Surveillance, Immunization, and Epidemiologic Research (M Rahman PhD), National Institute of Infectious Diseases, Tokyo, Japan (M Shigematsu PhD); Global Health Nursing Department, St. Luke's International University, Tokyo, Japan (M Rahman PhD); Department of Population Science and Human Resource Development, University of Rajshahi, Rajshahi, Bangladesh (M Rahman DrPH); School of Nursing and Healthcare Professions, Federation University Australia, Berwick, VIC, Australia (M Rahman PhD); Future Technology Research Center, National Yunlin University of Science and Technology, Yunlin, Taiwan (A Rahmani PhD); Department of Cardiology, Emory University, Atlanta, GA, USA (P Ram MD); Department of Epidemiology, Biostatistics and Occupational Health, McGill University, Montreal, QC, Canada (J Rana MPH); Research and Innovation Division, South Asian Institute for Social Transformation, Dhaka, Bangladesh (J Rana MPH); Department of Health Innovation, Manipal Academy of Higher Education, Udupi, India (U Rani PhD); Department of Oral Pathology, Sharavathi Dental College and Hospital, Shimogga, India (S Rao MDS); Department of Geography, Soran University, Soran, Iraq (A Rasul PhD); Department of Biomedical Engineering, Khulna University of Engineering and Technology, Khulna, Bangladesh (Z Ratan MSc); School of Health and Society, University of Wollongong, Wollongong, NSW, Australia (Z Ratan MSc); WHO Collaborating Centre for Public Health Education and Training (D L Rawaf MD), Department of Primary Care and Public Health (Prof S Rawaf MD), Imperial College London, London, UK; NIHR-Biomedical Research Centre (Prof A Zumla PhD), University College London Hospitals, London, UK (D L Rawaf MD); Academic Public Health England, Public Health England, London, UK (Prof S Rawaf MD); Department of Immunology and Laboratory Sciences (M Razeghinia MSc), Medical Laboratory Sciences (M Sahebazamani MSc), Sirjan School of Medical Sciences, Sirjan, Iran; Department of Biological Sciences, King Abdulaziz University, Jeddah, Egypt (Prof E M M Redwan PhD); Protein Research, Research and Academic Institution, Alexandria, Egypt (Prof E M M Redwan PhD); School of Medicine (Prof A M N Renzaho PhD), Translational Health Research Institute (Prof A M N Renzaho PhD), Western Sydney University, Campbelltown, NSW, Australia; Department of Epidemiology and Biostatistics (Prof M Rezaeian PhD), Department of Medical Biochemistry (M Sahebazamani MSc), Rafsanjan University of Medical Sciences, Rafsanjan, Iran; Department of Public Health (A Riad DDS), Czech National Centre for Evidence-based Healthcare and Knowledge Translation (A Riad DDS), Masaryk University, Brno, Czech Republic; Institute of Diagnostic and Interventional Radiology and Neuroradiology, School of Medicine, Essen University Hospital, Essen, Germany (R Rikhtegar MD); Department of Pharmacology and Toxicology, University of Antioquia, Medellin, Colombia (Prof J A B Rodriguez PhD); Department of Critical Care Medicine, University of Pittsburgh, Pittsburgh, PA, USA (K E Rudd MD); Research and Development, Islamabad Diagnostic Center Pakistan, Islamabad, Pakistan (Prof U Saeed PhD); Biological Production Division, National Institute of Health, Islamabad, Pakistan (Prof U Saeed PhD); Faculty of Medicine, Bioscience and Nursing, MAHSA University, Selangor, Malaysia (S Z Safi PhD); Interdisciplinary Research Centre in Biomedical Materials, COMSATS Institute of Information Technology, Lahore, Pakistan (S Z Safi PhD); Department of Clinical Medical Sciences, University of the West Indies, St Augustine, Trinidad and Tobago (S Sakhamuri MD); Thoracic Department, North Central Regional Health Authority, Champ Fleurs, Trinidad and Tobago (S Sakhamuri MD); Department of Entomology, Ain Shams University, Cairo, Egypt (A M Samy PhD); Faculty of Health & Social Sciences, Bournemouth University, Bournemouth, UK (B Sathian PhD); Department of Public Health Sciences, University of North Carolina at Charlotte, Charlotte, NC, USA (M Sawhney PhD); Department of Preventive and Social Medicine, Jawaharlal Institute of Postgraduate Medical Education and Research, Puducherry, India (G Saya MD); Department of Population and Health, University of Cape Coast, Cape Coast, Ghana (A Seidu MPhil); College of Public Health, Medical and Veterinary Sciences, James Cook University, Townsville, QLD, Australia (A Seidu MPhil); National Heart, Lung, and Blood Institute, National Institute of Health, Rockville, MD, USA (A Seylani BS); Public Health Division, An-Najah National University, Nablus, Palestine (A A Shaheen PhD); Independent Consultant, Karachi, Pakistan (M A Shaikh MD); Research Institute of Pharmaceutical Sciences (H Shamsad PhD), University of Karachi, Karachi, Pakistan; Nahavand School of Allied Medical Sciences, Hamadan University of Medical

Sciences, Hamadan, Nahavand, Iran (A Sharifi PhD); Department of Medical Oncology, Kent Hospital, Warwick, RI, USA (P Sharma MD); Department of Microbiology, Kasturba Medical College, Mangalore, Manipal Academy of Higher Education, Manipal, Mangalore, India (S M Shenoy MD); Public Health Dentistry Department, Krishna Institute of Medical Sciences Deemed to be University, Karad, India (Prof K M Shivakumar PhD); School of Health, Victoria University of Wellington, Wellington, New Zealand (Prof C R Simpson PhD); Usher Institute, University of Edinburgh, Edinburgh, UK (Prof C R Simpson PhD); Department of Pulmonary and Critical Care Medicine, Medical College of Wisconsin, Milwaukee, WI, USA (H Singh MD); School of Medicine, University of Alabama at Birmingham, Birmingham, AL, USA (Prof J A Singh MD); Medicine Service, US Department of Veterans Affairs, Birmingham, AL, USA (Prof J A Singh MD); Department of Chemistry, Maharishi Markandeshwar (Deemed to be University), Mullana, India (S S Siwal PhD); Department Number 16, Moscow Research and Practical Centre on Addictions, Moscow, Russia (V Y Skryabin MD); Department of Infectious Diseases and Epidemiology, Pirogov Russian National Research Medical University, Moscow, Russia (A A Skryabina MD); Taub Institute for Research on Alzheimer's Disease and the Aging Brain, Columbia University Medical Center, New York, NY, USA (S Song PhD); Department of Land Surveying and Geo-Informatics, Hong Kong Polytechnic University, Hong Kong, China (Y Song PhD); Department of Microbiology, All India Institute of Medical Sciences, Bilaspur, India (P Sood PhD); Division of Community Medicine, International Medical University, Kuala Lumpur, Malaysia (C T Sreeramareddy MD); Department of Medicine, Democritus University of Thrace, Alexandroupolis, Greece (P Steiropoulos MD); Center for Biotechnology and Microbiology, University of Swat, Mingora, Swat, Pakistan (M Suleman PhD); School of Life Sciences, Xiamen University, China, Xiamen, China (M Suleman PhD); Department of Biostatistics and Epidemiology, Shahid Sadoughi University of Medical Sciences, Yazd, Iran (M Taheri Soodejani PhD); Department of Dermato-Venereology, Dr Victor Babes Clinical Hospital of Infectious Diseases and Tropical Diseases, Bucharest, Romania (M Tampa PhD); Department of Science, Technology and Natural Resources, Policy Research Institute, Kathmandu, Nepal (S Tandukar PhD); Department of Economics, Rice University, Houston, TX, USA (N Y Tat MS); Research and Innovation Department, Enventure Medical Innovation, Houston, TX, USA (N Y Tat MS); Department of Pathology, University of Texas, Galveston, TX, USA (V Y Tat BS); Department of Public Health, Dire Dawa university, Dire Dawa, Ethiopia (Y M Tefera MPH); Pediatric Intensive Care Unit, King Saud University, Riyadh, Saudi Arabia (M Tamsah MD); Faculty of Public Health, Universitas Sam Ratulangi, Manado, Indonesia (J H V Ticoalu MPH); Department of Epidemiology and Biostatistics, Birjand University of Medical Sciences, Birjand, Iran (A Tiyuri MSc); Infectious Diseases Department, King Fahad Medical City, Riyadh, Saudi Arabia (Prof I I Tleyjeh MD); General Department of Surgery, National Center of Traumatology and Orthopedics, Ulaanbaatar, Mongolia (M Togtmol MD); Board, Mongolian Burns Association, Ulaanbaatar, Mongolia (M Togtmol MD); Department of Pathology and Legal Medicine, University of São Paulo, Ribeirão Preto, Brazil (M R Tovani-Palone PhD); Modestum, London, UK (M R Tovani-Palone PhD); Amity Institute of Biotechnology, Amity University Rajasthan, Jaipur, India (E Upadhyay PhD); Clinical Cancer Research Center, Milad General Hospital, Tehran, Iran (S Valadan Tahbaz PhD, S Yahyazadeh Jabbari MD); Department of Microbiology, Islamic Azad University, Tehran, Iran (S Valadan Tahbaz PhD); Velez Sarsfield Hospital, Buenos Aires, Argentina (Prof P R Valdez MD); Laboratory of Toxicology, University of Crete, Heraklion, Greece (C Vardavas PhD); UKK Institute, Tampere, Finland (Prof T J Vasankari MD); Faculty of Medicine and Health Technology, Tampere University, Tampere, Finland (Prof T J Vasankari MD); Faculty of Information Technology, HUTECH University, Ho Chi Minh City, Vietnam (B Vo PhD); Foundation University Medical College, Foundation University Islamabad, Islamabad, Pakistan (Prof Y Waheed PhD); Department of Otolaryngology, Peking University Third Hospital, Beijing, China (Y Wang MD); Department of Biomedical Sciences, City University of Hong Kong, Hong Kong, China (A Waris MS); Department of

Community Medicine, Rajarata University of Sri Lanka, Anuradhapura, Sri Lanka (N D Wickramasinghe MD); Cardiovascular Program, The George Institute for Global Health, Sydney, NSW, Australia (X Xu PhD); Department of Clinical Microbiology, Iranshahr University of Medical Sciences, Iranshahr, Iran (S Yaghoubi PhD); Department of Pediatrics, Kyung Hee University, Seoul, South Korea (D Yon MD); Department of Neuropsychopharmacology, National Center of Neurology and Psychiatry, Kodaira, Japan (N Yonemoto PhD); Department of Public Health, Juntendo University, Tokyo, Japan (N Yonemoto PhD); Department of Radiology, Children's Hospital of Philadelphia, Philadelphia, PA, USA (A Zandifar MD); Unit on Child & Adolescent Health, Medical Research Council South Africa, Cape Town, South Africa (Prof H J Zar PhD); Research and Development Department, Sina Medical Biochemistry Technologies, Shiraz, Iran (I Zare BSc); Addictology Department, Russian Medical Academy of Continuous Professional Education, Moscow, Russia (Prof M S Zastrozhin PhD); Department of International Health, Georgetown University, Washington, DC, USA (Prof W Zeng PhD); Department of Nutrition and Health Science, Ball State University, Muncie, IN, USA (M Zhang PhD); Department of Infection, University College London, London, UK (Prof A Zumla PhD).

Contributors

Please see appendix 1 (pp 89–94) for more detailed information about individual author contributions to the research, divided into the following categories: managing the overall research enterprise; writing the first draft of the manuscript; primary responsibility for applying analytical methods to produce estimates; primary responsibility for seeking, cataloguing, extracting, or cleaning data; designing or coding figures and tables; providing data or critical feedback on data sources; developing methods or computational machinery; providing critical feedback on methods or results; drafting the manuscript or revising it critically for important intellectual content; and managing the estimation or publications process. Members of the core research team (HHK, AV, SBS, AN, CET, MCD, RGB, JRL, MHB, SBA, KB, FBB, JTZ, WMG, HH, DB, RVD, MN, MB, and CJLM) for this topic area had full access to the underlying data used to generate the estimates presented in this article. All other authors had access to and reviewed the estimates as part of the research evaluation process, which included additional formal stages of review. The corresponding author had final responsibility for the decision to submit for publication.

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Data sharing

To download the data used in these analyses, please visit the Global Health Data Exchange GBD 2019 website.

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