

Ozone-related acute excess mortality projected to increase in the absence of climate and air quality controls consistent with the Paris Agreement

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104 **Summary (max 150 words)**

105 Short-term exposure to ground-level ozone in cities is associated with increased mortality and is
106 expected to worsen with climate and emission changes. However, no study has yet
107 comprehensively assessed future ozone-related acute mortality across diverse geographic
108 areas, various climate scenarios, and using CMIP6 multi-model ensembles, limiting our
109 knowledge on future changes in global ozone-related acute mortality and ability to design
110 targeted health policies. Here, we combine CMIP6 simulations and epidemiological data from
111 406 cities in 20 countries or regions. We find that ozone-related deaths in 406 cities will
112 increase by 45 to 6,200 deaths/year between 2010-2014 and 2050-2054, with attributable
113 fractions increasing in all climate scenarios (from 0.17% to 0.22% of total deaths), except the
114 single scenario consistent with the Paris Climate Agreement (declines from 0.17% to 0.15% of
115 total deaths). These findings stress the need for more stringent air quality regulations, as current
116 standards in many countries are inadequate.

117 **Introduction**

118 Poor air quality is the largest environmental risk to human health, accounting for 6.7 million
119 deaths of the total 9 million pollution-related deaths in 2019¹. Among the different types of air
120 pollutants, ground-level ozone is a highly reactive and oxidative gas that can trigger coughing
121 and shortness of breath, worsen asthma, and cause damage to airways². Short-term exposure
122 to ozone has been linked to heightened excess mortality due to respiratory and cardiovascular
123 disease² as well as non-injury and all-cause mortality^{3,4}. A recent multi-city multi-country study
124 found the ozone-mortality relationship is spatially heterogeneous, with different countries and
125 regions experiencing different health impacts from short-term ozone exposure⁵.

126 Climate change and changes in emissions of ozone precursors are projected to worsen ground-
127 level ozone concentrations in many areas around the world, but with the magnitude of change
128 varying across regions⁶. The spatial heterogeneity of future changes in ozone production and
129 loss are driven by spatial differences in anthropogenic and natural emissions of ozone
130 precursors as well as spatial differences in meteorological parameters, such as temperature,
131 water vapor, and radiation⁷. The spatial heterogeneity observed in both future ozone
132 concentrations and ozone-mortality relationships suggest the importance of conducting health

133 impact assessments with broad geographical coverage for designing nuanced policies linked to
134 health promotion and disease prevention.

135 Previous research has projected varied impacts on ozone-related acute excess mortality,
136 ranging from minor to significant increases in scenarios characterized by high global warming
137 and elevated emissions of ozone precursors. Conversely, in scenarios with low global warming
138 and reduced emissions of ozone precursors, the projected health impacts range from moderate
139 decreases to small increases^{8, 9, 10}. However, these studies have generally been limited in
140 geographical coverage, with particular focus on the United States and China, and considered
141 limited future climate and air quality scenarios^{8,9,10}. No study has yet estimated future changes
142 in ozone-related acute excess mortality across a broad geographical scope and across a
143 broader range of future climate and air quality scenarios. This limits our ability to understand the
144 magnitude of the future health burden and to design effective policies to protect public health.

145 Here, we assess the near-future (2050-2054) changes in ozone-related short-term excess
146 mortality in 406 cities in 20 countries or regions due to changes in emissions and climate,
147 baseline mortality rates, and population under four Shared Socioeconomics Pathways (SSP)
148 scenarios. The analysis is conducted using simulated maximum daily 8-hour average ozone
149 concentrations from five models from the most recent Coupled Model Intercomparison Project
150 Phase 6 (CMIP6), projections of population and baseline mortality rates from the SSP project,
151 and observed ozone concentrations and country-or region-specific ozone-mortality relationships
152 from a recent study by the Multi-Country Multi-City Collaborative (MCC) Research Network. We
153 find that ozone-related acute excess mortality increases across all studied scenarios, and are
154 accompanied by rising attributable mortality fractions in all instances, except for the scenario
155 that aligns with the objectives of the Paris Climate Agreement. These results underscore the
156 significance of developing urgent and location-specific air pollution and climate mitigations to
157 minimize ozone-related health burden.

158 **Results**

159 *Results overview*

160 This study estimates the near-future (2050-2054) changes in ozone-related short-term excess
161 mortality by following the causal pathway of ozone concentrations to population exposure (see
162 our methods in the “Experimental Procedures” section). In the following subsections, we
163 describe our findings at each stage of the analysis. To concisely present our findings, we
164 aggregate the results obtained from specific sites to the country or region level; however, it is

165 important to note that these aggregations should not be construed as being representative of
166 the entire country or region.

167 *Present and future ozone concentrations*

168 Between 2010 and 2014, the annual mean bias-corrected ozone concentration across the 406
169 studied cities was $73 \mu\text{g}/\text{m}^3$. As shown in Fig 1a, annual mean daily maximum 8-hr average
170 (MDA8) ozone concentrations varied spatially, with higher concentrations found in Mexico,
171 Taiwan, Japan, the United States, Canada, and Japan, while lower concentrations were found
172 in Australia, Northern Europe, and China. On average, ozone concentrations exceeded the
173 maximum background levels of $70 \mu\text{g}/\text{m}^3$, a threshold assumed by past health impact
174 assessments in which concentrations are largely attributed to non-anthropogenic sources⁵,
175 about 222 days per year in the studied cities.

176 Under a scenario with strong climate and air pollution controls (SSP1-2.6), ozone
177 concentrations are projected to decrease on average by $8 \mu\text{g}/\text{m}^3$ between the present and
178 future period. In this scenario, ozone concentrations are projected to decrease in nearly all
179 studied countries or regions, except in Canada and China. The middle-of-the-road scenario
180 (SSP 2-4.5) is linked to a slight overall increase of $1 \mu\text{g}/\text{m}^3$ in ozone concentrations, with
181 projected increases in Asia, North America, Czech Republic, Estonia, Germany, and the UK,
182 and projected decreases in South Africa and the remaining studied countries or regions in
183 Europe. The regional rivalry scenario (SSP 3-7.0), which has weak climate and air pollution
184 controls, is linked to an average $9 \mu\text{g}/\text{m}^3$ increase in ozone concentrations across all countries
185 or regions. Similarly, the scenario with weak climate mitigation measures but strong air pollution
186 measures (SSP 5-8.5), is linked to an average increase of $4 \mu\text{g}/\text{m}^3$ ozone concentrations across
187 all studied countries or regions (Table S1). Though the number of models available for each
188 SSP varies, in general, the UKESM-1-0-LL and MPI-ESM-1-2-HAM models project higher
189 ozone concentrations than the other models in the present period.

190 A bias-correction technique was performed on simulated ozone concentrations from five global
191 chemistry-climate models using observed concentrations from the MCC network to obtain city-
192 level concentrations for the present (2010-2014) and future (2050-2054) periods. In the present
193 period, simulated annual mean ozone concentrations on average overestimate observed ozone
194 concentrations by about $4 \mu\text{g}/\text{m}^3$ (range: -75 to $50 \mu\text{g}/\text{m}^3$) (Fig S1). Biases varied spatially,
195 however, with simulated ozone concentrations generally underestimating ozone concentrations
196 in North America and Taiwan (range: 6 to $41 \mu\text{g}/\text{m}^3$ below observed concentrations) and

197 overestimating ozone concentrations in all other studied countries or regions (range: 4 to 32
198 $\mu\text{g}/\text{m}^3$ above observed concentrations) (Fig S4).

199 The spatial distribution and sign of changes in ozone concentrations are generally consistent
200 across models for a given SSP, though there are a few key differences. For instance, under
201 SSP 3-7.0, estimated changes in ozone concentrations are larger in the MPI-ESM-1-2-HAM
202 (average increase of $8 \mu\text{g}/\text{m}^3$) than in the other four models (average increase of $4 \mu\text{g}/\text{m}^3$).
203 Second, there are noticeable spatial differences in the changes in ozone concentrations in
204 certain geographies such as in central Europe and eastern Asia across the models. These
205 differences across models may be attributed to factors such as differences in treatment of
206 physical and chemical interactions, grid resolution, and initial conditions for atmospheric
207 variables and chemical species.

208 *Ozone-related short-term excess mortality*

209 In the present period (2010-2014), short-term exposure to ozone concentrations above 70
210 $\mu\text{g}/\text{m}^3$ accounts for 0.17% (95% eCI: 0.13-0.21%) of total deaths in the 406 cities. This is
211 equivalent to about 6,600 deaths per year (95% eCI: 2,600-10,100 deaths per year), similar to
212 previous estimates from Vicedo-Cabrera et al.⁵. Large numbers of ozone-related deaths are
213 found in key cities—such as 557 (95% eCI: 83-1,057) deaths per year in the Valley of Mexico,
214 Mexico; 250 (95% eCI: 116-425) deaths per year in Los Angeles, United States; 157 (95% eCI:
215 74-268) deaths per year in Tokyo, Japan; 138 (95% eCI: 75-202) deaths per year in Riverside,
216 United States; 121 (95% eCI: 75-212) deaths per year in Guadalajara, Mexico; 118 (95% eCI:
217 65-183) deaths per year in Toronto, Canada; and 105 (95% eCI: 0-226) deaths per year in
218 Taipei, Taiwan. When mortality estimates are restricted to days where ozone concentrations are
219 above the WHO guideline of $100 \mu\text{g}/\text{m}^3$, health impacts are reduced to 3,600 ozone-related
220 deaths per year. Total estimates of ozone-related acute excess mortality vary moderately
221 (range: 5,500 to 7,500 deaths per year) across global chemistry-climate models in the present
222 period, with the highest total mortality estimates produced by GFDL-ESM4 and the lowest total
223 mortality estimates produced by CESM2.

224 Under SSP-1.2.6, a scenario with strong climate and air pollution controls that are aligned with
225 the Paris Climate Agreement, ozone-related acute excess mortality is estimated to account for
226 0.15% (95% eCI: 0.11 to 0.19%) of total deaths in the future period, with total ozone-related
227 mortality increasing by 0.7% (95% eCI: -5 to 7%) between the present (2010-2014) and future
228 (2050-2054) periods. Ozone-related mortality is projected to decrease in 70% of the studied

229 cities, but is notably increasing in several cities in Canada, China, Mexico, and the United
230 States (Fig 2a, Table 1).

231 Under SSP-2.4.5, the middle-of-the-road scenario, ozone-related acute excess mortality is
232 estimated to account for 0.21% (95% eCI: 0.15 to 0.26%) of total deaths in the future period,
233 with total ozone-related mortality increasing by 64% (95% eCI: 49 to 82%) between the present
234 (2010-2014) and future (2050-2054) periods. Ozone-related mortality is projected to increase in
235 75% of the studied locations, with large increases in cities in North America and Asia, and small
236 increases or decreases in cities in Europe and South Africa (Fig 2b, Table 1).

237 Under SSP-3.7.0, the regional rivalry scenario with weak climate and air pollution controls,
238 ozone-related acute excess mortality is estimated to account for 0.21% (95% eCI: 0.16 to
239 0.26%) of total deaths in the future period, with total ozone-related mortality increasing by 94%
240 (95% eCI: 73 to 116%) between the present (2010-2014) and future (2050-2054) periods.
241 Increases in ozone-related excess mortality were projected in 94% of cities included in the
242 analysis, with small to moderate increases in cities in eastern Europe, Japan and southeastern
243 United States, and large increases in cities in all other parts of North America, northwestern
244 Europe, China, Taiwan, and South Africa (Fig 2c, Table 1).

245 Under SSP-5.8.5, a scenario with weak climate controls but strong air pollution controls, ozone-
246 related acute excess mortality is estimated to account for 0.22% (95% eCI: 0.17 to 0.29%) of
247 total deaths in the future period, with total ozone-related mortality increasing by 56% (95% eCI:
248 42 to 87%) between the present (2010-2014) and future (2050-2054) periods. Increases in
249 ozone-related excess mortality were projected in 94% of cities included in the analysis, with
250 small to moderate increases in cities in China and Europe, and large increases in cities in North
251 America and all other parts of Asia (Fig 2d, Table 1).

252 *Drivers of changes in ozone-related excess mortality*

253 Changes in ozone-related mortality are driven by three key factors: changes in emissions and
254 climate, population size, and mortality rates. Here, we isolate the relative contribution of each
255 factor in influencing changes in ozone-related mortality between present and future periods.
256 Though national age-specific baseline mortality rates are generally projected to decline in the
257 studied countries or regions across all SSPs, we observe an increasing proportion of older
258 individuals between 65 and 74 years and 75 years and older (Fig S2). Across the SSPs, age-
259 specific baseline mortality rates for individuals above 75 years are on average 24 to 41 times

260 greater than individuals between 0 and 64 years, resulting in higher overall baseline mortality
261 rates.

262 Under SSP 1-2.6, total ozone-related mortality is estimated increase by 45 deaths/year (CI: -940
263 to 980 deaths/year) between the present period (2010-2014) and future period (2050-2054).
264 Changes in emissions and climate are linked to net decreases in ozone-related mortality
265 fractions (range: -97% to -3% decrease in deaths per year) in all studied countries or regions
266 except China (67% increase in deaths per year) and Canada (3% increase in deaths per year).
267 Changes in population size generally increase ozone-related mortality in North America, South
268 Africa, Sweden, France, Switzerland, Czech Republic, and Spain (range: 3-49% increase in
269 deaths per year) and decrease it in Asia and the remaining European countries or regions
270 (range: -23 to -2% deaths per year). Changes in baseline mortality rates are projected to
271 decrease ozone-related mortality in South Africa, Czech Republic, France, UK, Estonia, and
272 Sweden (range: -17% to -1% deaths per year) and increase it in all other countries or regions
273 (Fig 3a).

274 Under SSP 2-4.5, total ozone-related mortality is estimated to increase by 4,400 deaths/year
275 (CI: 2,300 to 6,600 deaths/year) between the present period (2010-2014) and future period
276 (2050-2054). Changes in emissions and climate are linked to net increases in ozone-related
277 mortality in Asia, North America, UK, Estonia, and Germany (range: 1% to 118% deaths per
278 year), and net decreases in all other countries or regions (range: -48% to -0.4% deaths per
279 year). Changes in population size generally decrease ozone-related mortality in Asia, Germany,
280 Estonia, Greece, Italy, and Portugal (range: -22% to -3% deaths per year), and increase it in
281 North America, South Africa, and the remaining European countries or regions (range: 1% to
282 49% deaths per year). Changes in baseline mortality rates are projected to increase ozone-
283 related mortality in all studied countries or regions (range: 4% to 144% deaths per year) except
284 Sweden (-4% deaths per year) (Fig 3b).

285 Under SSP 3-7.0, total ozone-related mortality is estimated to increase by 6,200 deaths/year
286 (CI: 3,100 to 9,300 deaths/year) between the present period (2010-2014) and future period
287 (2050-2054). Changes in emissions and climate are consistently associated with net increases
288 in ozone-related mortality (range: 4% to 47% increase in deaths per year) at a country- or
289 region-level. Changes in population size slightly decrease ozone-related mortality (-32% to -
290 0.02% decrease in deaths per year) due to the decreasing population size projected in most
291 countries or regions, except for Mexico, South Africa, and Sweden where populations are
292 increasing (0.5% to 46% increase in deaths per year). Decreases in population size are

293 consistently offset by increasing baseline mortality rates (32 to 105%), largely driven by
294 population aging (Fig 3c).

295 Under SSP 5-8.5, total ozone-related mortality is estimated to increase by 3,700 deaths/year
296 (CI: 2,000 to 5,800 deaths/year) between the present period (2010-2014) and future period
297 (2050-2054). Changes in emissions and climate are consistently linked to a net increase in
298 ozone-related mortality (range: 5% to 84% deaths per year). Changes in population size
299 generally decrease ozone-related mortality in Asia and Mexico (range: -23% to -2% deaths per
300 year) and increase it in all other studied countries or regions (range: 20% to 57% deaths per
301 year). Conversely, changes in baseline mortality rates generally increase ozone-related
302 mortality in Asia and Mexico (range: 8% to 90% deaths per year) and decrease it in all other
303 studied countries or regions (range: - 29% to -2% deaths per year) (Fig 3d).

304 **Discussion**

305 Ground-level ozone exposure, associated with premature mortality and morbidity^{3,4,5}, is
306 projected to worsen in many regions due to climate-driven meteorological shifts and changes in
307 ozone precursor emissions⁶. Despite this, a comprehensive estimation of future changes in
308 ozone-related acute excess mortality across diverse geographical locations and under various
309 climate and air quality scenarios using the most recent CMIP6 model ensembles has been
310 lacking. Our study addresses this gap, building on the largest epidemiological study to date⁵, by
311 quantifying future changes in ozone-related acute excess mortality in 406 cities across 20
312 countries or regions. We project an increase in ozone-related deaths in all studied scenarios,
313 ranging from 45 to 6,200 deaths/year in the studied 406 cities, accompanied by rising mortality
314 fractions (0.17% to 0.21-0.22% of total deaths), except in the scenario aligned with the Paris
315 Climate Agreement goals (decreasing from 0.17% to 0.15% of total deaths).

316 Our analysis projects about 6,600 ozone-related deaths per year in the 406 cities in the present
317 period, consistent with a previous study that estimated ozone-related deaths for the same MCC
318 network⁵. This work advances the prior MCC study by estimating future changes in ozone-
319 related acute excess mortality, employing multi-model ensembles under CMIP6 under four
320 distinct SSP scenarios to provide a novel and comprehensive projection of future ozone-related
321 mortality burden. Under SSPs 1-2.6, 2-4.5, 3-7.0, and 5-8.5, ozone-related acute excess
322 mortality in the studied 406 cities, is estimated to increase by 45 deaths/year, 4,400
323 deaths/year, 6,200 deaths/year, and 3,700 deaths/year, respectively. These future estimates
324 are broadly consistent with past research that show small to large increases in health impacts

325 under scenarios with high global warming and emissions of ozone precursors, and moderate
326 decreases to small increases in health impacts under scenarios with low global warming and
327 emissions of ozone precursors^{8,9,10}. However, these studies cannot be directly compared as
328 they were conducted over different geographical boundaries, climate scenarios, socioeconomic
329 pathways and threshold concentrations. We address gaps of previous studies that investigated
330 future trends in ozone-related acute excess mortality by expanding our analysis to a broader
331 geographical scope and considering a wider range of climate and air quality scenarios.

332 Previous studies that examined future ozone-related acute excess mortality were often
333 constrained to long-term exposure metrics linked to specific causes (e.g., respiratory mortality),
334 narrow geographical scopes, limited climate and air quality scenarios, and location-specific
335 ozone-mortality relationships that have been generalized to wider populations. Our study builds
336 on this prior work in several fundamental ways. First, we concentrate on short-term ozone-
337 mortality relationships that encompass a broader spectrum of mortality causes beyond
338 respiratory and cardiovascular disease. Second, we significantly broaden the geographic scope,
339 analyzing 406 cities in 20 countries or regions. Third, we estimate health impacts from ground-
340 level ozone concentrations under four SSP scenarios, which consider a wide array of climate
341 and air pollution controls. Fourth, we incorporate country- or region-specific ozone-mortality
342 relationships derived from a recent MCC study⁵, to address the uncertainty linked to location-
343 specific parameters in air pollution health impact assessments, as underscored previously¹¹.
344 These enhancements produce a more accurate estimate of the overall health impacts of short-
345 term exposure to ozone, which can inform the design of more nuanced health policies.

346 Four primary limitations are encountered in our study: First, while our analysis uses baseline
347 mortality rates that are adjusted for changes in demographic composition, the ozone-mortality
348 relationships we utilize are not age-specific. Performing age-stratified analysis may be key to
349 more fully accounting for ozone-related health burdens given the increased vulnerability of older
350 populations to air pollution exposure¹². Indeed, our study found that changes in baseline
351 mortality rates, largely influenced by changes in age demographics, are often the biggest driver
352 of changes in ozone-related deaths and underscore the need to conduct further analysis that
353 uses age-specific ozone-mortality relationships. Second, our approach of using the 70 $\mu\text{g}/\text{m}^3$
354 threshold based on a previous health impact assessment⁵ also does not incorporate the ozone
355 concentration changes below this threshold. Further epidemiological studies are needed to
356 examine if there are sufficient evidence on the adverse ozone effects on health below this
357 threshold. Third, though the CMIP6 project includes future ozone projections of at least 26

358 models across the four SSPs, most archived ozone fields at a coarser temporal resolution (e.g.,
359 monthly) that is not suitable to evaluate short-term exposure to ozone. Consequently, the
360 analysis is constrained to simulations with 1-5 available global chemistry-climate models,
361 depending on the SSP being modeled. Fourth, although our analysis largely expands on the
362 geographical scope of previous studies, the study cannot be considered a true global analysis
363 as it is limited to the 406 cities within the MCC network for which there are observed ozone
364 concentrations and ozone-mortality relationships.

365 Our findings underscore the need for more stringent air quality regulations, given that current
366 ozone standards in many of the studied countries or regions exceed the $70 \mu\text{g}/\text{m}^3$, a threshold in
367 which concentrations are largely attributed to non-anthropogenic sources and ozone-mortality
368 relationships are established⁵. For instance, in the United States, the Environmental Protection
369 Agency has set the primary and secondary 8-hour standard to approximately $137 \mu\text{g}/\text{m}^3$ ¹³.
370 Similarly, the European Union's Air Quality Directive prescribes a maximum daily 8-hour
371 average concentrations of $120 \mu\text{g}/\text{m}^3$ ¹⁴. Canada's ambient air quality standards are established
372 at $122 \mu\text{g}/\text{m}^3$ ¹⁵, while in Korea, the Ministry of environment regulates ozone concentrations are
373 set to $118 \mu\text{g}/\text{m}^3$ ¹⁶. Higher thresholds of daily maximum 8-hour average concentrations are
374 implemented in China at $160 \mu\text{g}/\text{m}^3$ ¹⁷ and in Mexico at $137 \mu\text{g}/\text{m}^3$ ¹⁸. Our study highlights the
375 need for more rigorous ozone standards. Beyond mitigating ozone-related acute excess
376 mortality, the implementation of stricter air quality regulations will likely yield additional benefits
377 in terms of reducing long-term ozone-related mortality and conferring climate benefits.

378 **Experimental Procedures**

379 *Resource availability*

380 *Lead contact*

381 Further information and requests for resources should be directed to and will be fulfilled by the
382 lead contact, Dr. Nina G. G. Domingo.

383 *Materials availability*

384 This study did not generate new unique materials.

385 *Data and code availability*

386 The projected data on temperature and ozone concentration can be obtained from the CMIP6
387 database (<https://esgf-node.llnl.gov/search/cmip6/>). The projected baseline mortality and

388 population data can be obtained from the Socioeconomic Data and Applications Center
389 ([https://sedac.ciesin.columbia.edu/data/set/popdynamics1-km-downscaled-pop-base-year-](https://sedac.ciesin.columbia.edu/data/set/popdynamics1-km-downscaled-pop-base-year-projection-ssp-2000-2100-rev01/maps/services)
390 [projection-ssp-2000-2100-rev01/maps/services](https://sedac.ciesin.columbia.edu/data/set/popdynamics1-km-downscaled-pop-base-year-projection-ssp-2000-2100-rev01/maps/services)). The historical baseline mortality and
391 population data can be obtained from the United Nations' World Population Prospects 2019
392 Report (<https://population.un.org/wpp/Download/Standard/MostUsed/>). Code used to generate
393 the results are publicly available on Github (https://github.com/CHENlab-Yale/MCC_FutureO3).
394 Any additional information required for reanalyzing the data reported in this paper is available
395 from the lead contact upon reasonable request.

396 *Study design*

397 This analysis estimated future excess mortality from short-term exposure to ozone in 406 cities
398 in 20 countries or regions. The 406 cities in 20 countries or regions are distributed over North
399 America, Europe, Asia, Australia, and Africa.

400 *Historical ozone observations and baseline mortality*

401 Historical ambient ozone observations and baseline mortality counts for 406 cities in 20
402 countries or regions were obtained from the Multi-Country Multi-City (MCC) Collaborative
403 Research Network. Daily observations of historical ozone concentrations from at least one
404 monitor were obtained for each city and presented in terms of maximum daily 8-hour average
405 concentrations. For cities that had more than one monitor, historical ozone concentrations were
406 computed as the average of the observed concentrations from all available monitors. It is also
407 important to note that ozone data in Japan was derived from the measurements of
408 photochemical oxidant, which is primarily ozone ($\geq 90\%$), followed by other species such as
409 peroxy acetyl nitrate (PAN), hydrogen peroxide and organic hydroperoxides. Baseline mortality
410 counts were measured in terms of daily all-cause deaths for cities located in Canada, the Czech
411 Republic, Estonia, France, Germany, Greece, Italy, Japan, Mexico, Portugal, South Africa,
412 South Korea, Sweden, Taiwan, UK, and US; daily deaths due to non-external causes for cities
413 in Australia, China, and Spain; and daily deaths due to non-external causes other than
414 unintentional injuries for cities in Switzerland.

415 Historical ozone observations and baseline mortality counts were available in largely
416 overlapping periods between January 1, 1985, to December 31, 2015, across the 406 cities. To
417 minimize computational demands of the analysis, we utilized data from the last three full and
418 consecutive years for each city to bias correct ozone concentrations and estimated ozone-
419 related acute excess mortality in present (2010-2014) and future (2050-2054) periods. The mid-

420 century timeframe was selected to represent the future period to ensure the relevancy of results
421 to present-day policy formation.

422 *Present and future ozone projections*

423 We obtained global ozone simulations performed with five global chemistry-climate models from
424 the Aerosols and Chemistry Model Intercomparison Project (AerChemMIP) under the Coupled
425 Model Intercomparison Project Phase 6 (CMIP6). These five chemistry-climate models include
426 the Community Earth System Model version 2 (CESM2), EC-Earth3-AerChem, Geophysical
427 Fluid Dynamics Laboratory Earth System Model version 4 (GFDL-ESM4), Max Planck Institute
428 Earth System Model (MPI-ESM1-2-HAM), and U.K. Earth System Model (UKESM-1-0-LL).
429 Surface ozone concentrations were available at an hourly temporal scale, which we used to
430 calculate MDA8 ozone concentrations. We then applied a constant scaling factor of (1 ppb =
431 $1.96 \mu\text{g}/\text{m}^3$) to convert the molar mixing ratios to mass densities. Further information on each of
432 the models can be found in Table S3. For this analysis, historical simulations for the years 2010-
433 2014 were utilized to represent the present period and simulations corresponding to SSPs 1-
434 2.6, 2-4.5, 3-7.0 and 5-8.5 for the years 2050-2054 were utilized to represent the future period.
435 While it would be ideal to model time periods exceeding five years, our analysis is limited to a
436 five-year time frame due to the constraints in computational resources. Different numbers of
437 models are available for each SSP, and even when a model simulated multiple SSPs, they
438 provided different numbers of ensemble members. Despite this limitation, we chose to use
439 ozone projections from all available models and ensemble members in our study to maximize
440 the number of models we could include.

441 The SSPs were developed for the CMIP6 in 2017¹⁹ and thus do not explicitly incorporate climate
442 and air quality policies or market changes (e.g., electric vehicles) that have occurred since then.
443 Nevertheless, these changes fall within the wide range of possible future spanned by the SSPs,
444 which were designed to cover different possible future socioeconomic pathways¹⁹.

445 As applied in a previous study⁸, we performed a bias-correction technique on the modeled
446 ozone concentrations from the five global chemistry-climate models to obtain ozone
447 concentrations for each of the 406 cities in both present and future periods. Bias correction
448 techniques increase the utility of publicly available models, which are still imperfect
449 mathematical representations of the Earth's climate system. Biases in climate models arise due
450 to simplified representations of atmospheric chemistry and physics, coarse spatial resolution,
451 and incomplete understanding of the global climate system²⁰. A recent global study

452 demonstrated that after bias correction using a similar quantile mapping approach, the data
453 accuracy of ozone concentrations improved pronouncedly with increases in the correlation
454 coefficient and decreases in the root-mean-square error between the observational and
455 modeled data²¹.

456 To perform the bias-correction step, we first computed monthly biases between the observed
457 ozone concentrations and modeled ozone concentrations. Specifically, monthly biases were
458 computed for each quantile of the modeled ozone concentrations within the boundaries of the
459 grid cell from each chemistry-climate model. Assuming historical monthly biases persist
460 throughout time, we then corrected for these biases in present and future time periods.

461 *Changes in baseline mortality counts*

462 We computed daily baseline mortality counts in each of the 406 cities in present and future
463 periods as follows:

$$464 \quad M_{p/f} = M_h \times \Delta Y_b \times \Delta POP \quad (\text{Equation 1})$$

465 Where $M_{p/f}$ is the city-level daily baseline mortality count in the present or future periods, M_h is
466 the city-level daily baseline mortality count corresponding to the last full and consecutive three
467 years of available historical data, ΔY_b is the change in country- or region-level annual baseline
468 mortality rate between the historical period and present or future periods, and ΔPOP is the
469 change in country- or region-level population between the historical period and present or future
470 periods. We trimmed the daily baseline mortality count to the last full and consecutive three
471 years from the MCC network to be consistent with the criteria we used for the observed ozone
472 concentration data in the bias correction stage.

473 Country- or region-level projections of baseline mortality rates and population across the four
474 SSPs were obtained from the Shared Socioeconomics Pathway project and historical estimates
475 of baseline mortality and population were obtained from the United Nations' World Population
476 Prospects 2019 Report.

477 *Health impact assessment*

478 We estimated the daily ozone-related excess mortality in present and future periods using the
479 Attributable Fraction (AF) approach. The AF is defined as the share of the baseline mortality
480 that is attributable to short-term exposure and is computed as follows:

$$481 \quad AF = 1 - \exp^{-\beta \times C} \quad (\text{Equation 2})$$

482 Where C is the difference between the city-level maximum daily 8-hour average concentration
483 for ozone and the maximum background levels of $70 \mu\text{g}/\text{m}^3$, and β corresponds to the logarithm
484 of the relative risk at ozone concentration C . In this analysis, country- or region-specific
485 exposure-response functions are obtained from⁵ for ozone concentrations above maximum
486 background levels. Exposure-response functions in this study were obtained through a two-
487 stage time-series analysis of about 45 million deaths. In the first stage, the study runs separate
488 time series regression models to obtain city-specific ozone-mortality risks. In the second stage,
489 the study pools the city-specific estimates through a meta-analysis to derive best linear
490 unbiased predictions of the risks at a country-or region-level. We prioritize using country- or
491 region-specific ozone-mortality relationships in our study as Vicedo-Cabrera et al. found that
492 these relationships were spatially heterogenous (range: 1.0008 to 1.0035 relative risk per 10
493 $\mu\text{g}/\text{m}^3$ increase in ozone)⁵. These spatial differences in ozone-mortality relationships may be
494 driven by local conditions, including concentrations and sources of ozone precursors, and
495 population characteristics—which we would not be able to capture by extrapolating a single
496 ozone-mortality relationship to the entire globe.

497 For global chemistry-climate models that have multiple ensemble members, and thus multiple
498 estimates of concentrations for the same period and city, we use the average of all bias-
499 corrected concentrations across all available ensemble members.

500 The attributable daily deaths due to short-term ozone exposure in each city and period is then
501 calculated as follows:

$$502 \quad ADD = M_{p/f} \times AF \text{ (Equation 3)}$$

503 Where ADD is the estimated city-level number of daily deaths attributed in present and future
504 periods, and $M_{p/f}$ and AF are as defined above. Annual attributable mortality is then computed
505 by summing the ADD throughout the year in each period. Lastly, we computed the absolute and
506 percentage change in ozone-related excess mortality between present and future periods.

507 Uncertainty in mortality estimates were quantified using Monte Carlo simulations that
508 incorporated uncertainty from coefficients in the exposure-response functions and inter-model
509 variability. For absolute mortality estimates, we derived confidence intervals at a 90%
510 confidence level from the distribution across 1,000 coefficient samples, assuming a normal
511 distribution, for each of the five global chemistry-climate models. To derive confidence intervals
512 for changes in mortality over time, we subtract the mean mortality estimate in the present period
513 from the lower or upper bound mortality estimate in the future period.

514 As a sensitivity test, we compared the ozone-related excess mortality in the present period
515 using both the raw simulated ozone concentrations and bias-corrected ozone concentrations.
516 To minimize computational requirements, we limit the sensitivity test to the four cities that are
517 associated with the highest ozone-related mortality in the main analysis. Across the four cities,
518 ozone-related excess mortality differed by -11 to 550% between the two ozone datasets (Table
519 S2), therefore, we elected to limit our main analysis to the bias-corrected ozone concentrations.

520 Changes in estimated ozone-related mortality between present and future periods can be
521 attributed to three factors—emissions and climate, population size, and mortality rates. We
522 decomposed the impacts of each of these three factors by first running a scenario with changing
523 emissions and climate and fixed population and mortality rates to isolate the impact of
524 emissions and climate. All else held constant, changes in ozone-related mortality are directly
525 proportional to changes in population size and changes in mortality rates and could be readily
526 isolated without further computation.

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538 **Declaration of interests**

539 The authors declare no competing interests.

540 **Declaration of Generative AI and AI-assisted technologies in the writing process**

541 To improve the readability and clarity of the manuscript, the authors incorporated suggestions
542 from the language models, Chat GPT-3.5 and Bard. After using this tool/service, the author(s)
543 reviewed and edited the content as needed and takes full responsibility for the content of the
544 publication.

545

546 **References**

- 547 1. Fuller R., Landrigan P.J., Balakrishnan K., Bathan G., Bose-O'Reilly S., Brauer M.,
548 Caravanos, J., Chiles, T., Cohen, A., Corra, L., et al. (2022). Pollution and health: a
549 progress update. *Lancet Planet. Health* 6, E535-E547. [https://doi.org/10.1016/S2542-](https://doi.org/10.1016/S2542-5196(22)00090-0)
550 [5196\(22\)00090-0](https://doi.org/10.1016/S2542-5196(22)00090-0).
- 551 2. Zhang J., Wei Y., and Fang Z. (2019). Ozone Pollution: A Major Health Hazard
552 Worldwide. *Front Immunol.* 10, 2518. <https://doi.org/10.3389/fimmu.2019.02518>.
- 553 3. Orellano P., Reynoso J., Quaranta N., Bardach A., and Ciapponi A. (2020). Short-term
554 exposure to particulate matter (PM10 and PM2.5), nitrogen dioxide (NO2), and ozone
555 (O3) and all-cause and cause-specific mortality: Systematic review and meta-analysis.
556 *Environ Int.* 142, 105876. <https://doi.org/10.1016/j.envint.2020.105876>.
- 557 4. Bell M.L. (2004). Ozone and Short-term Mortality in 95 US Urban Communities, 1987-
558 2000. *JAMA* 292, 2372-2378. <https://doi.org/10.1001/jama.292.19.2372>.
- 559 5. Vicedo-Cabrera A.M., Sera F., Liu C., Armstrong B., Milojevic A., Guo Y., Tong, S.,
560 Lavigne, E., Kysely, J., Urban, A., et al. (2020). Short term association between ozone
561 and mortality: global two stage time series study in 406 locations in 20 countries. *Br.*
562 *Med. J.* 368, m108. <https://doi.org/10.1136/bmj.m108>.
- 563 6. Fiore A.M., Naik V., and Leibensperger E.M. (2015). Air Quality and Climate
564 Connections. *J Air Waste Manag Assoc.* 65, 645-685.
565 <https://doi.org/10.1080/10962247.2015.1040526>.
- 566 7. Zanis P., Dimitris A., Turnock S.T., Naik V., Szopa S., Georgoulas A.K., Bauer, S.,
567 Deushi, M., Horowitz, L., and Keeble J. (2022). Climate change penalty and benefit on
568 surface ozone: a global perspective based on CMIP6 earth system models. *Environ.*
569 *Res. Lett.* 17, 024014. <https://doi.org/10.1088/1748-9326/ac4a34>.
- 570 8. Chen K., Fiore A.M., Chen R., Jiang L., Jones B., Schneider A., Peters, A., Bi, J., Kan,
571 H., and Kinney, P. (2018). Future ozone-related acute excess mortality under climate
572 and population change scenarios in China: A modeling study. *PLoS Med.* 15, e1002598.
573 <https://doi.org/10.1371/journal.pmed.1002598>.
- 574 9. Fann N.L., Nolte C.G., Sarofim M.C., Martinich J., and Nassikas N.J. (2021).
575 Associations Between Simulated Future Changes in Climate, Air Quality, and Human
576 Health. *JAMA* 4, e2032064. <https://doi.org/10.1001/jamanetworkopen.2020.32064>.
- 577 10. Chang H.H., Zhou J., and Fuentes M. (2010). Impact of Climate Change on Ambient
578 Ozone Level and Mortality in Southeastern United States. *Int. J. Environ. Res. Public*
579 *Health* 7, 2866-2880. <https://doi.org/10.3390/ijerph7072866>.
- 580 11. Pozzer A., Anenberg S.C., Dey S., Haines A., Lelieveld J., and Chowdhury S. (2022).
581 Mortality attributable to ambient air pollution: A review of global estimates. *GeoHealth* 7,
582 e2022GH000711. <https://doi.org/10.1029/2022GH000711>.
- 583 12. Chen K., Vicedo-Cabrera A.M., and Dubrow R. (2020). Projections of Ambient
584 Temperature- and Air Pollution-Related Mortality Burden Under Combined Climate
585 Change and Population Aging Scenarios: a Review. *Curr. Environ. Health Rep.* 7, 243-
586 255. <https://doi.org/10.1007/s40572-020-00281-6>.
- 587 13. Environmental Protection Agency (2020). Review of the Ozone National Ambient Air
588 Quality Standards. [https://www.govinfo.gov/content/pkg/FR-2020-12-31/pdf/2020-](https://www.govinfo.gov/content/pkg/FR-2020-12-31/pdf/2020-28871.pdf)
589 [28871.pdf](https://www.govinfo.gov/content/pkg/FR-2020-12-31/pdf/2020-28871.pdf).
- 590 14. Official Journal of the European Union (2008). Directive 2008/50/EC of the European
591 Parliament and of the Council of 21 May 2008 on Ambient Air Quality and Cleaner Air for

- 592 Europe. <https://eur-lex.europa.eu/legal->
593 [content/EN/TXT/PDF/?uri=CELEX:32008L0050&qid=1696086918600](https://eur-lex.europa.eu/legal-content/EN/TXT/PDF/?uri=CELEX:32008L0050&qid=1696086918600).
- 594 15. State of Environment Reporting, Ministry of Environment and Climate Change Strategy,
595 British Columbia, Canada (2022). Status of Ground-Level Ozone in B.C. (2018-2020).
596 [https://www.env.gov.bc.ca/soe/indicators/air/ozone.html#:~:text=Ground%2DLevel%20O](https://www.env.gov.bc.ca/soe/indicators/air/ozone.html#:~:text=Ground%2DLevel%20Ozone:%20Canadian%20Ambient)
597 [zone:%20Canadian%20Ambient](https://www.env.gov.bc.ca/soe/indicators/air/ozone.html#:~:text=Ground%2DLevel%20Ozone:%20Canadian%20Ambient).
- 598 16. Yeo, M.J., and Kim, Y.P. (2021). Long-term trends of surface ozone in Korea. *J. Clean.*
599 *Prod.* 1, 125352. [https://doi: 10.1016/j.jclepro.2020.125352](https://doi.org/10.1016/j.jclepro.2020.125352).
- 600 17. Wang Z., Tan Y., Guo M., Cheng M., Gu Y., Chen S., Wu, X., and Chai, F.
601 (2022). Prospect of China's ambient air quality standards. *J Environ. Sci.* 1, 255–
602 269. [https:// doi: 10.1016/j.jes.2022.03.036](https://doi.org/10.1016/j.jes.2022.03.036).
- 603 18. Diario Oficial de la Federación (2014). NORMA Oficial Mexicana NOM-020-SSA1-2014,
604 Salud ambiental. Valor límite permisible para la concentración de ozono (O3) en el aire
605 ambiente y criterios para su evaluación.
606 https://dof.gob.mx/nota_detalle.php?codigo=5356801&fecha=19/08/2014#gsc.tab=0.
- 607 19. O'Neill B.C., Kriegler E., Riahi K., Ebi K.L., Hallegatte S., Carter T.R., Mathur, R.,
608 and van Vuuren, D.P. (2013). A new scenario framework for climate change
609 research: the concept of shared socioeconomic pathways. *Clim. Change.* 122,
610 387-400. <https://doi.org/10.1007/s10584-013-0905-2>.
- 611 20. IPCC (2021). Summary for Policymakers. *Climate Change 2021: The Physical Science*
612 *Basis. Contribution of Working Group I to the Sixth Assessment Report of the*
613 *Intergovernmental Panel on Climate Change (Cambridge University Press)*, pp. 3–32.
- 614 21. Ban J., Lu K., Wang Q., and Li T. (2022). Climate change will amplify the inequitable
615 exposure to compound heatwave and ozone pollution. *One Earth* 5, 677–686.
616 <https://doi.org/10.1016/j.oneear.2022.05.007>.

617

618 **Fig 1. Annual average MDA8 O₃ concentration at 406 locations in 20 countries or regions**
619 **in present (2010-2014) and future (2050-2054) time periods in the AerChemMIP models.**
620 (A) present O₃ concentrations in each city, (B) absolute change in O₃ concentrations under SSP
621 1-2.6, (C) absolute change in O₃ concentrations under SSP 2-4.5, (D) absolute change in O₃
622 concentrations under SSP 3-7.0, and (E) absolute change in O₃ concentrations under SSP 5-
623 8.5. O₃ concentration is the maximum daily 8-hour average.

624 **Fig 2. Change in O₃-related excess mortality between present (2010-2014) and future**
625 **(2050-2054) time periods in 406 locations in 20 countries or regions.** Absolute changes in
626 O₃-related excess mortality under (A) SSP 1-2.6, (B) SSP 2-4.5, (C) SSP 3-7.0, and (D) SSP 5-
627 8.5.

628

629 **Fig 3. Changes in O₃-related mortality between present (2010-2014) and future (2050-**
630 **2054) time periods allocated to changes in emissions and climate, mortality rates, and**
631 **population.** City-level changes in O₃-related mortality under (A) SSP 1-2.6, (B) SSP 2-4.5, (C)
632 SSP 3-7.0, and (D) SSP 5-8.5 are aggregated to the country- or region-level.

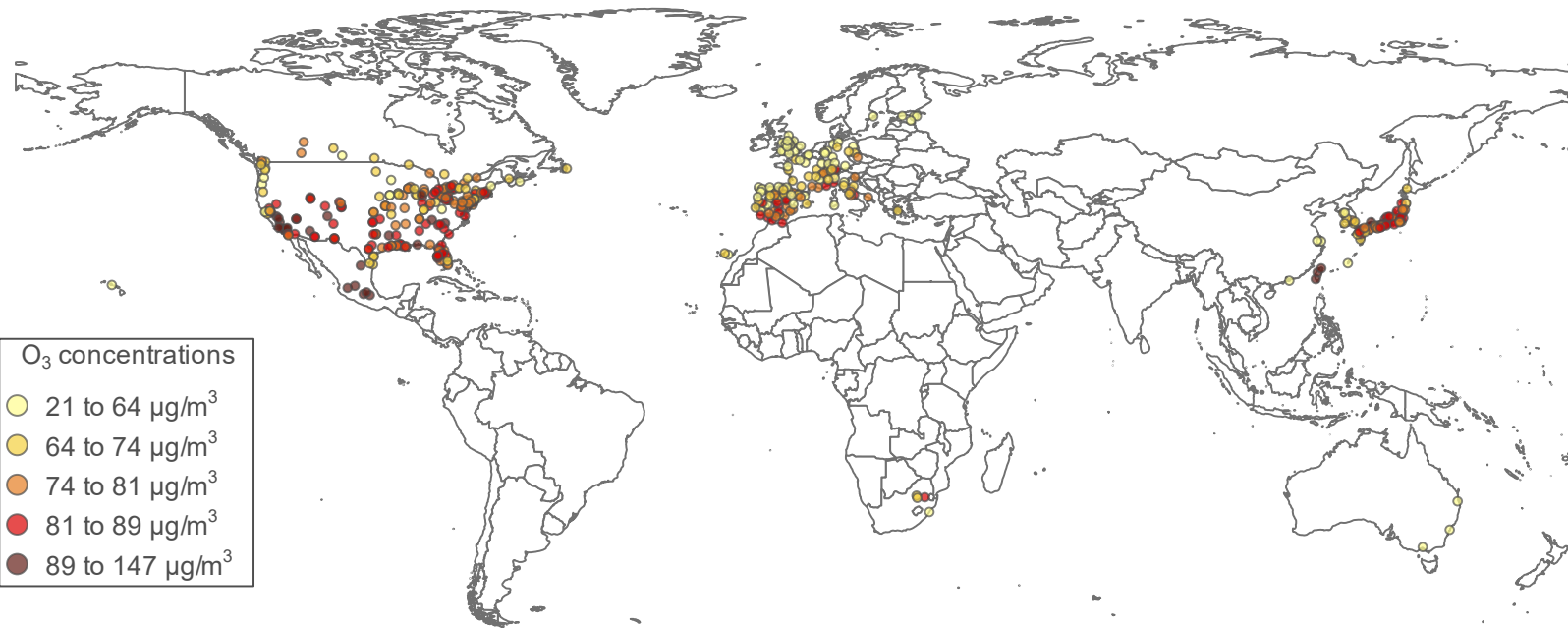
Table 1. Change in O₃-related mortality between present (2010-2014) and future (2050-2054) time periods under each SSP. City-level changes in O₃-related mortality are aggregated to the country-or region-level. Values are rounded to the nearest whole number.

Country or region	Number of cities	Mean change in O ₃ -related mortality (deaths/yr)				Change in O ₃ -related mortality (%)			
		SSP 1-2.6	SSP 2-4.5	SSP 3-7.0	SSP 5-8.5	SSP 1-2.6	SSP 2-4.5	SSP 3-7.0	SSP 5-8.5
Australia	3	0 (CI: 0 to 0)	0 (CI: 0 to 0)	0 (CI: 0 to 0)	0 (CI: 0 to 0)	-	-	-	-
Canada	26	157 (CI: 101 to 215)	415 (CI: 271 to 568)	441 (CI: 284 to 603)	323 (CI: 209 to 454)	46% (CI: 30% to 64%)	120% (CI: 78% to 164%)	132% (CI: 85% to 181%)	93% (CI: 60% to 131%)
China	3	59 (CI: 19 to 98)	115 (CI: 37 to 188)	104 (CI: 32 to 178)	80 (CI: 26 to 136)	142% (CI: 44% to 233%)	274% (CI: 88% to 448%)	179% (CI: 56% to 310%)	168% (CI: 54% to 285%)
Czech Republic	1	-19 (CI: -33 to -5)	5 (CI: 2 to 9)	17 (CI: 5 to 28)	6 (CI: 2 to 11)	-59% (CI: -100% to -16%)	16% (CI: 5% to 28%)	54% (CI: 15% to 93%)	21% (CI: 6% to 37%)
Estonia	4	0 (CI: 0 to 0)	0 (CI: 0 to 0)	0 (CI: 0 to 0)	0 (CI: 0 to 0)	-98% (CI: -157% to -35%)	-2% (CI: -5% to 6%)	4% (CI: 1% to 31%)	19% (CI: 6% to 37%)
France	18	-78 (CI: -122 to -32)	8 (CI: 3 to 19)	98 (CI: 40 to 153)	61 (CI: 25 to 104)	-68% (CI: -107% to -28%)	6% (CI: 2% to 16%)	75% (CI: 31% to 118%)	40% (CI: 17% to 68%)
Germany	12	-63 (CI: -97 to -29)	13 (CI: 6 to 21)	78 (CI: 37 to 122)	28 (CI: 12 to 46)	-64% (CI: -100% to -30%)	14% (CI: 6% to 22%)	77% (CI: 36% to 121%)	24% (CI: 11% to 40%)
Greece	1	-31 (CI: -69 to 0)	-12 (CI: -26 to 0)	12 (CI: 1 to 27)	9 (CI: 0 to 21)	-79% (CI: -176% to 0%)	-28% (CI: -62% to 0%)	17% (CI: 2% to 63%)	18% (CI: 0% to 44%)
Italy	13	-118 (CI: -203 to -35)	-35 (CI: -61 to -9)	59 (CI: 17 to 102)	45 (CI: 13 to 85)	-83% (CI: -144% to -24%)	-25% (CI: -43% to -7%)	37% (CI: 11% to 67%)	30% (CI: 9% to 57%)
Japan	43	-339 (CI: -489 to -201)	318 (CI: 187 to 470)	614 (CI: 366 to 897)	320 (CI: 185 to 499)	-35% (CI: -50% to -20%)	32% (CI: 19% to 47%)	64% (CI: 38% to 93%)	31% (CI: 18% to 48%)
Mexico	8	378 (CI: 90 to 661)	800 (CI: 183 to 1409)	1385 (CI: 329 to 2433)	415 (CI: 87 to 770)	44% (CI: 10% to 77%)	93% (CI: 21% to 164%)	160% (CI: 39% to 288%)	51% (CI: 11% to 94%)
Portugal	6	-26 (CI: -54 to 0)	0 (CI: 0 to 3)	18 (CI: 1 to 37)	8 (CI: 0 to 17)	-88% (CI: -185% to 0%)	1% (CI: 0% to 9%)	45% (CI: 4% to 116%)	26% (CI: 0% to 56%)

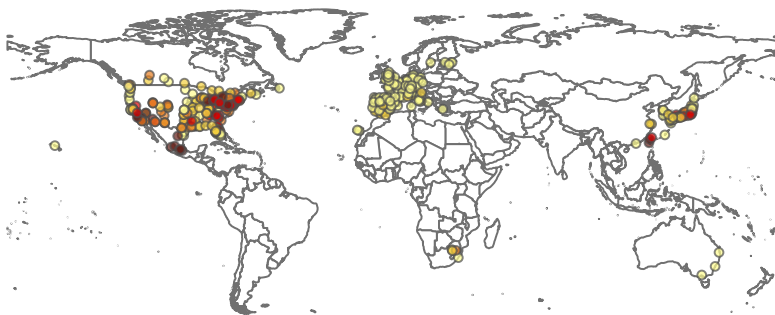
South Africa	4	-43 (CI: -62 to -24)	-23 (CI: -34 to -13)	105 (CI: 57 to 158)	30 (CI: 15 to 59)	-45% (CI: -64% to -24%)	-24% (CI: -35% to -13%)	97% (CI: 53% to 146%)	28% (CI: 14% to 55%)
South Korea	7	-30 (CI: -49 to -10)	86 (CI: 29 to 142)	188 (CI: 65 to 310)	116 (CI: 38 to 193)	-35% (CI: -58% to -12%)	102% (CI: 34% to 167%)	204% (CI: 71% to 339%)	133% (CI: 43% to 221%)
Spain	47	-32 (CI: -83 to 0)	2 (CI: 0 to 5)	29 (CI: 1 to 74)	19 (CI: 0 to 50)	-81% (CI: -212% to 0%)	5% (CI: 0% to 13%)	41% (CI: 3% to 193%)	49% (CI: 0% to 129%)
Sweden	1	-4 (CI: -7 to -1)	0 (CI: 0 to 0)	4 (CI: 1 to 6)	4 (CI: 1 to 6)	-91% (CI: -151% to -29%)	1% (CI: 0% to 7%)	80% (CI: 25% to 134%)	64% (CI: 20% to 109%)
Switzerland	8	-20 (CI: -33 to -6)	8 (CI: 2 to 13)	30 (CI: 9 to 52)	16 (CI: 5 to 27)	-70% (CI: -119% to -22%)	28% (CI: 9% to 47%)	104% (CI: 33% to 181%)	56% (CI: 17% to 95%)
Taiwan	3	-4 (CI: -8 to 0)	180 (CI: 7 to 335)	283 (CI: 19 to 529)	147 (CI: 6 to 279)	-2% (CI: -3% to 0%)	79% (CI: 3% to 146%)	106% (CI: 8% to 216%)	76% (CI: 3% to 145%)
UK	15	-17 (CI: -21 to -12)	10 (CI: 8 to 14)	30 (CI: 22 to 38)	18 (CI: 13 to 26)	-85% (CI: -108% to -64%)	51% (CI: 38% to 68%)	97% (CI: 72% to 123%)	82% (CI: 60% to 117%)
USA	183	273 (CI: 183 to 364)	2516 (CI: 1687 to 3408)	2689 (CI: 1833 to 3579)	2076 (CI: 1362 to 2973)	8% (CI: 5% to 10%)	69% (CI: 46% to 93%)	80% (CI: 55% to 106%)	60% (CI: 40% to 87%)

Figure

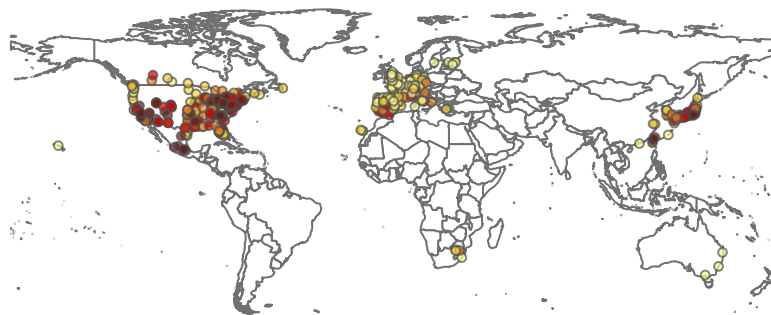
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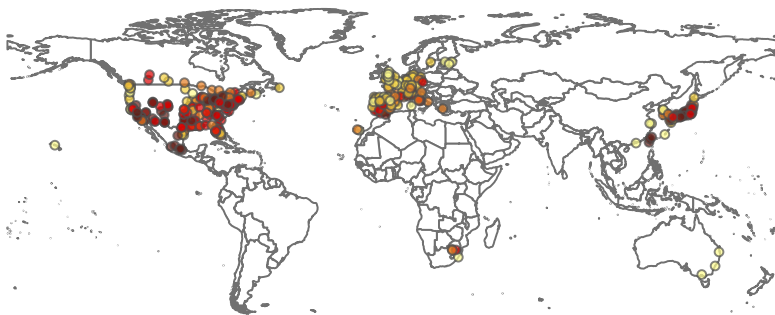
B



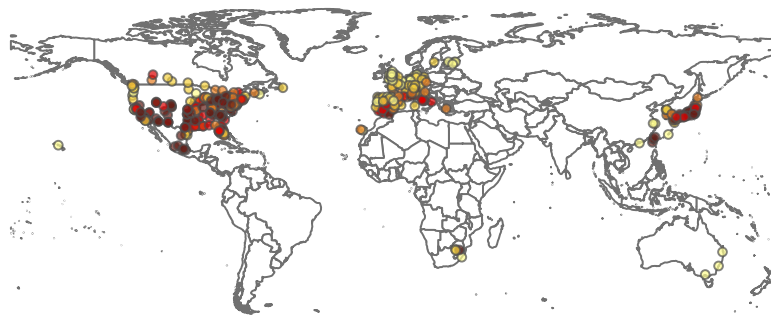
C



D

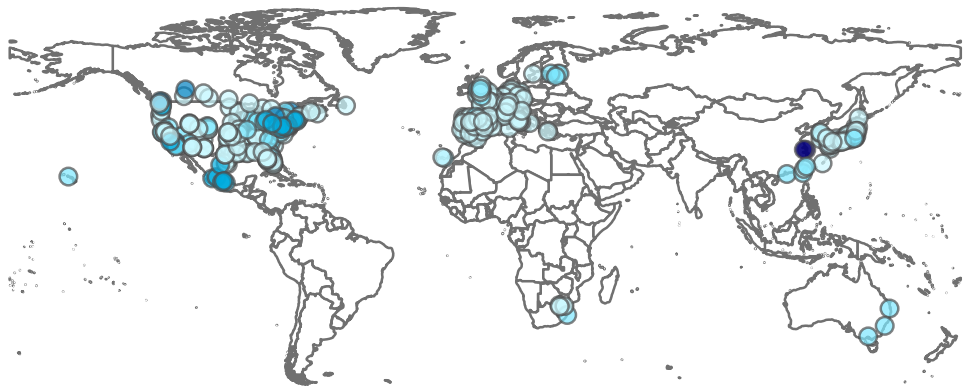


E

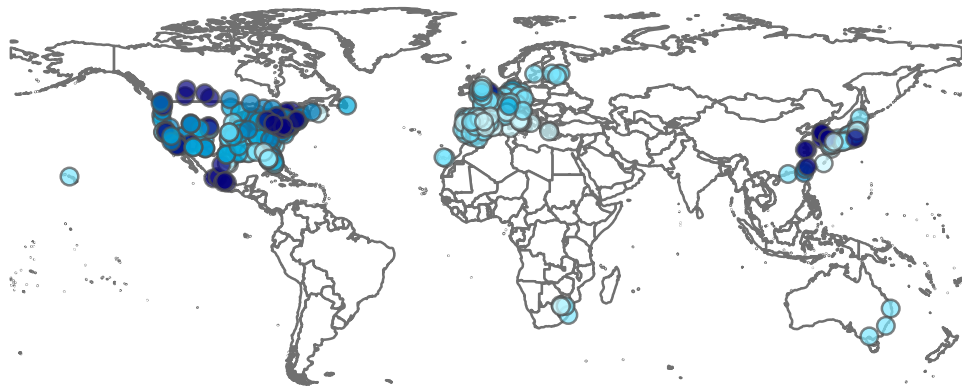


Figure

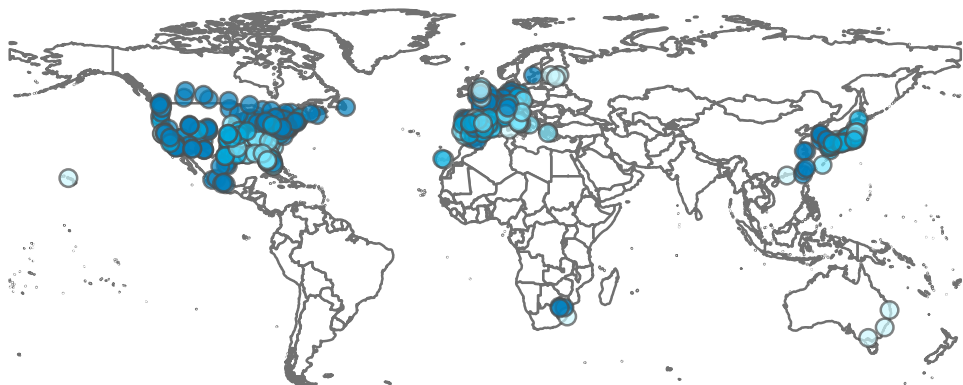
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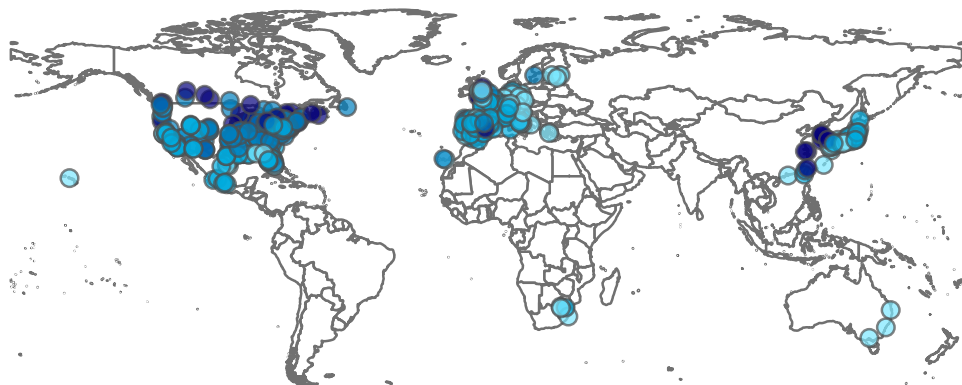
B



C



D



O₃-related mortality ○ -109 to -8% ● -8 to 27% ● 27 to 56% ● 56 to 79% ● 79 to 442%

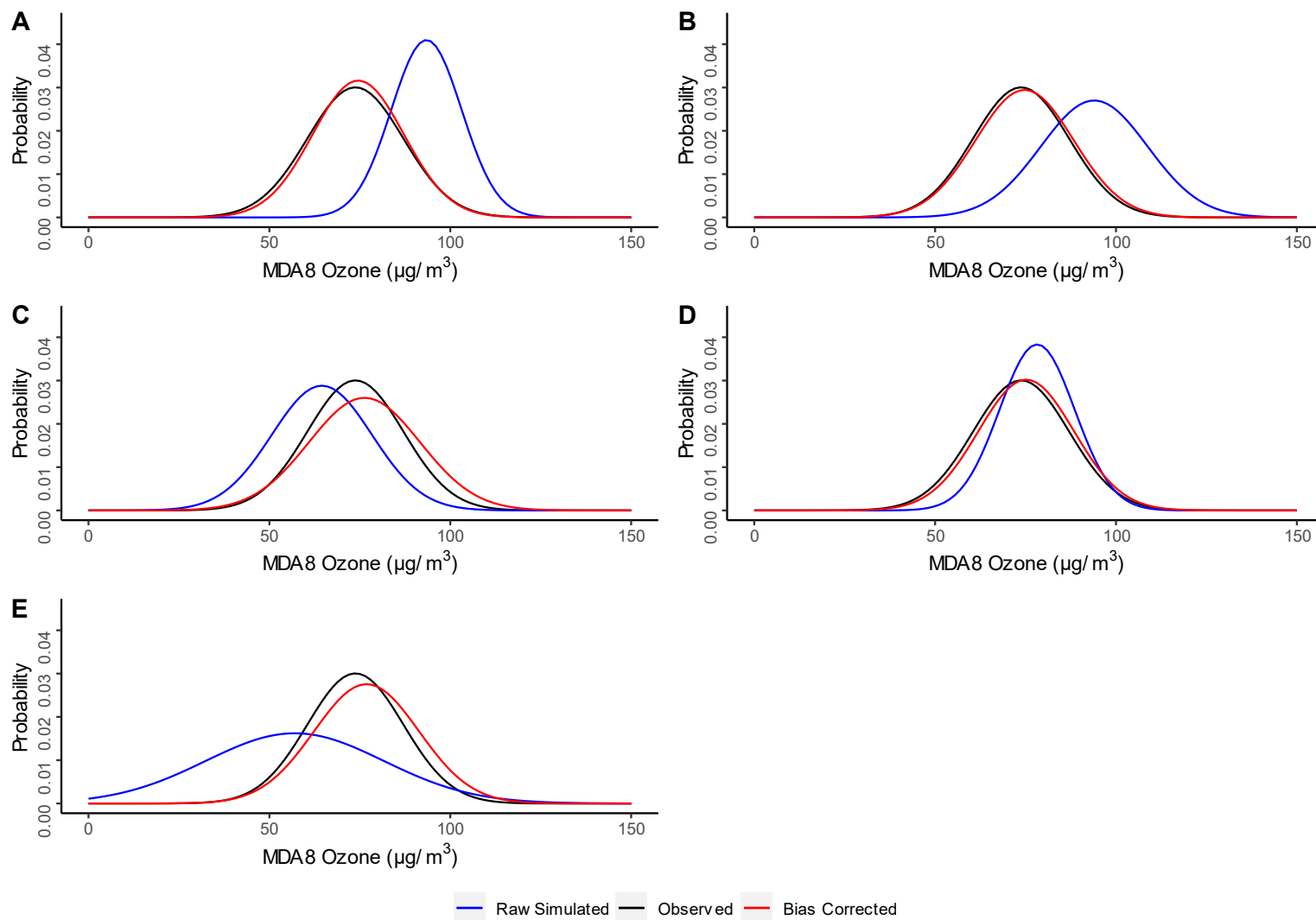


Fig S1. Raw simulated, observed, and bias corrected O_3 concentrations averaged over the 406 cities for each global chemistry-climate model in the present period. (A) CESM2, (B) EC-Earth3-AerChem, (C) GFDL-ESM4, (D) MPI-ESM-1-2-HAM, and (E) UKESM-1-0-LL.

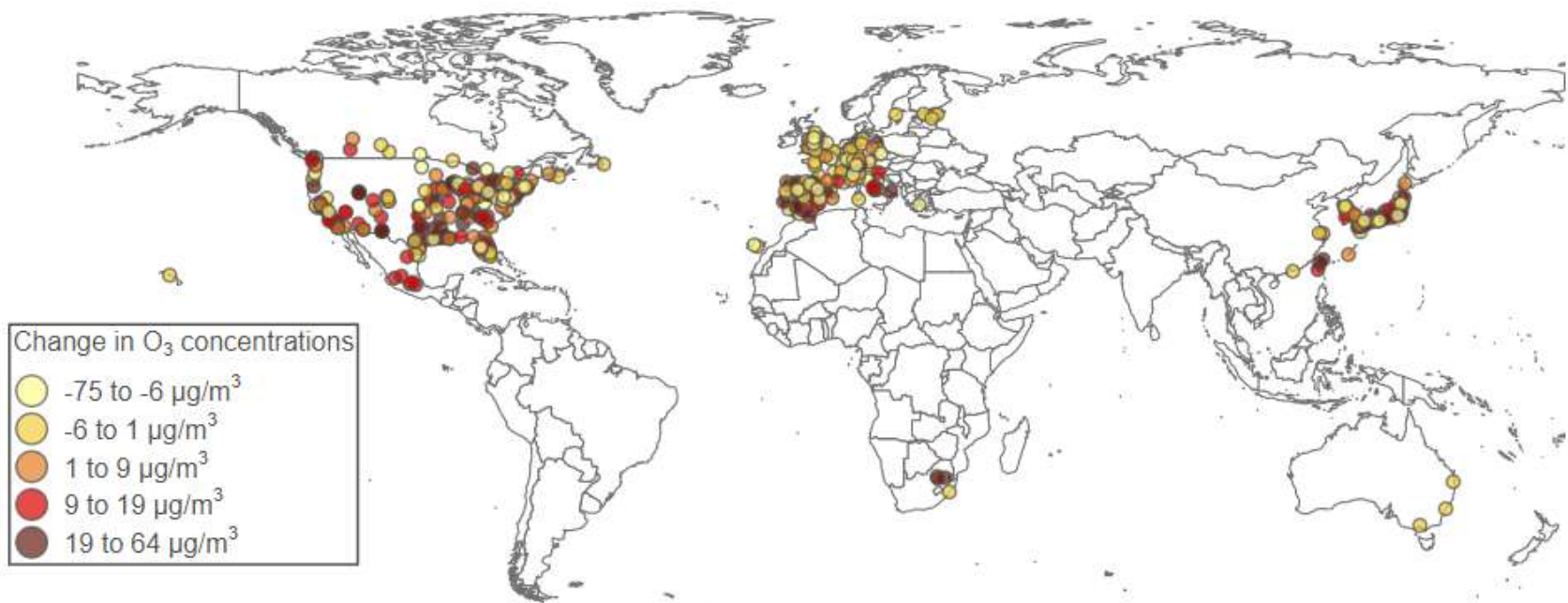


Fig S2. O₃ bias over the 406 cities average across the global-chemistry climate models in the present period. Global chemistry-climate models include CESM2, EC-Earth3-AerChem, GFDL-ESM4, MPI-ESM-1-2-HAM, and UKESM-1-0-LL.

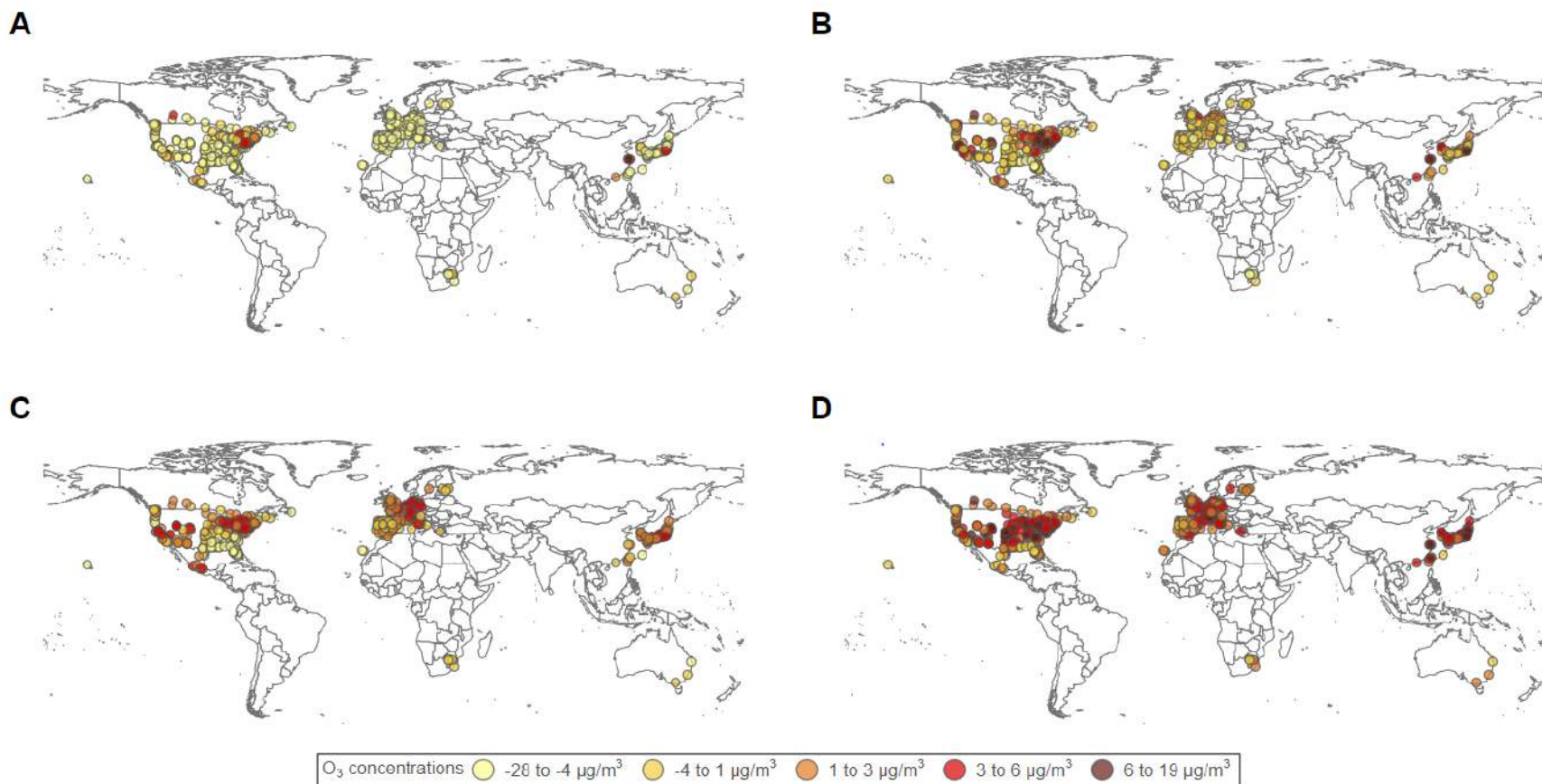


Fig S3. Absolute change in MDA8 O₃ concentration at 406 locations in 20 countries between the present (2010-2014) and future (2050-2054) time periods. (A) absolute change in O₃ concentrations under SSP 1-2.6, (B) absolute change in O₃ concentrations under SSP 2-4.5, (C) absolute change in O₃ concentrations under SSP 3-7.0, and (D) absolute change in O₃ concentrations under SSP 5-8.5. O₃ concentration is the maximum daily 8-hour average.

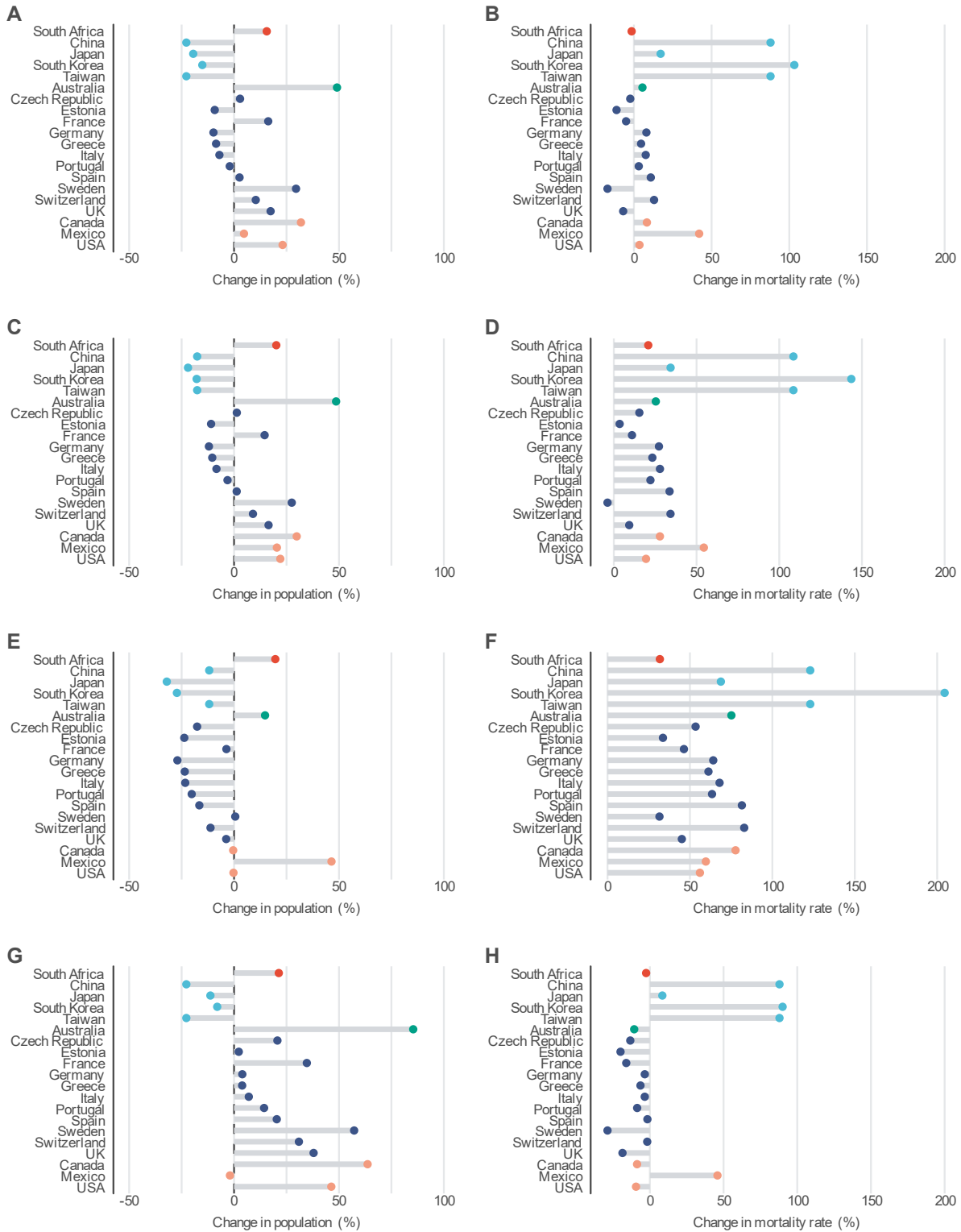


Fig S4. Change in population and mortality rates in 20 countries. Change in national population under (A) SSP 1-2.6, (C) SSP 2-4.5, (E) SSP 3-7.0, and (G) SSP 5-8.5. Change in national mortality rates under (B) SSP 1-2.6, (D) SSP 2-4.5, (F) SSP3-7.0, and (H) SSP 5-8.5.

Table S1. Change in O₃ concentrations between present (2010-2014) and future (2050-2054) time periods. City-level changes in O₃ concentrations are aggregated to the country level and rounded to the nearest whole number.

Country	Number of cities	O ₃ concentrations (µg/m ³)				
		Present data	SSP 1-2.6	SSP 2-4.5	SSP 3-7.0	SSP 5-8.5
Australia	3	34	29	33	36	35
Canada	26	81	84	88	87	89
China	3	58	68	69	63	63
Czech Republic	1	75	58	76	82	79
Estonia	4	57	50	56	60	59
France	18	70	53	66	75	75
Germany	12	62	52	63	71	68
Greece	1	74	54	69	78	81
Italy	13	72	49	66	79	79
Japan	43	80	77	87	88	90
Mexico	8	133	132	135	141	131
Portugal	6	76	59	73	79	78
South Africa	4	78	73	73	81	81
South Korea	7	69	63	72	74	75
Spain	47	73	56	71	77	77
Sweden	1	61	53	62	65	67
Switzerland	8	74	54	70	82	78
Taiwan	3	109	95	108	113	107
UK	15	59	47	59	64	61

USA	183	84	85	92	89	92
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Table S2. O₃-related mortality by global climate model in 4 cities in the present (2010-2014) period.

Global climate model	City	O ₃ -related mortality (deaths/yr)	
		Raw simulated O ₃	Bias corrected O ₃
CESM2	Valley of Mexico, Mexico	323	535
EC-Earth3-AerChem	Valley of Mexico, Mexico	67	530
GFDL-ESM4	Valley of Mexico, Mexico	11	614
MPI-ESM-1-2-HAM	Valley of Mexico, Mexico	28	584
UKESM-1-0-LL	Valley of Mexico, Mexico	0	524
CESM2	Los Angeles, USA	253	223
EC-Earth3-AerChem	Los Angeles, USA	227	320
GFDL-ESM4	Los Angeles, USA	31	303
MPI-ESM-1-2-HAM	Los Angeles, USA	227	178
UKESM-1-0-LL	Los Angeles, USA	0	225
CESM2	Tokyo, Japan	203	133
EC-Earth3-AerChem	Tokyo, Japan	312	151
GFDL-ESM4	Tokyo, Japan	218	144
MPI-ESM-1-2-HAM	Tokyo, Japan	82	162
UKESM-1-0-LL	Tokyo, Japan	63	193
CESM2	Riverside, USA	118	132
EC-Earth3-AerChem	Riverside, USA	97	123
GFDL-ESM4	Riverside, USA	6	157
MPI-ESM-1-2-HAM	Riverside, USA	105	124
UKESM-1-0-LL	Riverside, USA	0	151

Table S3. Global climate model and ensemble members.

Global climate model	Scenario	Ensemble member	Citation
CESM2	Historical	ic1 – 001, ic1 – 002, ic1 – 003, ic1 – 004, ic2 – 001, ic2 – 002, ic2 – 003, ic2 – 004, ic3 – 001, ic3 – 002, ic3 – 003, ic3 – 004, ic4 – 001, ic4 – 002, ic4 – 003, ic4 – 004	1
CESM2	SSP 3-7.0	001, 002, 003, 004	2
EC-Earth3-AerChem	Historical	r1i1p1f1, r4i1p1f1	3
EC-Earth3-AerChem	SSP 3-7.0	r1i1p1f1, r4i1p1f1	4
GFDL-ESM4	Historical	r1i1p1f1	5
GFDL-ESM4	SSP 1-2.6	r1i1p1f1	6
GFDL-ESM4	SSP 2-4.5	r2i1p1f1, r3i1p1f1	7
GFDL-ESM4	SSP 3-7.0	r1i1p1f1	8
MPI-ESM-1-2-HAM	Historical	r1i1p1f1, r2i1p1f1, r3i1p1f1	9
MPI-ESM-1-2-HAM	SSP 3-7.0	r1i1p1f1, r2i1p1f1, r3i1p1f1	10
UKESM-1-0-LL	Historical	r1i1p1f2	11
UKESM-1-0-LL	SSP 1-2.6	r1i1p1f2	12
UKESM-1-0-LL	SSP 2-4.5	r1i1p1f2	13
UKESM-1-0-LL	SSP 3-7.0	r1i1p1f2	14
UKESM-1-0-LL	SSP 5-8.5	r1i1p1f2	15

References

1. Fiore, A. M., Hancock, S., Lamarque, J. F., Correa, G., Chang, K. L., Ru, M., Cooper, O., Gaudel, A., Polvani, L., Sauvage, B., and Ziemke, J. (2022). Understanding recent tropospheric ozone trends in the context of large internal variability: A new perspective from chemistry-climate model ensembles. *Environ. Res.: Climate* 1, 025008.
2. Personal communication with Dr. Jean-Francois Lamarque.
3. EC-Earth-Consortium EC-Earth3-AerChem model output prepared for CMIP6 CMIP historical (2020). Version 20200624 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.4701>.
4. EC-Earth-Consortium EC-Earth3-AerChem model output prepared for CMIP6 ScenarioMIP ssp370 (2020). Version 20220630 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.4885>
5. Krasting, J.P., John, J.G., Blanton, C., McHugh, C., Nikonov, S., Radhakrishnan, A., Rand, K., Zadeh, N.T., Balaji, V., Durachta, J., et al. (2018). NOAA-GFDL GFDL-ESM4 model output prepared for CMIP6 CMIP historical. Version 20190726 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.8597>.
6. John, J.G., Blanton, C., McHugh, C., Radhakrishnan, A., Rand, K., Vahlenkamp, H., Wilson, C., Zadeh, N.T., Dunne, J.P., et al. (2018). NOAA-GFDL GFDL-ESM4 model output prepared for CMIP6 ScenarioMIP ssp126. Version 20221210 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.8684>.
7. John, J.G., Blanton, C., McHugh, C., Radhakrishnan, A., Rand, K., Vahlenkamp, H., Wilson, C., Zadeh, N.T., Dunne, J.P., et al. (2018). NOAA-GFDL GFDL-ESM4 model output prepared for CMIP6 ScenarioMIP ssp245 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.8686>.
8. John, J.G., Blanton, C., McHugh, C., Radhakrishnan, A., Rand, K., Vahlenkamp, H., Wilson, C., Zadeh, N.T., Dunne, J.P., et al. (2018). NOAA-GFDL GFDL-ESM4 model output prepared for CMIP6 ScenarioMIP ssp370 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.8691>.
9. Neubauer, D., Ferrachat, S., Siegenthaler-Le Drian, C., Stoll, J., Folini, D.S., Tegen, I., Wieners, K.-H.; Mauritsen, T., Stemmler, I., Barthel, S., et al. (2019). HAMMOZ-Consortium MPI-ESM1.2-HAM model output prepared for CMIP6 CMIP historical (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.5016>.
10. Keeble, J., Hassler, B., Banerjee, A., Checa-Garcia, R., Chiodo, G., Davis, S., Eyring, V., Griffiths, P.T., Morgenstern, O., Nowack, P., et al. (2021). Evaluating stratospheric ozone and water vapour changes in CMIP6 models from 1850 to 2100. *Atmospheric Chem. Phys.* 21, 5015-5061. <https://doi.org/10.5194/acp-21-5015-2021>.
11. Tang, Y., Rumbold, S., Ellis, R., Kelley, D., Mulcahy, J., Sellar, A., Walton, J., and Jones, C. (2019). MOHC UKESM1.0-LL model output prepared for CMIP6 CMIP historical (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.6113>.
12. Good, P., Sellar, A., Tang, Y., Rumbold, S., Ellis, R., Kelley, D., and Kuhlbrodt, T. (2019). MOHC UKESM1.0-LL model output prepared for CMIP6 ScenarioMIP ssp126 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.6339>.
13. Good, P., Sellar, A., Tang, Y., Rumbold, S., Ellis, R., Kelley, D., and Kuhlbrodt, T. (2019). MOHC UKESM1.0-LL model output prepared for CMIP6 ScenarioMIP ssp245 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.6339>.
14. Good, P., Sellar, A., Tang, Y., Rumbold, S., Ellis, R., Kelley, D., and Kuhlbrodt, T. (2019). MOHC UKESM1.0-LL model output prepared for CMIP6 ScenarioMIP ssp370 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.6347>.
15. Good, P., Sellar, A., Tang, Y., Rumbold, S., Ellis, R., Kelley, D., and Kuhlbrodt, T. (2019). MOHC UKESM1.0-LL model output prepared for CMIP6 ScenarioMIP ssp585 (Earth System Grid Federation). <https://doi.org/10.22033/ESGF/CMIP6.6339>.