

# 1 Long-term trends in urban NO<sub>2</sub> concentrations and associated pediatric 2 asthma cases: estimates from global datasets

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## 17 Abstract

18 **Background:** Levels of nitrogen dioxide (NO<sub>2</sub>), a combustion-related air pollutant largely associated with  
19 traffic in urban areas, have been changing rapidly due to competing influences of regulation and  
20 population and fossil fuel-powered economic expansion. Traffic-related NO<sub>2</sub> is associated with pediatric  
21 asthma incidence in epidemiological studies around the world. We aim to assess long-term trends in  
22 NO<sub>2</sub> concentrations and NO<sub>2</sub>-attributable pediatric asthma incidence in cities globally.

23 **Methods:** We estimate global annual average surface NO<sub>2</sub> concentrations at 1km resolution for 1990-  
24 2019 by combining land use regression model predictions with NO<sub>2</sub> column densities from the Ozone  
25 Monitoring Instrument satellite sensor. We use these concentrations with an epidemiologically-derived  
26 concentration-response factor, population, and baseline disease rates to estimate NO<sub>2</sub>-attributable  
27 pediatric asthma incidence. We explore trends over the last two decades.

28 **Findings:** We found diverging regional trends leading to an emerging global convergence in urban NO<sub>2</sub>  
29 concentrations globally from 2000-2019. Concentrations are high but declining in high-income countries  
30 and low but rising elsewhere. Estimated NO<sub>2</sub>-attributable pediatric asthma incidence shows similar  
31 trends with decreases of 28-56% in North America, Western and Central Europe, and Australasia, but  
32 increases of >50% in Central and South Asia and >100% in Sub-Saharan Africa.

33 **Interpretation:** Traffic-related air pollution continues to be an important contributor to pediatric asthma  
34 incidence in cities in both developed and developing countries. Divergent experiences of different world  
35 regions show that while population growth is worsening NO<sub>2</sub> levels with substantial implications for

36 children's health in Asia and Africa, rapid and substantial NO<sub>2</sub> declines are possible with effective  
37 regulations.

38 **Funding:** Health Effects Institute and NASA

39

40 **Research in context:**

41 **Evidence before this study:** We searched PubMed and Google Scholar databases for studies published  
42 in English from the database inception until March 11, 2021, using the search terms (“NO<sub>2</sub>” OR “nitrogen  
43 dioxide”) AND “asthma” AND “trends”. Previous studies have reported epidemiological analyses linking  
44 changes with asthma to changes in NO<sub>2</sub>, or have assessed long-term trends in NO<sub>2</sub> concentrations in  
45 some countries or world regions. However, these studies provide little information about how NO<sub>2</sub>  
46 concentrations are changing in urban areas all around the world, and the influence those changes have  
47 on pediatric asthma incidence. One study published in 2019 showed that over 4 million new pediatric  
48 asthma cases, representing ~13% of all pediatric asthma cases worldwide in 2015, could be attributed to  
49 NO<sub>2</sub> pollution. Understanding temporal trends in NO<sub>2</sub>-attributable pediatric asthma incidence could help  
50 inform asthma and air pollution mitigation strategies.

51 **Added value of this study:** To our knowledge, this is the first study to have estimated long-term trends  
52 in NO<sub>2</sub> concentrations and NO<sub>2</sub>-attributable pediatric asthma incidence in urban areas worldwide. We  
53 focus on the last two decades, from 2000 to 2019, a period in which NO<sub>2</sub> levels have been changing  
54 rapidly around the world due to competing influences of regulation versus population growth and fossil  
55 fuel-powered economic expansion. We show that regional trends are diverging, with high but declining  
56 concentrations in high-income countries that have regulated NO<sub>2</sub> and low but rising concentrations in  
57 other parts of the world where population is expanding. The effect of these contrasting trends is an  
58 emerging convergence in urban NO<sub>2</sub> concentrations, and, to a lesser extent, NO<sub>2</sub>-attributable asthma  
59 incidence, globally.

60 **Implications of all the available evidence:** Urban NO<sub>2</sub> concentrations have been trending downward for  
61 decades in places that have effective air quality regulations, with benefits for improved children's  
62 respiratory health. Even with these improvements, the current levels of NO<sub>2</sub> contribute substantially to  
63 pediatric asthma incidence, highlighting that mitigating air pollution should be a critical element of  
64 children's public health strategies. For cities that have not benefited from strong local or national air  
65 quality management programs, the experience of cities that have such programs demonstrates that  
66 addressing combustion-related air pollution can lead to dramatic improvements in air quality and public  
67 health over short time frames. Our study also demonstrates the utility of satellite remote sensing for  
68 environmental and public health surveillance in urban areas worldwide.

69 **Acknowledgements:** This work was supported by grants from the Health Effects Institute (Research  
70 Agreement #4977/20-11) and NASA (Grant #80NSSC19K0193). We gratefully acknowledge the  
71 developers of the OMI NO<sub>2</sub> concentration products, GHS-SMOD urban area dataset, GBD disease rate  
72 datasets, and Worldpop population dataset. We appreciate helpful discussions with Bryan Duncan.  
73 Research described in this article was conducted under contract to the Health Effects Institute (HEI), an  
74 organization jointly funded by the United States Environmental Protection Agency (EPA) (Assistance  
75 Award No. R-82811201) and certain motor vehicle and engine manufacturers. The contents of this

76 article do not necessarily reflect the views of HEI, or its sponsors, nor do they necessarily reflect the  
77 views and policies of the EPA or motor vehicle and engine manufacturers.

78 **Data availability:** NO<sub>2</sub> concentrations are available at:

79 [https://figshare.com/articles/dataset/Global\\_surface\\_no2\\_concentrations\\_1990-2020/12968114](https://figshare.com/articles/dataset/Global_surface_no2_concentrations_1990-2020/12968114).

80 Estimated NO<sub>2</sub>-attributable asthma incidence results are available at:

81 [https://github.com/AMohegh/Asthma\\_no2\\_urban](https://github.com/AMohegh/Asthma_no2_urban)

82 **Author contributions:** SCA, AM, and DG designed the study; PH, AL, and LL provided data; AM carried  
83 out the calculations, SCA, AM, DG, MB, PH, AL, and SW contributed input on methods development; SCA  
84 and AM wrote the paper; all authors helped interpret results and reviewed the manuscript. AM and DG  
85 have verified the underlying data.

86 **Declaration of interests:** The authors declare no conflict of interest.

87

## 88 Introduction

89 Nitrogen dioxide (NO<sub>2</sub>), a component of nitrogen oxides (NO<sub>x</sub>), is a pervasive air pollutant that is  
90 regulated by the U.S. Environmental Protection Agency. NO<sub>2</sub> is a precursor for ground-level ozone and  
91 fine particulate matter (PM<sub>2.5</sub>), other regulated air pollutants that are the leading contributors to air  
92 pollution-related mortality.<sup>1</sup> NO<sub>2</sub> is an effective tracer for anthropogenic fuel combustion generally, and  
93 traffic specifically.<sup>2-5</sup> Major anthropogenic NO<sub>2</sub> sources include on-road and non-road transportation  
94 tailpipe emissions (including heavy-, medium-, and light-duty vehicles, shipping, and aviation), power  
95 plants, industrial manufacturing, and agriculture.<sup>6-9</sup> In urban and industrialized areas, a lower fraction of  
96 NO<sub>2</sub> comes from natural sources like soil and lightning. NO<sub>2</sub> concentration trends can be used to  
97 evaluate the efficacy of air pollution regulations, as well as effects of abrupt emission changes (e.g.  
98 power plant closures, new oil and gas fields, COVID-19 lockdowns).<sup>10-13</sup>

99 In addition to its role in PM<sub>2.5</sub> and ozone formation, NO<sub>2</sub> itself (often as a marker for the broader traffic-  
100 related air pollution mixture) has been linked to several adverse health outcomes including asthma  
101 exacerbation.<sup>14,15</sup> Recent epidemiological studies have also found associations between transportation  
102 related air pollutants with new asthma development in children.<sup>16,17</sup> While the putative agent in the  
103 traffic-related air pollution mixture remains unknown, epidemiological studies are relatively consistent  
104 in their finding that NO<sub>2</sub> is significantly associated with pediatric asthma incidence, while the evidence  
105 for other traffic-related air pollutants (e.g. PM<sub>2.5</sub>) is more mixed.<sup>16,17</sup> Previous health impact assessments  
106 have linked NO<sub>2</sub> with over four million new pediatric asthma cases each year globally, ~13% of the global  
107 pediatric asthma burden.<sup>18,19</sup> The fraction of new pediatric asthma cases that were attributed to NO<sub>2</sub>  
108 ranged substantially higher, up to ~50%, in urban areas.

109 NO<sub>x</sub> emissions and NO<sub>2</sub> concentrations have changed dramatically in response to socioeconomic  
110 changes.<sup>20-24</sup> In the U.S., NO<sub>2</sub> concentrations have dropped ~50% on average from the 1980s to 2010s,<sup>25</sup>  
111 with even larger drops near major roadways<sup>25</sup> and point sources.<sup>10</sup> In the last two decades, U.S. NO<sub>x</sub>  
112 emissions have been falling ~5% per year as vehicles get more fuel efficient and cleaner and as power  
113 plants have been shifting from coal to relatively cleaner fuels (e.g. natural gas).<sup>11,26,27</sup> NO<sub>2</sub> concentrations  
114 have also decreased in Europe, though more slowly.<sup>28,29</sup> In contrast, NO<sub>2</sub> has been increasing in India,<sup>30</sup>

115 the Middle East,<sup>31</sup> and eastern Europe.<sup>22</sup> In China, NO<sub>x</sub> emissions peaked around 2011/2012 and  
116 subsequently declined.<sup>32–34</sup>

117 Previous research on NO<sub>2</sub> temporal trends in urban areas have focused only on a small subset of cities  
118 and have not considered its health impacts, precluding the ability to compare urban trends in NO<sub>2</sub>  
119 concentrations and associated health burdens in a consistent manner across the world. The global  
120 coverage and long continuous record of satellite remote sensing since the 1990s make it possible to  
121 track urban NO<sub>2</sub> concentrations globally.<sup>35–38</sup> Additionally, current satellite instruments operate with  
122 substantially higher spatial resolution compared with their predecessors, enabling assessments of intra-  
123 urban NO<sub>2</sub> variation and identification of small NO<sub>x</sub> emissions sources.<sup>39,40</sup>

124 Here we investigate long-term trends of annual average NO<sub>2</sub> concentrations and associated pediatric  
125 asthma burdens in urban areas over the past two decades (2000-2019) globally. We first generate a new  
126 gridded global surface annual average NO<sub>2</sub> concentration dataset from 1990 to 2019 using satellite data  
127 from the Ozone Monitoring Instrument (OMI) to scale NO<sub>2</sub> concentrations estimated from a previously  
128 published land use regression (LUR) model<sup>41</sup> to different years. We then use epidemiological  
129 concentration-response relationships to estimate the temporal trend in urban NO<sub>2</sub>-attributable pediatric  
130 asthma cases globally over the last two decades (2000-2019). Finally, we deconstruct the drivers of  
131 these trends to explore the influence of NO<sub>2</sub> versus demographic changes.

132

## 133 **Methods**

134 We integrated global environmental and demographic datasets available from different sources to  
135 generate new estimates of surface NO<sub>2</sub> concentrations and NO<sub>2</sub>-attributable pediatric asthma incidence  
136 in cities globally. The analysis was done in Python (version 3.6.7).

### 137 *NO<sub>2</sub> concentrations*

138 We estimated surface annual average NO<sub>2</sub> concentrations at 0.0083 degree resolution (~1 km<sup>2</sup>) in five-  
139 year increments from 1990-2010 and annually from 2010-2019. To begin, we used a previously  
140 published NO<sub>2</sub> concentration dataset for the average of 2010-2012, which used land use regression  
141 modeling with inputs from road networks and other land use variables, as well as satellite NO<sub>2</sub> column  
142 observations from SCIAMACHY and GOME-2.<sup>4,41</sup> We aggregated NO<sub>2</sub> concentrations from this dataset  
143 from its native 100m x 100m resolution globally to 0.0083 degree resolution, which is still a high enough  
144 resolution to avoid substantially underestimating NO<sub>2</sub>-attributable asthma impacts.<sup>19</sup>

145 Due to the lack of ground measurements in rural areas, the Larkin et al.<sup>41</sup> NO<sub>2</sub> dataset is fine-tuned  
146 towards the urban areas and overestimates NO<sub>2</sub> concentrations in rural areas. We therefore applied the  
147 Larkin et al.<sup>41</sup> NO<sub>2</sub> concentrations in all 1km x 1km grid cells globally that are categorized as “urban”  
148 according to the Global Human Settlement Model grid,<sup>42</sup> as well as those grid cells that include major  
149 roadways<sup>41</sup>. For grid cells >5km away from roadways and in rural areas, we developed new NO<sub>2</sub>  
150 concentration estimates using NO<sub>2</sub> column observations from the OMI satellite instrument with some  
151 adjustments to fill spatial and temporal gaps in the OMI satellite record, and to estimate 24 hour  
152 averages from the early afternoon OMI overpass time. The datasets, correction factors, and evaluation  
153 against ground monitor observations are described in the Supplemental Material and in Table S1 and  
154 Figures S1 through S5.

155 While the global NO<sub>2</sub> concentration dataset is available for 1990-2019, the estimates for 1990 and 1995  
156 have more uncertainty. Here, we use the results for 2000 to 2019 to explore trends and NO<sub>2</sub>-attributable  
157 pediatric asthma incidence, as the long-term satellite observational record enhances confidence in  
158 results for this period.

159

### 160 *NO<sub>2</sub>-attributable pediatric asthma incidence*

161 We estimated NO<sub>2</sub>-attributable cases of pediatric asthma incidence for 2000-2019 using an  
162 epidemiologically-derived concentration response function, following the method used by Achakulwisut  
163 et al.<sup>18</sup> and Mohegh et al.<sup>19</sup>, as shown in Equation 1:

$$\text{Equation 1: } Burden_{C, AG} = \sum_{\text{Grid cells in } C} Inc_{C, AG} \times Pop_{\text{Grid cell } (i,j), AG} \times (1 - e^{-\beta_{AG} X_{\text{Grid cell } (i,j)}})$$

164

165 where C is the country, AG is the age group, Inc is the baseline asthma incidence rate for the age group  
166 and country, Pop is the grid-cell population,  $\beta$  is the concentration response factor relating the NO<sub>2</sub>  
167 concentration with increased risk of asthma incidence, and X is the grid-cell annual average NO<sub>2</sub>  
168 concentration. We perform the health impact calculations at a spatial resolution close to the resolution  
169 of the population dataset (1km), since previous work indicates that 1km resolution achieves  
170 computational efficiency while still capturing co-location of concentrations and population at the city-  
171 scale. Therefore, all other datasets have been re-gridded to match it, including the concentration  
172 dataset at ~100m resolution and asthma baseline rates at the country/territory level). We assumed a  
173 theoretical minimum risk exposure level (TMREL) of 2 ppb annual average NO<sub>2</sub> concentration, and only  
174 estimated NO<sub>2</sub>-attributable asthma incidence above that level. We applied a relative risk (RR) of 1.26 per  
175 10ppb increase in annual average NO<sub>2</sub> concentration from Khreis et al.<sup>16</sup>

176 We use population estimates from Worldpop (Tatem 2017) in four different age groups (1-4, 5-9, 10-14,  
177 and 15-18 years old) from 2000-2019 at ~1km resolution. National baseline asthma incidence rates are  
178 from the Institute for Health Metrics and Evaluation's Global Burden of Disease (GBD) 2019 Study for  
179 the same age groups. Boundaries of urban areas are from the GHS-SMOD Urban Centre dataset which  
180 provides shapefiles of boundaries of over 13,000 urban clusters globally for the year 2015 (latest year).<sup>42</sup>  
181 We use the 2015 urban boundaries for all years of our analysis. Here, we consider grid-cells to be part of  
182 an "urban cluster" if they are included in "urban" and "sub-urban" areas in the GHS-SMOD dataset,  
183 which narrows the selection to areas with more than 300 people per km<sup>2</sup> that are part of clusters with  
184 more than 5000 population. We consider all other gridcells to be "rural". Regional definitions are  
185 consistent with the Global Burden of Disease 2019 Study.<sup>1</sup>

186

### 187 *Drivers of change*

188 To disentangle the contribution of exposure, population, and disease rates to NO<sub>2</sub>-attributable pediatric  
189 asthma cases, we performed four sets of simulations: a control scenario, where we calculate the  
190 estimated asthma cases for each year, and three sets of scenarios in which we revert one contributing  
191 parameter back to the base year of 2000. Using a combination of these simulations, we estimate the  
192 contribution of each parameter in the overall changes in estimated asthma cases between 2000 and

193 each year of analysis through 2019. The contribution of each parameter to that change in NO<sub>2</sub>-  
194 attributable asthma cases between years is calculated using the formulas presented in the supplemental  
195 document. This method allows for calculating ratios for each parameter that add up to the scaling factor  
196 between the year 2019 and 2000, and due to its logarithmic nature, it is consistent with the original  
197 health impact function, which is a multiplication of three parameters. This approach is described in more  
198 detail in the Supplemental Material.

199

#### 200 *Role of the funding source*

201 The funders of the study had no role in study design, data collection, data analysis, data interpretation,  
202 or writing of the report. SA and AM had full access to all the data in the study and had final responsibility  
203 for the decision to submit for publication.

204

#### 205 **Results**

206 When averaged globally, urban annual average NO<sub>2</sub> concentrations decreased by 13% between 2000  
207 and 2019 (Figure 1). We estimate that in 2000, the global urban population weighted average  
208 concentration was 12.2 ppb, it increased between 2006 and 2011, and then slowly declined from 2012  
209 to a level of 10.6 ppb in 2019. However, these global trends mask steeper, and contrasting trends in  
210 different world regions. NO<sub>2</sub> concentrations in high-income cities were the highest among all regions in  
211 2000, and remained above the global average throughout the time period, even after the steady, nearly  
212 monotonic decline of 35% in these cities from 2000 to 2019. In contrast, concentrations have risen in  
213 South Asia by 16% and in Sub-Saharan Africa by 11% over this period, though concentrations in these  
214 regions remain the lowest of cities in all other world regions in 2019. Cities in other world regions  
215 experienced only modest changes in NO<sub>2</sub> concentrations from 2000 to 2019. Even these large regional  
216 groupings of cities obscure contrasting temporal trends between sub-regions. For example, within the  
217 high-income region, concentrations dropped by nearly 53% in North America, but only by 35% and 39%  
218 in Western Europe and Australasia, respectively. In high-income Asia Pacific cities, concentrations rose  
219 dramatically in the 2000s and then declined steadily from 2005 to 2019, and are now at approximately  
220 the same level as in 2000. Another region with contrasting sub-regional trends is Central and Eastern  
221 Europe and Central Asia, where concentrations have dropped by a large amount in Central and Eastern  
222 Europe but have risen steadily in Central Asia. Similar trends can be seen when including all urban and  
223 rural areas together (Figure S7).

224 We estimate that 1.85 million new pediatric asthma cases were attributed to NO<sub>2</sub> pollution globally in  
225 2019 (in both urban and rural areas, Figure S8), approximately 65% of which occurred in urban areas  
226 (Figure S9). Similar to concentrations, estimated NO<sub>2</sub>-attributable pediatric asthma cases changed very  
227 little from 2000 to 2019 (Figure 2). Regional trends in estimated NO<sub>2</sub>-attributable asthma incidence  
228 largely follow the NO<sub>2</sub> concentration trends, though with less cross-over in rankings between the world  
229 regions over time. For example, the high-income region had the most NO<sub>2</sub>-attributable asthma incidence  
230 in 2000, and it still had the most in 2019, despite declining trends. Similarly, Central and Eastern Europe  
231 and Central Asia, South Asia, and Sub-Saharan Africa had the least NO<sub>2</sub>-attributable asthma cases  
232 throughout 2000-2019, despite increasing trends over time. Sub-regional trends in NO<sub>2</sub>-attributable

233 asthma incidence show a similar pattern as sub-regional trends in NO<sub>2</sub> concentrations. Considering  
234 individual countries, the 2019 to 2000 ratio of urban NO<sub>2</sub>-attributable asthma cases is relatively  
235 consistently positive throughout India, generally negative with increases in some areas within the United  
236 States and Western European countries, and very mixed within China, South American countries, and  
237 South Africa (Figure 3). While approximately two-thirds of NO<sub>2</sub>-attributable asthma cases occurred in  
238 urban areas, there is wide variation across regions driven largely by differences in urban vs rural  
239 population in these regions (Figure S4).

240 The estimated trends in urban NO<sub>2</sub>-attributable pediatric asthma incidence are driven by simultaneous  
241 and often competing changes in NO<sub>2</sub> concentrations, pediatric population size, and asthma incidence  
242 rates over time (Figure 4). We find that these drivers of the NO<sub>2</sub>-attributable asthma trends are  
243 inconsistent globally, though some world regions show similar influences. No region that has  
244 experienced a decline in estimated NO<sub>2</sub>-attributable asthma cases has done so without a drop in NO<sub>2</sub>  
245 concentrations. Similarly, all regions that experienced an increase in NO<sub>2</sub> concentrations saw an increase  
246 in estimated NO<sub>2</sub>-attributable asthma cases. Australasia and high-income Asia Pacific are the only  
247 regions where concentrations, population, and disease rates all contribute to the overall decline in NO<sub>2</sub>-  
248 attributable asthma cases. The opposite occurred in Central, South, and Southeast Asia, as well as in  
249 North Africa and the Middle East, where concentrations, population, and disease rates all contributed to  
250 an increasing trend in NO<sub>2</sub>-attributable asthma cases. High-income North America, Western Europe, and  
251 several other regions experienced competing effects of reduced NO<sub>2</sub> concentrations but increases in  
252 asthma incidence rates and/or pediatric population size. These competing influences changed over time,  
253 such that declining concentrations have a larger influence over time in North America and Southern  
254 Latin America, while population growth becomes more of a factor in North Africa and Middle East and  
255 Southern Sub-Saharan Africa (Figure S10). Changing baseline pediatric asthma rates also influence the  
256 trends in some regions, in particular East Asia and Southern Sub-Saharan Africa.

257

## 258 **Discussion**

259 We found an emerging convergence in NO<sub>2</sub> concentrations globally, with high but declining  
260 concentrations in high-income countries that have regulated NO<sub>2</sub> and low but rising concentrations in  
261 other parts of the world where population is expanding. Estimated urban NO<sub>2</sub>-attributable pediatric  
262 asthma incidence also appears to be beginning to converge globally, though with wider separation  
263 between world regions compared with regional average concentrations, driven by differences in  
264 population and asthma rates between countries. In no region did NO<sub>2</sub>-attributable asthma cases decline  
265 without declining NO<sub>2</sub> concentrations, while population growth almost always contributed to rising  
266 trends in NO<sub>2</sub> asthma impacts. The influence of baseline pediatric asthma incidence rates was  
267 inconsistent between regions.

268 Our estimated NO<sub>2</sub> concentration trends are consistent with recent studies that used satellite data to  
269 investigate global NO<sub>2</sub> concentration trends during 2004-2018 for the U.S., Europe, China, India, and  
270 Japan<sup>21,22,43</sup>. Qu et al.<sup>44</sup> also showed a similar decrease in U.S. NO<sub>2</sub> concentrations based on ground  
271 observations, satellite data, and modeling outputs from 2006-2016, and Henneman et al.<sup>25</sup> showed a  
272 similar decrease using ground observations from 1980-2020.

273 Our estimate of 1.84 million NO<sub>2</sub>-attributable new pediatric asthma cases globally (in both urban and  
274 rural areas) in 2019 is less than half of the 4.2 million found by Achakulwisut et al.<sup>18</sup> for 2015. Our  
275 estimates for India and the U.S. are 35% lower and 53% higher than our previous estimates,  
276 respectively<sup>19</sup>. Several factors explain this discrepancy. First, our new NO<sub>2</sub> concentration dataset  
277 corrects for a high NO<sub>2</sub> concentration bias in rural areas. This change results in lower contributions of  
278 NO<sub>2</sub> to asthma incidence in rural areas, particularly in countries with larger rural populations like India.  
279 Second, compared with previous GBD versions, GBD 2019 baseline asthma rates are much lower, except  
280 for high income areas such as the U.S. For example, the global baseline asthma incidence rate for people  
281 <20 years old in 2015 (the year analyzed by Mohegheh et al.<sup>19</sup>) decreased from 1055 to 828 cases per  
282 100,000 in the GBD 2017 (GBD 2017) vs. GBD 2019 Study (GBD 2019). This difference is even larger in  
283 China and India, where pediatric asthma incidence rates in the GBD 2019 Study were ~72% of their value  
284 in the GBD 2017 Study. U.S. baseline pediatric asthma incidence rates increased from 1257 to 2750  
285 cases per 100,000 in the GBD 2017 vs. GBD 2019 Study. Changes in baseline asthma rates have  
286 approximately proportional effects on estimated NO<sub>2</sub>-attributable pediatric asthma cases. Finally, as we  
287 have shown, NO<sub>2</sub> concentrations are changing rapidly in different directions depending on the region  
288 and city, with the same directional effect on estimated NO<sub>2</sub>-attributable pediatric asthma cases.

289 While we sought to create a new NO<sub>2</sub> dataset that leverages the advantages of different data sources,  
290 our new NO<sub>2</sub> concentrations are still uncertain. First, many cities are still lacking ground NO<sub>2</sub> monitors,  
291 limiting our ability to calibrate and evaluate concentrations in these regions.<sup>41</sup> Urban NO<sub>2</sub> concentrations  
292 therefore have more certainty in North America, Europe, and Asia, compared with Africa and South  
293 America. Second, rural NO<sub>2</sub> concentrations are uncertain in all regions globally due to the  
294 disproportionate location of ground monitors in urban areas. Third, our approach for scaling NO<sub>2</sub>  
295 concentrations from the 2010-2012 average LUR model to other years assumes that the land use  
296 predictors of that regression model are unchanged over time. This assumption is likely supported by  
297 slow changes in road density and volume and urban form from one year to the next, but over the two  
298 decades time frame explored here, some land use evolution is likely. The directional impact of these  
299 uncertainties and limitations on results is currently unknown.

300 Estimated NO<sub>2</sub>-attributable asthma incidence is heavily influenced by baseline disease rates, which vary  
301 considerably within individual countries.<sup>45</sup> While we used national pediatric asthma rates, asthma  
302 prevalence differs between urban and rural areas. Living in urban areas has been associated with  
303 increased risk of asthma prevalence in low- and middle-income countries<sup>46</sup> and asthma-related  
304 emergency department visits and hospitalizations in the U.S.<sup>47</sup> Our method also does not capture  
305 temporal trends in asthma rates in cities specifically, which may differ from the broader national rate  
306 changes. If asthma prevalence tends to be higher in urban areas compared with national averages, we  
307 may have underestimated NO<sub>2</sub>-attributable asthma cases.

308 Despite these uncertainties and limitations, our results demonstrate the important influence of  
309 combustion-related air pollution on children's health in cities globally. In places that have effective air  
310 quality management programs, NO<sub>2</sub> concentrations have been trending downward for decades, with  
311 benefits for improved children's respiratory health. Even with these improvements, the current levels of  
312 NO<sub>2</sub> contribute substantially to pediatric asthma incidence, highlighting that mitigating air pollution  
313 should be a critical element of children's public health strategies. For cities that have not benefited from  
314 strong local or national-scale air quality management programs, the experience of cities that have such  
315 programs demonstrates that addressing combustion-related air pollution can lead to dramatic

316 improvements in air quality and public health over short time frames. These air quality improvements  
317 can be achieved through either end-of-pipe emission control technologies or avoiding the combustion in  
318 the first place, which would have additional benefits from reduced greenhouse gas emissions.

319 Our study also highlights the value of satellite remote sensing and statistical models for tracking NO<sub>2</sub>  
320 pollution and for environmental health surveillance at local, national, and global scales. The combination  
321 of methods offers strengths beyond what is possible from each technique alone: a long and consistent  
322 observational record of NO<sub>2</sub> column densities from satellite remote sensing with the high spatial  
323 resolution of the surface concentration predictions from the LUR model. Future studies may seek to  
324 leverage these data sources and others, including mobile monitoring, distributed networks of ground  
325 sensors, and chemical transport models, to further improve upon the accuracy and spatiotemporal  
326 resolution of NO<sub>2</sub> concentration estimates. Further, our study shows the importance of considering not  
327 just concentrations, but how demographics change over time, for understanding air pollution health  
328 risks. Improved and more widely accessible information about disease rates, and capturing population  
329 distribution and movement, will enable more accurate and highly resolved air pollution health impact  
330 assessments.

331

332

333 **References**

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460

461

462 **Figure captions**

463

464 Figure 1. Trends in annual average NO<sub>2</sub> surface concentrations in urban areas in various world regions.  
465 The figure presents regional average concentrations aggregated for GBD super regions in form of NO<sub>2</sub>  
466 concentrations (main panel) and aggregated results for GBD regions in each super region in form of  
467 relative changes from the year 2000 (side panels). Regional definitions are shown in Figure S5. China is  
468 included in East Asia and India is included in South Asia. H.I. = High Income; L.A.C. = Latin America &  
469 Caribbean; S.S.A. = Sub-Saharan Africa; N.A.,M.E. = North Africa and Middle East; S.A. = South Asia;  
470 SE.A., E.A., O. = Southeast Asia, East Asia, and Oceania; C.E., E.E., C.A. = Central Europe, Eastern Europe,  
471 and Central Asia.

472

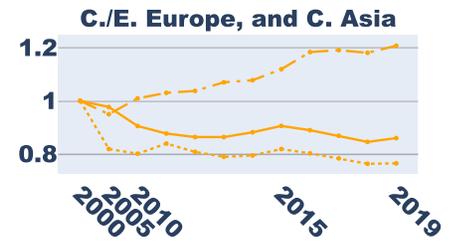
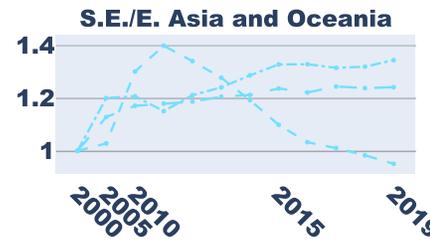
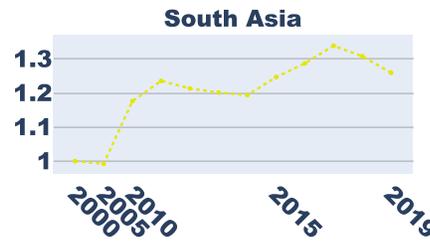
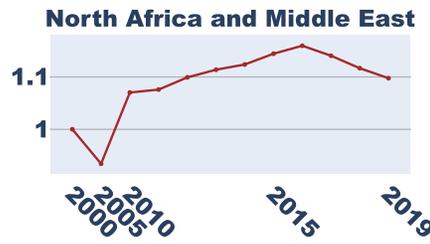
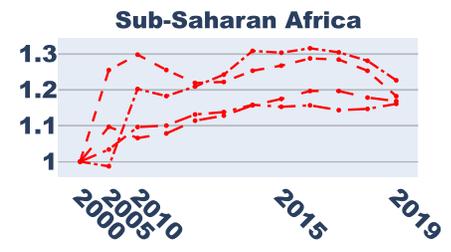
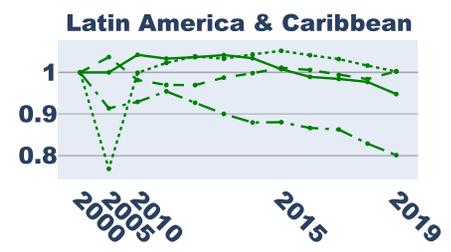
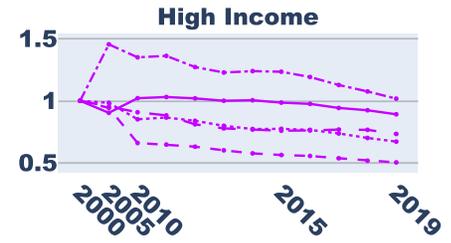
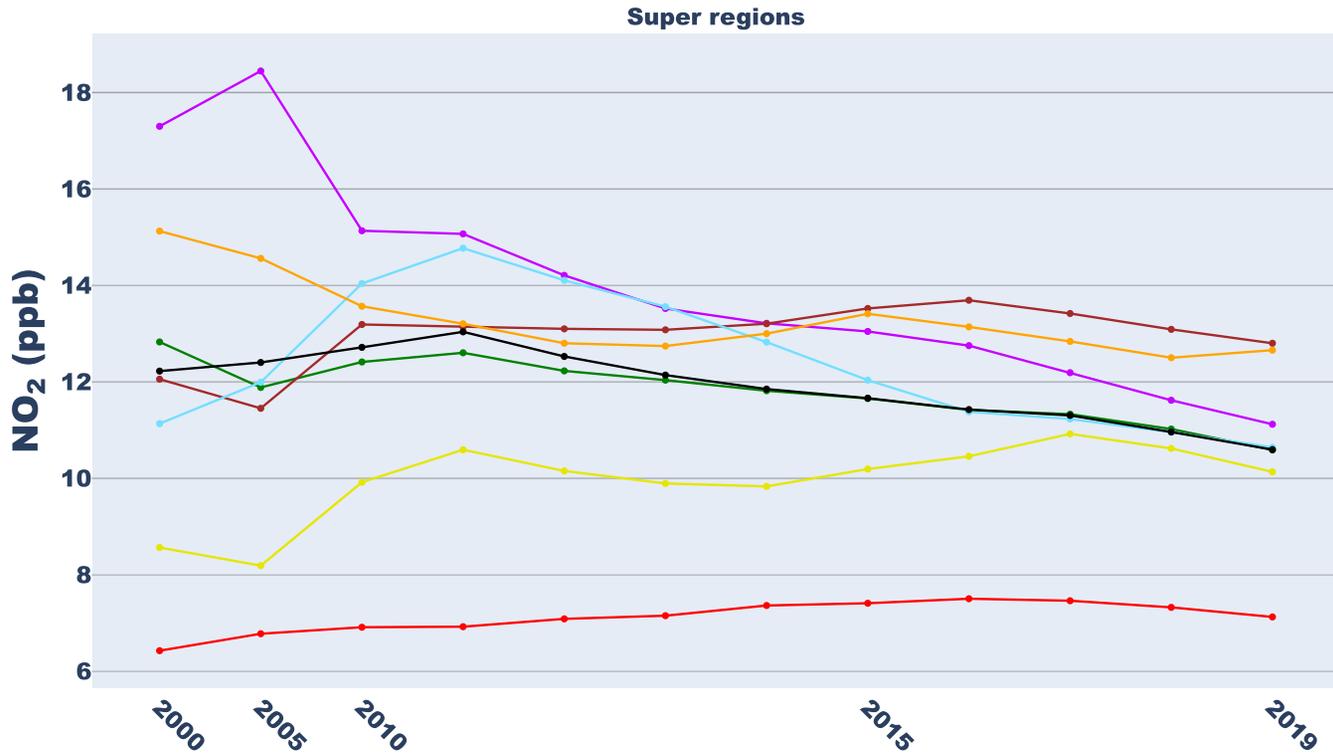
473 Figure 2. Trends in estimated urban NO<sub>2</sub>-attributable pediatric asthma incidence in each super-region  
474 (absolute magnitude, main panel) and sub-region (relative changes from 2000, side panels). The legends  
475 are the same as Figure 1.

476

477 Figure 3. Relative changes in estimated NO<sub>2</sub>-attributable pediatric asthma incidence between 2000 and  
478 2019 globally (including both urban and rural areas). Lack of color indicates no NO<sub>2</sub>-attributable  
479 pediatric asthma incidence as NO<sub>2</sub> concentrations were <2 ppb, the low concentration threshold for  
480 estimating attributable asthma incidence.

481

482 Figure 4. Contribution of population, baseline asthma rates, and NO<sub>2</sub> concentrations to changes in  
483 estimated NO<sub>2</sub>-attributable pediatric asthma incidence between the year 2000 and 2019 for each sub-  
484 region. Since meteorological factors can influence comparisons of concentration contributions when  
485 comparing two years, we show contributions by year for several regions in Figure S3. Percentage labels  
486 indicate the net change.



- Global    —●— H.I., Southern Latin America    -.-.- H.I., Western Europe    -.-.- H.I., High-income North America    —●— H.I., Australasia
- H.I., High-income Asia Pacific    -.-.- L.A.C, Central Latin America    —●— L.A.C, Tropical Latin America
- .-.- L.A.C, Andean Latin America    -.-.- L.A.C, Caribbean    -.-.- S.S.A, Southern Sub-Saharan Africa
- .-.- S.S.A, Western Sub-Saharan Africa    -.-.- S.S.A, Central Sub-Saharan Africa    -.-.- S.S.A, Eastern Sub-Saharan Africa
- N.A., M.E., North Africa and Middle East    -.-.- S.A., South Asia    -.-.- SE.A., E.A., O., East Asia    —●— SE.A., E.A., O., Southeast Asia
- .-.- SE.A., E.A., O., Oceania    —●— C.E., E.E., C.A., Central Asia    —●— C.E., E.E., C.A., Eastern Europe    -.-.- C.E., E.E., C.A., Central Europe

