Health Impact Assessment of Fine Particle Pollution at the Regional Level

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Since the year 2000, evaluation of the impact of air pollution on people’s health has drawn the attention of the general public and has led decision-makers to develop specific health policies. In most of the health impact assessment literature, investigators have reported on long- and short-term effects of air pollution. Here the authors present results of a health impact assessment of short-term effects of particulate matter ≤10 μm in diameter (PM10) in the Lombardy region of Italy (2003–2006). The impact was evaluated in terms of numbers of attributable deaths under several counterfactual scenarios of air pollution reduction based on World Health Organization guidelines and European Union limits. The authors found that annual average PM10 levels exceeding the World Health Organization threshold of 20 μg/m³ and the European Union limit of 40 μg/m³ were responsible for 302 and 109 attributable deaths per year, corresponding to attributable community rates of 13 and 5 deaths per 100,000 inhabitants per year, respectively. A 20% reduction in existing PM10 levels could reduce by more than 30% the burden of short-term deaths linked to ambient air pollution. Therefore, policies for air pollution reduction appear to be necessary in order to protect and improve individual and community health.

air pollution; attributable risk; Bayesian analysis; particulate matter; public health; shrunken estimator

Abbreviations: CrI, credibility interval; EU, European Union; MISA, Meta-Analysis of the Italian Studies on Short-Term Effects of Air Pollution; PM10, particulate matter ≤10 μm in diameter; RS, reduction scenario; WHO, World Health Organization.
The region can be geographically and economically divided into 3 zones: the mountain range of the Alps; the sloping foothills; and the immediate facing plains, where one finds the highly industrialized provinces of Varese, Como, Lecco, Bergamo, and Brescia. The service sector is concentrated in the Milan area, and the large Po River plains have a rich agricultural sector (16). The population is concentrated in the Milan conurbation and the Alpine foothills of the above-mentioned provinces (6.5 million inhabitants, a density of 1,200 inhabitants/km²). Since the 1980s, while remaining the most important industrial area of the country, Lombardy has shown marked growth in the service sector.

The area’s climate depends on altitude and the presence of inland waters. The temperature shows high annual variations (in Milan, the average temperature is 1.5°C in January and 24°C in July), and thick fog is frequent between October and February. The basin of the Po River, where most of the major cities are located, is bordered on 3 sides by mountains, and Atlantic weather disturbances are frequently unable to cross the Alpine barrier, consequently providing no air mass exchange. Wind speed measured in the Po River basin is among the lowest in Europe. This causes frequent phenomena of thermal inversion, with smog and pollution being trapped close to the ground. This unfavorable context and these climate characteristics create a high level of air pollution. Lombardy has exceeded the EU PM₁₀ limit since 1996, when the limit was first issued. Road transport can be considered the main source of air pollution in the urban areas considered in this study. For example, in the district of Milan, road transport is estimated to be responsible for 63% of total PM₁₀ emissions (17).

In the present study, we assessed the health impact of PM₁₀ in Lombardy during the period 2003–2006. We evaluated the impact of PM₁₀ in terms of number of attributable deaths, assuming several counterfactual scenarios of air pollution reduction. We focused on short-term evaluation because it allows for immediate appraisal of deaths prevented by reduction policies, while for long-term effects the impact is highly speculative in nature because of latency time and the role of cumulative exposure.

We propose an original approach to assess the health impact of air pollution at the regional level that is characterized by the use of local data for estimation of the concentration-response function and by the specification of several counterfactual scenarios defined in terms of both annual average PM₁₀ concentrations and daily concentrations. In particular, we applied 2 different methods for calculation of attributable deaths. The first, which we named the “macro” approach, is appropriate for evaluating impacts under scenarios defined in terms of annual mean air pollutant concentration, and the second, named the “micro” approach, is used when the counterfactual scenarios are defined in terms of daily levels of PM₁₀.
MATERIALS AND METHODS

Data

We considered air pollution and mortality data for the period covering 2003–2006 for 13 areas in Lombardy: 11 cities with more than 50,000 inhabitants, 1 smaller town (Sondrio) that is the capital of an Alpine administrative province, and all of the municipalities belonging to the administrative agricultural district of Lodi, collapsed into a single epidemiologic time series. We did not consider the smaller municipalities in estimating air pollutant effects because precision is poor in the presence of small daily death counts.

The air quality monitoring network of the Lombardy Regional Environmental Protection Agency provided the daily time series of PM$_{10}$ measurements, temperature, and relative humidity values. All of the monitoring stations were located at sites not greatly influenced by local traffic and therefore provided measurements of the background levels of PM$_{10}$. For each municipality, missing daily values at one monitor were imputed using concentrations measured by the remaining monitors, and a daily time series of PM$_{10}$ levels was obtained by averaging data over the available monitors (10).

Death certificates were obtained from the regional mortality register. We considered mortality from all causes, excluding external causes (International Classification of Diseases, Ninth Revision, codes below 800). For each area, we focused on daily numbers of deaths occurring in the resident population inside the area and in municipalities within a 10-km radius from the border.

Effect estimates

Health impact was evaluated for each area in terms of the number of attributable deaths, combining the concentration-response functions derived from a Bayesian meta-analysis of the 13 study areas and the observed daily time series of PM$_{10}$ concentrations and mortality over the study period. Inference on the concentration-response function was made up of 2 steps: an area-specific analysis and a Bayesian combined analysis. At the first stage, we estimated the short-term effect of PM$_{10}$ separately for each area. For specificity reasons, in estimating the air pollutant effect we considered only deaths of the resident population occurring inside the area. By excluding deaths occurring outside the area, we obtained less precise estimates, but we avoided possible bias due to classifying as exposed those persons who did not experience the PM$_{10}$ levels observed in their residence area.

We specified a Poisson regression model for the daily number of deaths (10, 18). Analysis was age-adjusted (<65, 65–74 years, and ≥75 years). We controlled for time-related confounding, effects of temperature and humidity, and influenza epidemics. The average of current-day and previous-day concentrations (lag 0–1) was used as the indicator of PM$_{10}$ exposure. We modeled the air pollution effect using a linear term. Analyses were performed with R 2.11.0 software (19).

At the second stage of the analysis, a Bayesian random-effects meta-analysis of the first-stage area-specific estimates was performed (20). Random-effects meta-analysis allows one to obtain estimates of the second-stage area-specific effects ($\hat{\beta}_i$), or shrunken estimates. Under the classical approach, shrunken estimates ($\hat{\beta}_i$) combine the area-specific estimates obtained in the first step of the analysis ($\hat{\beta}_i$) and the overall estimate from meta-analysis ($\hat{\beta}$). Each location-specific estimate is pulled towards the overall effect estimate, proportionally to its variance $\hat{\sigma}_i^2$:

$$\hat{\beta}_i = \frac{\hat{\sigma}_i^2}{\hat{\sigma}_i^2 + \tau^2} \hat{\beta} + \left(1 - \frac{\hat{\sigma}_i^2}{\hat{\sigma}_i^2 + \tau^2}\right) \hat{\lambda}_i.$$ 

These estimates are more stable than the area-specific estimates because they borrow strength from all locations while reflecting heterogeneity among areas. The overall amount of shrinkage depends on the heterogeneity parameter $\tau^2$. Under the Bayesian approach, the posterior distribution of the area-specific parameters $\beta_i$ can be obtained by updating the first-stage area-specific estimates using information from all analyzed areas (21–23).

Posterior distributions of the model parameters were obtained with WinBugs (24).

Health impact assessment

The impact of air pollution on mortality was quantified in terms of the number of attributable deaths. The model assumes that the exposure level within each area is homogeneous. For each area, we considered the deaths of residents occurring inside the area or in municipalities within a 10-km radius from the border of the area. We excluded events occurring elsewhere. The main rationale for this choice is that a person experiencing the air pollution level of his or her residence area on a certain day could be admitted to the hospital of a neighboring municipality and then die within few days due to the effects of this exposure.

We used 2 different approaches, which we called the “macro” and “micro” approaches, for health impact assessment. The macro approach relies on the yearly average of the number of deaths and the yearly average of PM$_{10}$ levels (1). Under the micro approach, impact calculation was made day by day using the series of daily death counts and daily PM$_{10}$ levels.

If the distribution of the PM$_{10}$ daily concentrations is not strongly asymmetric and the degree of correlation between outcome and exposure is not large, the macro approach approximates the results we would obtain by applying the day-by-day, micro approach. Despite this correspondence, the opportunity to use one approach or the other depends on the chosen counterfactual scenario. While using macro modeling is the simplest way to quantify the impact under counterfactual scenarios defined in terms of yearly average, the micro approach also allows evaluation of the impact under counterfactual scenarios defined in terms of daily concentration.

The effect of air pollution is assumed to be linear without any threshold on a logarithmic scale. We counted the events attributable to exposure levels exceeding a given threshold $T_0$. Let $y_i$ be the yearly average of the number of deaths (under the macro approach) or the daily number of deaths (under the micro approach) for the $i$th area. Let $x_i$ be the yearly
average of the PM$_{10}$ concentrations (under the macro approach) or the daily concentration of PM$_{10}$ (under the micro approach) for the $i$th area. Given a certain value of the coefficient which describes the effect of air pollution in the $i$th area (for simplicity, we indicated this value with $\hat{\beta}_i$), we assumed that above the threshold $T_0$, 

$$y_i = y_0 \exp(\hat{\beta}_i(x_i - T_0)), \quad x_i > T_0,$$

where $y_0$ is the baseline number of deaths, that is, the expected number of events (yearly average or daily count, depending on the approach) we would observe for $x_i = T_0$.

The number of events attributable to PM$_{10}$ levels exceeding the threshold (i.e., attributable deaths (AD)) was then calculated as the difference between the observed number of events and the baseline number of events:

$$AD_i = y_i - y_0 = y_i \left(1 - \frac{1}{\exp(\hat{\beta}_i(x_i - T_0))}\right) = y_i \text{AF}_i,$$

with $\text{AF}_i$ representing the attributable fraction of deaths. When making a calculation under the micro approach, the total number of attributable deaths over a time span was obtained by summing the number of attributable deaths from each day.

We evaluated the impact using different values of $\hat{\beta}_i$ sampled from the posterior distribution of the area-specific effect, obtaining a whole posterior distribution of the number of attributable deaths for each area.

The percentage of deaths attributable to the exposure and the attributable community rate per 100,000 inhabitants per year were also calculated for each area (25, 26). These measures allowed relative evaluation of the PM$_{10}$ impact and comparison among areas.

**Counterfactuals**

The health impact assessment was conducted by specifying different reduction scenarios (RS). We used the following scenarios under the macro approach:

- **RS0**: $T_0 = 20$ \text{g/m}^3, that is, the WHO Air Quality Guideline threshold (11) for PM$_{10}$ annual average;
- **RS1**: $T_0 = 40$ \text{g/m}^3, that is, the EU limit for PM$_{10}$ annual average (12);
- **RS2**: $T_0$ equal to a reduction of 20% in the observed concentration of PM$_{10}$, provided it is greater than 20 \text{g/m}^3; and
- **RS3**: $T_0$ equal to a reduction of 20% in the observed concentration of PM$_{10}$, provided it is greater than 40 \text{g/m}^3.

We chose the RS2 and RS3 counterfactual scenarios according to the EU regulation, which asks for a progressive reduction of 20% in the observed concentrations until the limit is reached for all of the contexts in which the limit cannot currently be reached (12).

When applying the micro approach, we considered a further scenario:

- **RS4**: PM$_{10}$ concentrations not exceeding 50 \text{g/m}^3 more than 35 days per year, that is, the EU limit for daily averages (12).

We estimated the number of attributable deaths under the RS4 scenario by averaging 100 different pseudodata obtained by constraining to the threshold a different random set of days exceeding the 50 \text{g/m}^3 level, so that the number of days with a PM$_{10}$ concentration above the limit was set equal to 35 per year.

**RESULTS**

The 13 study areas represent approximately 35% of the Lombardy population (Table 1). The municipality of Milan has 1,299,633 inhabitants, which is 14% of the regional population. The average PM$_{10}$ level across the 13 areas was 45.4 \text{g/m}^3, with the highest values being observed in Cremona, Milan, and Lodi. In Figure 1, the areas considered in the meta-analysis are highlighted in white. These areas are the most densely populated in the region and hence the most polluted.

**Effect estimates**

Posterior distributions of the effect measures were summarized as posterior mean values and credibility intervals. Lower and upper bounds of a $(1 - \alpha)\%$ credibility interval are defined as the $(\alpha/2)$th and $(1 - \alpha/2)$th percentiles of the posterior distribution, respectively. For example, the 25th and 75th percentiles define the 50\% credibility interval for the parameter of interest. Effect estimates were expressed in terms of the percentage of variation in mortality associated with an increase of 10 \text{g/m}^3 in PM$_{10}$ concentrations at lag 0–1.

In Table 2 we show, for each of the examined areas, the posterior mean of the percent variation with its 50\% and 90\% credibility intervals (27, 28). The pooled meta-analytic estimate of percent variation in natural mortality was 0.30 (90\% credibility interval (CrI): $-0.21, 0.70$; 50\% CrI: 0.14, 0.50). We obtained the posterior distribution of the heterogeneity statistic $I^2$ (the percentage of total variability due to heterogeneity between areas) (29). There was little evidence of heterogeneity across zones ($I^2 = 4.23, 90\%$ CrI: 0.10; 28.17). Still, the posterior effect estimate was higher for the Milan area (0.63, 90\% CrI: 0.28, 1.02; 50\% CrI: 0.48, 0.78) than for the other areas, with the posterior distribution for this city being shifted upward from the overall marginal posterior effect distribution (Figure 2).

**Health impact estimates**

The health impact of PM$_{10}$ is presented by area in Table 3. Over the course of the study period, 11 areas exceeded the EU limit of 40 \text{g/m}^3 (RS1), and all 13 exceeded the WHO threshold of 20 \text{g/m}^3 (RS0). The number of attributable deaths under the counterfactual scenario RS0 was approximately 3–4 times the number of attributable deaths under the counterfactual scenario RS1. The total number of attributable deaths per year in the 13 study areas was 302 assuming the 20 \text{g/m}^3 limit, while it was 109 assuming the 40 \text{g/m}^3 limit. In both cases, most of the attributable deaths (76\% and 82\% of the total, respectively) were observed in Milan, which contained approximately 50\% of the entire study population.

Assuming a reduction of 20\% in the observed concentrations in all areas (RS2), the total number of prevented deaths...
was 101. The scenario RS3 involved only the 11 areas in which the 40 \( \mu g/m^3 \) limit was not currently being met. Assuming a 20% reduction in these areas (or a smaller reduction when this was sufficient to reach the 40 \( \mu g/m^3 \) limit), the total number of prevented deaths per year was 93.

Table 4 shows the percentage of attributable deaths and the attributable community rate under the RS0 and RS1 scenarios. Areas with the largest risk attributable to PM10 were Milan, Cremona, Lodi, and Mantova. Deaths attributable to exceeding the WHO limit of 20 \( \mu g/m^3 \) amounted to 1.4% of all natural deaths among the studied areas. The percentage of deaths attributable to exceeding the EU limit of 40 \( \mu g/m^3 \) was 0.5.

In terms of attributable community rate, 13 and 5 deaths per 100,000 inhabitants per year were attributable to yearly average PM10 levels exceeding 20 \( \mu g/m^3 \) and 40 \( \mu g/m^3 \), respectively.

Satisfying the condition of the PM10 daily limit of 50 \( \mu g/m^3 \) not being exceeded more than 35 days per year (RS4) would have resulted in an annual average slightly above 40 \( \mu g/m^3 \) for the Milan area but notably would have avoided some PM10-related deaths even in areas with lower yearly average PM10 concentrations (Table 5).

DISCUSSION

We related PM10 levels to immediate health effects on the populations of the most urbanized cities in the Lombardy region and the agricultural province of Lodi. It is reasonable to expect that the health impact of air pollution in all of Lombardy is mainly produced in these areas.

We evaluated the number of deaths we would have observed in the years 2003–2006 had the air pollution levels been different. Note that because no threshold has been proven for the toxic effects of air pollutants, the counterfactuals are arbitrary. A natural background level (i.e., from nonanthropogenic sources) of 7.5 \( \mu g/m^3 \) was used in previous impact calculation (1). However, it is questionable whether any such
Figure 2. Multiple box-and-whiskers plots of the posterior distributions of the area-specific effects of particulate matter ≤10 μm in diameter (PM_{10}) and of the overall PM_{10} effect (in gray), Lombardy, Italy, 2003–2006. The gray lines indicate a symmetric interval of 2 standard deviations around the posterior mean of the overall effect. Effects are expressed in terms of percent variation in natural mortality associated with an increase of 10 μg/m³ in PM_{10} concentrations at lag 0–1.

Table 3. Annual Average Concentration of Particulate Matter ≤10 μm in Diameter (PM_{10}) and Expected Number of Deaths Attributable to PM_{10} Evaluated With the “Macro Approach” Under Different Counterfactual Reduction Scenarios, by Area, Lombardy, Italy, 2003–2006

<table>
<thead>
<tr>
<th>Area</th>
<th>Average PM_{10} Concentration, μg/m³</th>
<th>Scenario RS0b</th>
<th>Scenario RS1c</th>
<th>Scenario RS2d</th>
<th>Scenario RS3e</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>AD 50% CrI</td>
<td>AD 50% CrI</td>
<td>AD 50% CrI</td>
<td>AD 50% CrI</td>
<td>AD 50% CrI</td>
</tr>
<tr>
<td>Bergamo</td>
<td>46.1</td>
<td>10.2</td>
<td>4.0, 17.1</td>
<td>36.9</td>
<td>3.6</td>
</tr>
<tr>
<td>Brescia</td>
<td>49.4</td>
<td>4.7</td>
<td>4.7, 16.8</td>
<td>39.5</td>
<td>1.6</td>
</tr>
<tr>
<td>Busto Arsizio</td>
<td>44.7</td>
<td>4.6</td>
<td>0.8, 9.1</td>
<td>35.8</td>
<td>1.7</td>
</tr>
<tr>
<td>Como</td>
<td>43.6</td>
<td>5.7</td>
<td>1.7, 10.5</td>
<td>34.9</td>
<td>2.1</td>
</tr>
<tr>
<td>Cremona</td>
<td>53.5</td>
<td>6.4</td>
<td>1.0, 13.5</td>
<td>42.8</td>
<td>2.0</td>
</tr>
<tr>
<td>Lecco</td>
<td>38.4</td>
<td>1.6</td>
<td>0.4, 4.4</td>
<td>30.7</td>
<td>0.7</td>
</tr>
<tr>
<td>Lodi district</td>
<td>52.6</td>
<td>19.9</td>
<td>7.7, 33.5</td>
<td>42.1</td>
<td>6.4</td>
</tr>
<tr>
<td>Mantova</td>
<td>50.6</td>
<td>5.6</td>
<td>2.1, 9.6</td>
<td>40.5</td>
<td>1.9</td>
</tr>
<tr>
<td>Milan</td>
<td>52.5</td>
<td>231.3</td>
<td>174.7, 284.3</td>
<td>42.0</td>
<td>75.3</td>
</tr>
<tr>
<td>Pavia</td>
<td>44.4</td>
<td>5.2</td>
<td>1.5, 9.9</td>
<td>35.5</td>
<td>1.9</td>
</tr>
<tr>
<td>Sondrio</td>
<td>42.8</td>
<td>1.2</td>
<td>0.2, 2.3</td>
<td>34.2</td>
<td>0.4</td>
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<td>Varese</td>
<td>29.6</td>
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<td>1.3, 4.8</td>
<td>23.7</td>
<td>0.9</td>
</tr>
<tr>
<td>Vigevano</td>
<td>42.2</td>
<td>3.0</td>
<td>0.2, 6.7</td>
<td>33.8</td>
<td>1.2</td>
</tr>
</tbody>
</table>

Abbreviations: AD, attributable deaths; CrI, credibility interval; PM_{10}, particulate matter ≤10 μm in diameter; RS, reduction scenario.
b Annual average PM_{10} concentration observed over the study period (2003–2006).  
c Assumes that the 20 μg/m³ World Health Organization limit for annual average PM_{10} concentration is not exceeded.  
d Assumes that the 40 μg/m³ European Union limit for annual average PM_{10} concentration is not exceeded.  
e Assumes a 20% reduction in the annual average concentration of PM_{10}, provided it is greater than 20 μg/m³.  
f Assumes a 20% reduction in the annual average concentration of PM_{10}, provided it is greater than 40 μg/m³.  
g Threshold for the annual average concentration of PM_{10} under scenario RS2.  
h Threshold for the annual average concentration of PM_{10} under scenario RS3.
level could reasonably be reached through any reduction strategy in urbanized areas. We used the WHO value of 20 \( \mu g/m^3 \) as a background threshold value for the annual average PM\(_{10}\) concentration. This represents a target level to be reached in the future but is not reachable in the next few years (11). The EU limit of 40 \( \mu g/m^3 \) was chosen as a second counterfactual. This is greatly above the PM\(_{10}\) concentrations considered dangerous. By setting this limit, the EU delineated a point above which arguments on health protection should prevail over arguments based on economic considerations. We also evaluated 2 other counterfactual scenarios. We calculated the number of prevented deaths due to decreasing pollution concentrations of 20% of the actual measured values, provided that the annual average was above 20 \( \mu g/m^3 \) or 40 \( \mu g/m^3 \).

Since a 20% reduction of emissions is a realistic target, as is also stated in the European legislation, these scenarios allowed us to appraise how far we are from the 2 targets.

In the design of epidemiologic time-series studies, only the present population is usually considered, because any immediate effect of air pollution on persons not present is questionable. We followed this approach in order to avoid exposure misspecification bias in the estimated concentration-response relation, while in health impact assessment we applied the effect estimates to a larger set of events: deaths among the resident population occurring inside the area or in municipalities within a 10-km radius from the border of the area. In fact, considering only the deaths occurring inside the municipality of interest could cause considerable underestimation of impact, linked to the catchment area of the nearest hospitals and to the fact that in Lombardy 40%–50% of deaths occur in hospitals (30). In the analysis, we used the lag 0–1 concentration for comparability with the current literature and consistency with the evidence that PM\(_{10}\) has an immediate effect on mortality in comparable populations (31, 32). However, note that this choice can give rise to a certain degree of underestimation of the air pollutant impact if the PM\(_{10}\) effect 2 days or a few more days after the concentration peak is not negligible (18, 32).

We calculated the number of deaths attributable to PM\(_{10}\) exposure, regardless of the life expectancy of those who died. Although we cannot exclude the possibility that PM\(_{10}\) exposure can partly precipitate death in very frail people by a period of several days to a few weeks, there is evidence that the observed effects are not due primarily to this short-term mortality displacement (10, 33). Therefore, we expect that the observed impact would not be negligible, even in terms of loss of life expectancy.

The model adopted assumes a linear effect of air pollutants on a logarithmic scale, relying on previous findings from studies carried out in large cities, which are frequently characterized by high air pollution levels (15). In the study by Cohen et al. (2), the burden of disease attributable to air

<table>
<thead>
<tr>
<th>Area</th>
<th>Average PM(_{10}) Concentration,a ( \mu g/m^3 )</th>
<th>RS0(^b)</th>
<th>RS1(^c)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>AD ( %AD ) ACR</td>
<td>AD ( %AD ) ACR</td>
<td></td>
</tr>
<tr>
<td>Bergamo</td>
<td>46.1 10.2 0.87 8.8 2.4 0.21 2.1</td>
<td></td>
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</tr>
<tr>
<td>Brescia</td>
<td>49.4 4.7 0.38 2.5 1.5 0.12 0.8</td>
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<tr>
<td>Busto Arsizio</td>
<td>44.7 4.6 0.68 5.7 0.9 0.13 1.1</td>
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</tr>
<tr>
<td>Como</td>
<td>43.6 5.7 0.73 6.8 0.9 0.11 1.1</td>
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</tr>
<tr>
<td>Cremona</td>
<td>53.5 6.4 0.83 8.9 2.6 0.34 3.6</td>
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<tr>
<td>Lecco</td>
<td>38.4 1.6 0.34 3.4 0 0.00 0.0</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Lodi district</td>
<td>52.6 19.9 1.02 9.1 7.7 0.40 3.5</td>
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<td></td>
</tr>
<tr>
<td>Mantova</td>
<td>50.6 5.6 1.03 11.7 1.9 0.35 4.0</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Milan</td>
<td>52.5 231.3 2.03 17.8 89.5 0.78 6.9</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Pavia</td>
<td>44.4 5.2 0.70 7.4 0.9 0.12 1.3</td>
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</tr>
<tr>
<td>Sondrio</td>
<td>42.8 1.2 0.62 5.4 0.1 0.05 0.4</td>
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<tr>
<td>Varese</td>
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<tr>
<td>Vigevano</td>
<td>42.2 3.0 0.52 4.9 0.3 0.05 0.5</td>
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<tr>
<td>Total</td>
<td>302.4 1.42 12.6 108.7 0.51 4.6</td>
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</tr>
</tbody>
</table>

Abbreviations: ACR, attributable community rate; AD, attributable deaths; PM\(_{10}\), particulate matter \( \leq 10 \mu m \) in diameter; RS, reduction scenario.

- \(^a\) Annual average PM\(_{10}\) concentration observed over the study period (2003–2006).
- \(^b\) Assumes that the 20 \( \mu g/m^3 \) World Health Organization limit for annual average PM\(_{10}\) concentration is not exceeded.
- \(^c\) Assumes that the 40 \( \mu g/m^3 \) European Union limit for annual average PM\(_{10}\) concentration is not exceeded.
pollution was evaluated defining an upper bound for the annual average concentration. For cities with annual average PM$_{10}$ concentrations exceeding 100 $\mu$g/m$^3$, maximum concentration equal to 100 $\mu$g/m$^3$ was used for health impact assessment, because a linear exposure model could produce unrealistically large estimates of attributable mortality in the most extremely polluted regions. The annual average concentrations of PM$_{10}$ observed in Lombardy are not so high as to justify the use of an upper bound.

Because of a lack of statistical power, we did not assess the impact of PM$_{10}$ by age class. In fact, in most of the included cities, the age-specific estimates of the air pollutant effect relied on a very small number of daily events, particularly in the younger age classes. Using the age-adjusted effect estimates, we probably underestimated the impact of PM$_{10}$, because we did not account for the likely larger vulnerability and higher baseline risk of elderly persons. For the same reason, we did not assess the impact by specific cause of death, even if this could have, in principle, improved the specificity of our evaluation.

Large national/continental meta-analyses on short-term effects of air pollutants usually show a substantial heterogeneity of the effect. This implies that the portability of the effect estimates is questionable and that when performing impact

<table>
<thead>
<tr>
<th>Area</th>
<th>Average PM$_{10}$ Concentration$^{b}$ $\mu$g/m$^3$</th>
<th>Average PM$_{10}$ Concentration Expected Under RS4, $\mu$g/m$^3$</th>
<th>No. of Attributable Deaths</th>
<th>50% Credibility Interval</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bergamo</td>
<td>46.1</td>
<td>38.9</td>
<td>3.0</td>
<td>1.2, 5.0</td>
</tr>
<tr>
<td>Brescia</td>
<td>49.4</td>
<td>40.9</td>
<td>1.4</td>
<td>−1.4, 5.0</td>
</tr>
<tr>
<td>Busto Arsizio</td>
<td>44.7</td>
<td>37.9</td>
<td>1.3</td>
<td>0.2, 2.6</td>
</tr>
<tr>
<td>Como</td>
<td>43.6</td>
<td>38.6</td>
<td>1.3</td>
<td>0.4, 2.4</td>
</tr>
<tr>
<td>Cremona</td>
<td>53.5</td>
<td>43.2</td>
<td>2.1</td>
<td>0.3, 4.5</td>
</tr>
<tr>
<td>Lecco</td>
<td>38.4</td>
<td>35.3</td>
<td>0.3</td>
<td>−0.1, 0.7</td>
</tr>
<tr>
<td>Lodi district</td>
<td>52.6</td>
<td>42.8</td>
<td>6.5</td>
<td>2.5, 11.0</td>
</tr>
<tr>
<td>Mantova</td>
<td>50.6</td>
<td>42.6</td>
<td>1.5</td>
<td>0.6, 2.6</td>
</tr>
<tr>
<td>Milan</td>
<td>52.5</td>
<td>40.2</td>
<td>96.6</td>
<td>73.1, 118.4</td>
</tr>
<tr>
<td>Pavia</td>
<td>44.4</td>
<td>38.2</td>
<td>1.4</td>
<td>0.4, 2.7</td>
</tr>
<tr>
<td>Sondrio</td>
<td>42.8</td>
<td>37.6</td>
<td>0.3</td>
<td>0.1, 0.6</td>
</tr>
<tr>
<td>Varese</td>
<td>29.6</td>
<td>29.0</td>
<td>0.2</td>
<td>0.1, 0.4</td>
</tr>
<tr>
<td>Vigevano</td>
<td>42.2</td>
<td>36.3</td>
<td>0.8</td>
<td>0.0, 1.8</td>
</tr>
</tbody>
</table>

Abbreviations: PM$_{10}$, particulate matter $\leq 10 \mu$m in diameter; RS, reduction scenario.

$^{a}$ Assumes that the daily limit of 50 $\mu$g/m$^3$ for PM$_{10}$ is not exceeded more than 35 days per year in the European Union limit for daily averages.

$^{b}$ Annual average PM$_{10}$ concentration observed over the study period (2003–2006).

Table 6. Daily Average Number of Deaths, Concentration of Particulate Matter $\leq 10 \mu$m in Diameter (PM$_{10}$), Posterior Mean of the City-Specific PM$_{10}$ Effect, and Expected Number of Attributable Deaths Under Different PM$_{10}$ Reduction Scenarios for the City of Milan as Compared With the MISA Study, Lombardy, Italy$^{a}$

<table>
<thead>
<tr>
<th>Daily Average No. of Deaths</th>
<th>PM$_{10}$ Concentration, $\mu$g/m$^3$</th>
<th>City-Specific Distribution of % Variation</th>
<th>No. of Attributable Deaths Under Different Scenarios</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Mean</td>
<td>95th Percentile</td>
<td>Posterior Mean</td>
</tr>
<tr>
<td>MISA (1996–2002)</td>
<td>29.1</td>
<td>56.3</td>
<td>135.1</td>
</tr>
<tr>
<td>Lombardy health impact assessment (2003–2006)</td>
<td>31.3</td>
<td>52.5</td>
<td>120.8</td>
</tr>
</tbody>
</table>

Abbreviations: MISA, Meta-Analysis of the Italian Studies on Short-Term Effects of Air Pollution; PM$_{10}$, particulate matter $\leq 10 \mu$m in diameter; RS, reduction scenario.

$^{a}$ Effects are expressed in terms of percent variation in natural mortality associated with an increase of 10 $\mu$g/m$^3$ in PM$_{10}$ concentrations at lag 0–1.

$^{b}$ Assumes that the 20 $\mu$g/m$^3$ World Health Organization limit for annual average PM$_{10}$ concentration is not exceeded.

$^{c}$ Assumes that the 40 $\mu$g/m3 European Union limit for annual average PM$_{10}$ concentration is not exceeded.
analysis we should use specific estimates for the population considered (25). This is the reason why we derived the effect estimates to be used for attributable death calculation from the same population. However, even within Lombardy, we found some evidence of heterogeneity, and we found a larger effect for the city of Milan than for the other areas. For this reason, we used posterior area-specific distributions, which appropriately reflect differences among locations.

The effect estimates appeared to be increasing for the metropolitan area of Milan from the period 1996–2002 to the more recent time span of 2003–2006 (Table 6). The percent variation in total natural mortality associated with an increase in estimated PM10 effect was 0.36 in the first period (MISA study (10)) and 0.63 in the second period. The range of PM10 concentrations did not change greatly enough to support the hypothesis that the observed difference was due to a nonlinear concentration-response curve (the mean and 95th percentile of daily PM10 levels were 56.3 and 135.1, respectively, in 1996–2002 vs. 52.5 and 120.8, respectively, in 2003–2006). The observed discrepancy could simply be due to sampling variability, but it was shown elsewhere that PM10 effect estimates have increased over time even in other Italian cities (31); hence, we cannot exclude more complex explanations. The increasing trend in the effect could have been induced by the improvement in PM10 measurement in the most recent years, which probably reduced exposure misclassification (instruments for measuring particulate concentrations in the Lombardy air quality monitoring network were upgraded after 2002, with a gain in precision and accuracy). A more interesting hypothesis is that the observed change in the estimated effect was caused by variation in PM10 composition over time (for example, an increasing proportion of particulate matter ≤2.5 μm in diameter in total suspended particles) or by effect modification involving meteorologic conditions. Finally, the observed discrepancy could be due to an increase in the vulnerable fraction of exposed persons related to sociodemographic phenomena such as aging, changes in the economic status of the population, or immigration. Regarding aging, the aging index for the Lombardy region, calculated as the number of persons aged 65 years or over per hundred persons under age 14, increased from 123.5 in 1995 to 143.1 in 2008 (34).

Concerning the impact, the number of attributable deaths found in Milan was larger than the number of attributable deaths calculated with the same approach for the 1996–2002 period, despite the fact that the PM10 levels decreased over time (Table 6): The reduction in average PM10 concentration seems to have been compensated for by the larger effect estimate in the more recent period. This result, coupled with the likelihood that an aging population will increase vulnerability to air pollution, makes further development of policies for pollution reduction both crucial and urgent.

The goal of the WHO threshold of 20 μg/m³ is still out of reach, but we showed that even less stringent policies could bring about a relevant reduction in the number of attributable deaths. The health impact in the 13 study areas was 1.4% of the total number of natural deaths when considering the annual PM10 concentration limit of 20 μg/m³ and 0.5% when considering the limit of 40 μg/m³ (which was met by only 2 cities). Given the mortality rate and the air pollutant level observed during the study period, approaching the 20 μg/m³ limit through a reduction of current annual PM10 concentrations by 20% in all municipalities with an annual average above 20 μg/m³ would have prevented about 33% of the mortality burden (31%) if the 20% reduction policy had been restricted to only those municipalities with an annual average above 40 μg/m³.

The mortality burden attributable to exceeding the EU yearly limit of 40 μg/m³ for the annual average contributed to 93% of the mortality burden under the RS4 scenario (in which the daily limit of 50 μg/m³ for PM10 was not exceeded more than 35 days per year). In this specific situation, these 2 counterfactual scenarios provided quite similar results, but this could not be true under different distributions of daily PM10 levels.

In conclusion, we estimated that in the major cities of Lombardy, annual average PM10 levels exceeding the WHO limit of 20 μg/m³ and the EU limit of 40 μg/m³ were responsible for 13 and 5 deaths per 100,000 inhabitants per year, respectively. At the same time, we estimated that a 20% reduction in the existing PM10 levels could reduce by more than 30% the burden of short-term deaths linked to ambient air pollution exposure. Empirical studies showing an increase in life expectancy in parallel with the decrease in ambient fine-particle air pollution add credibility to these estimates and underline their relevance to public health goals (35). Therefore, policies for the reduction of air pollution appear to be necessary, and their implementation will be rewarding in terms of the protection and improvement of individual and community health.

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REFERENCES


