Effect of air pollution on pediatric respiratory emergency room visits and hospital admissions

S.C.L. Farhat1,2, R.L.P. Paulo2, T.M. Shimoda2, G.M.S. Conceição1, C.A. Lin1,2, A.L.F. Braga1,4, M.P.N. Warth3 and P.H.N. Saldiva1

1Laboratório de Poluição Atmosférica Experimental, Departamento de Patologia (LIM05), 2Departamento de Pediatria, Pronto Socorro do Instituto da Criança, and 3Departamento de Clínica Médica, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brasil 4Programa de Pediatria Ambiental, Departamento de Pediatria, Faculdade de Medicina, Universidade de Santo Amaro, Santo Amaro, SP, Brasil

Abstract

In order to assess the effect of air pollution on pediatric respiratory morbidity, we carried out a time series study using daily levels of PM10, SO2, NO2, ozone, and CO and daily numbers of pediatric respiratory emergency room visits and hospital admissions at the Children’s Institute of the University of São Paulo Medical School, from August 1996 to August 1997. In this period there were 43,635 hospital emergency room visits, 4534 of which were due to lower respiratory tract disease. The total number of hospital admissions was 6785, 1021 of which were due to lower respiratory tract infectious and/or obstructive diseases. The three health end-points under investigation were the daily number of emergency room visits due to lower respiratory tract diseases, hospital admissions due to pneumonia, and hospital admissions due to asthma or bronchiolitis. Generalized additive Poisson regression models were fitted, controlling for smooth functions of time, temperature and humidity, and an indicator of weekdays. NO2 was positively associated with all outcomes. Inter-quartile range increases (65.04 µg/m3) in NO2 moving averages were associated with an 18.4% increase (95% confidence interval, 95% CI = 12.5-24.3) in emergency room visits due to lower respiratory tract disease (4-day moving average), a 17.6% increase (95% CI = 3.3-32.7) in hospital admissions due to pneumonia or bronchopneumonia (3-day moving average), and a 31.4% increase (95% CI = 7.2-55.7) in hospital admissions due to asthma or bronchiolitis (2-day moving average). The study showed that air pollution considerably affects children’s respiratory morbidity, deserving attention from the health authorities.

Key words
- Time-series
- Poisson
- Air pollution
- Children
- Respiratory morbidity

Introduction

A large number of studies conducted worldwide by different research groups have demonstrated excessive mortality (1-8) and morbidity (9-18) associated with increases in air pollution. Some have pointed out children as a group highly susceptible to the effects of air pollution, mainly in terms of respiratory diseases (6,9,12-14,16-18).

Despite the increasing number of studies in this area, there are still points that deserve...
further clarification. While air pollution has decreased in urban areas of developed countries, high air pollutant concentrations have been recorded in large urban conglomerates of the developing world (6,7,19-21). In megacities of Asia, Africa and Latin America, air pollution has become a permanent component of the environment.

Time-series analysis seems to be the best approach to detect effects (22,23) in areas where pollution tends to increase. However, the proper application of the time-series approach requires high quality data regarding air pollution and the daily number of a given health outcome such as mortality counts or number of hospital admissions, conditions that are not always satisfied in areas with less developed economies.

São Paulo, the largest city in South America and the third most populated in the world, presents adequate conditions to be a good scenario for this kind of analysis. Presently, São Paulo has reliable air pollution and mortality data, and this information was used by our group in previous studies (6,7,24) indicating that air pollution affects mortality in our environment. The objective of our research was to evaluate the impact of air pollution on morbidity based on data gathered from a large emergency hospital used as an “environmental probe” (25) and the daily counts of hospital admissions of children in the public health system due to respiratory diseases (26,27). These investigations indicated that the effects of air pollution, when expressed in terms of morbidity, are higher than those of mortality, supporting the association between quality of the air and quality of life.

In the present study, we prospectively collected data on children’s lower respiratory tract morbidity (upper airway diseases excluded), expressed in terms of emergency room visits and admissions to our teaching hospital. The associations between these two specific health end-points (emergency room visits and hospital admissions) and air pollution were investigated in order to determine the pollutants more significantly associated with respiratory morbidity and the lag between worsening in the air quality and the necessity of hospital care and, considering only the hospital admission population, to compare the effects of air pollution on two categories of respiratory diseases, i.e., parenchymal disease (pneumonia and bronchopneumonia) and airway disease (asthma and bronchiolitis).

Material and Methods

Data collection

We collected data on respiratory morbidity of children under 13 years of age, from August 1996 to August 1997, at the Children’s Institute of the University of São Paulo Medical School. The hospital is located in a central area in downtown São Paulo and serves as reference center for the entire city. The physician staff of the Children’s Institute attended the patients and a highly experienced pediatrician always checked the diagnoses. Additionally, Dr. Farhat double-checked the coding of the diseases. Two data sets were collected. One data set included the total number of emergency room visits due to lower tract respiratory diseases (International Code of Diseases (ICD) 9th Revision 466; 480-519). The emergency service provides practically obligatory medical care even if it is necessary to keep a child under provisional bedding conditions. Thus, the number of respiratory emergency room visits in one hospital may represent an estimator of the total respiratory morbidity because of the limitations of the hospital’s capacity. This data set was similar to that employed by Lin et al. (25), i.e., hospital admissions due to pneumonia or bronchopneumonia (ICD 9th Revision 480-486), asthma (ICD 9th Revision 493), and bronchiolitis (ICD 9th Revision 466) and these respiratory diseases were coded
based on history, physical examination and radiology. Because we are aware of the potential misclassification of wheezing-related diseases, we included both asthma and bronchiolitis in the same disease group called obstructive diseases. Some of the children who are admitted to the emergency room require more prolonged medical care and are admitted to the infirmaries of the Children’s Institute. The number of hospital admissions is smaller than that of emergency room visits, but more detailed information can be collected about these patients. This second data set, which includes only the most severe respiratory cases, is similar to that used by Braga and colleagues (26,27).

During the study period, the São Paulo State Sanitary Agency (CETESB) had 13 monitoring stations located in different regions of the city. CETESB provides daily values of particulate matter with aerodynamic diameter smaller than 10 µm (PM$_{10}$) and sulfate dioxide (SO$_2$), measured at 13 stations, and nitrogen dioxide (NO$_2$) measured at 6 stations. For ozone (O$_3$), CETESB provides the highest hourly averages measured at 6 stations, and for carbon monoxide (CO) the highest 8-h moving average measured at 8 stations. The average of the available measurements was calculated for each day and was considered to be representative of citywide conditions. The lowest temperature of the day and the daily relative humidity were obtained from the Institute of Astronomy and Geophysics of the University of São Paulo.

Statistical analysis

We defined three outcomes: lower respiratory emergency room visits and hospital admissions due to pneumonia or bronchopneumonia and to asthma or bronchiolitis. Poisson regression techniques were adopted in this analysis because daily numbers of both emergency room visits and hospital admissions are count events and, therefore, present Poisson distribution. Generalized additive models (28) were used because they better fit epidemiologic time-series studies of the effects of air pollution on health. They allow a more flexible modeling of the association between health end points and the predictor variables due to the possibility to include linear and smooth functions for the confounders in the same model. We adopted the loess, a locally weighted running line smoother, for time, temperature, and humidity and day of the week. In each model, the span for the smooth function of time was chosen to remove long-term trends from the data, removing seasonality and minimizing the autocorrelation of the residuals. In models in which serial correlation of the residuals remained after the adjustment of time span we incorporated autoregressive terms (29).

Spans of smooth functions for temperature and humidity variables were chosen in order to minimize Akaike’s Information Criteria (30). In addition, indicators for each day of the week were included in the models. Robust regression (M-estimation) was used in order to reduce sensitivity to outliers in the dependent variable.

To determine the possible lag between the increases in air pollution and emergency room visits or hospital admissions, we adopted regression models considering different lag structures for each pollutant, ranging from the concurrent day to 7 days. Analyses were conducted using single-, 2-pollutant, and multi-pollutant models. The results are reported in terms of percent increase in the outcomes and the 95% confidence intervals were estimated assuming normal distribution of the estimated regression coefficients.

These regression models were estimated using the S-PLUS software (31).

Results

During the 13-month study period, PM$_{10}$
values (average concentration = 62.6 µg/m³) exceeded the annual standard adopted for inhalable particles (yearly mean of 50 µg/m³). The daily standard of PM₁₀ (24 h mean of 150 µg/m³) was exceeded three times. The yearly mean for NO₂ (average concentration = 125.3 µg/m³) exceeded the standard adopted of 100 µg/m³ and the daily air quality standard level (320 µg/m³) was exceeded three times. The 1-h standard for O₃ (160 µg/m³) was exceeded fourteen times, and the 8-h primary standard for CO (9 ppm) was exceeded four times. SO₂ levels remained below the regulatory standards.

Descriptive analyses of the variables employed in the study are presented in Table 1. During the study period there were 43,635 hospital emergency room visits, 4534 of which were due to lower respiratory tract disease. The total number of admissions to the Children’s Institute was 6785, 1021 of which were due to lower respiratory tract infectious and/or obstructive diseases.

The Pearson correlation coefficients between air pollutants and weather variables are presented in Table 2. The pollutants were highly correlated to each other except CO and O₃. Also, all pollutants were negatively correlated with temperature and humidity.

Figure 1 shows the percent increases and 95% confidence intervals in lower respiratory tract emergency room visits due to interquartile range increases in 5-day moving average of PM₁₀, 4-day moving average of NO₂, 5-day moving average of SO₂, 4-day moving average of O₃, and 2-day moving average of CO estimated by single-pollutant models. All of them contributed to an increase in emergency room visits. Table 3 presents the effects estimated using co-pollutant and multi-pollutant models. Only the effect of NO₂ did not change substantially in the two approaches, remaining positive and statistically significant, whereas the others presented a highly unstable behavior. PM₁₀ and O₃ lost their significance in co-pollutant models when they were included with NO₂, and CO and SO₂ effects only resisted the inclusion of O₃ and CO, respectively. Unexpected and implausible protective effects were observed for SO₂ when it was included in a co-pollutant model with NO₂ or in the multi-pollutant model.

Figure 2 shows the percent increases and 95% confidence intervals in pneumonia or bronchopneumonia hospital admissions due to interquartile range increases in 6-day moving average of PM₁₀, 3-day moving average of NO₂, 6-day moving average of SO₂, 7-day moving average of CO, and 6-day moving average of O₃ estimated by single-pollutant models.
Table 3. Increases of lower respiratory tract disease emergency room visits due to interquartile range increases in 5-day moving average of PM10, 4-day moving average of NO2, 5-day moving average of SO2, 4-day moving average of O3, and 2-day moving average of CO in single-pollutant models. PM10 = particulate matter with aerodynamic diameter smaller than 10 µm.

<table>
<thead>
<tr>
<th></th>
<th>Co-pollutant models</th>
<th>Multi-pollutant model</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>PM10</td>
<td>NO2</td>
</tr>
<tr>
<td>PM10</td>
<td>-</td>
<td>2.1</td>
</tr>
<tr>
<td>NO2</td>
<td>16.1*</td>
<td>-</td>
</tr>
<tr>
<td>SO2</td>
<td>(-10.8,6.2)</td>
<td>(-7.1,11.3)</td>
</tr>
<tr>
<td>O3</td>
<td>-3.44</td>
<td>-7.0*</td>
</tr>
<tr>
<td>CO</td>
<td>(-5.6,5.3)</td>
<td>(-6.7,4.2)</td>
</tr>
</tbody>
</table>

Data are reported as percent increase and 95% confidence interval. PM10 = particulate matter with aerodynamic diameter smaller than 10 µm.

*P < 0.05 (likelihood-ratio test: full model vs reduced model).
moving average of $O_3$, and 2-day moving average of CO in single-pollutant models. The five pollutants presented positive and statistically significant associations with the outcome and similar effect sizes. Table 4 shows the effects of air pollutants on pneumonia or bronchopneumonia hospital admissions using co-pollutant and multi-pollutant models. The inclusion of co-pollutants reduced all effects observed in single-pollutant models. PM$_{10}$ and SO$_2$ resisted the inclusion of $O_3$ and CO, while the effect of $O_3$ was not substantially affected by CO. Despite these decreases, no protective effect was detected. In a multi-pollutant model all pollutants remained positively associated with the outcome although losing statistical significance.

**Table 4.** Increases of pneumonia or bronchopneumonia hospital admissions due to interquartile range increases in 6-day moving average of PM$_{10}$, 3-day moving average of NO$_2$, 6-day moving average of SO$_2$, 7-day moving average of $O_3$, and 2-day moving average of CO in a multi-pollutant and co-pollutant models.

<table>
<thead>
<tr>
<th>Co-pollutant models</th>
<th>Multi-pollutant model</th>
</tr>
</thead>
<tbody>
<tr>
<td>PM$_{10}$</td>
<td>NO$_2$</td>
</tr>
<tr>
<td>PM$_{10}$</td>
<td>-</td>
</tr>
<tr>
<td></td>
<td>(-3.8,33.4)</td>
</tr>
<tr>
<td>NO$_2$</td>
<td>8.11</td>
</tr>
<tr>
<td></td>
<td>(-11.4,27.6)</td>
</tr>
<tr>
<td>SO$_2$</td>
<td>13.3</td>
</tr>
<tr>
<td></td>
<td>(-5.7,32.3)</td>
</tr>
<tr>
<td>O$_3$</td>
<td>10.9</td>
</tr>
<tr>
<td></td>
<td>(-10.4,32.2)</td>
</tr>
<tr>
<td>CO</td>
<td>4.4</td>
</tr>
<tr>
<td></td>
<td>(-7.9,16.7)</td>
</tr>
</tbody>
</table>

Data are reported as percent increase and 95% confidence interval. PM$_{10}$ = particulate matter with aerodynamic diameter smaller than 10 µm. *P < 0.05 (likelihood-ratio test: full model vs reduced model).

**Table 5.** Increases of asthma or bronchiolitis hospital admissions due to interquartile range increases in 2-day moving average of PM$_{10}$, 2-day moving average of NO$_2$, 2-day moving average of SO$_2$, 3-day moving average of O$_3$, and 2-day moving average of CO in a multi-pollutant and co-pollutant models.

<table>
<thead>
<tr>
<th>Co-pollutant models</th>
<th>Multi-pollutant model</th>
</tr>
</thead>
<tbody>
<tr>
<td>PM$_{10}$</td>
<td>NO$_2$</td>
</tr>
<tr>
<td>PM$_{10}$</td>
<td>-</td>
</tr>
<tr>
<td></td>
<td>(-50.0,28.0)</td>
</tr>
<tr>
<td>NO$_2$</td>
<td>47.7*</td>
</tr>
<tr>
<td></td>
<td>(1.15,94.2)</td>
</tr>
<tr>
<td>SO$_2$</td>
<td>3.8</td>
</tr>
<tr>
<td></td>
<td>(-23.3,21.0)</td>
</tr>
<tr>
<td>O$_3$</td>
<td>14.2</td>
</tr>
<tr>
<td></td>
<td>(-11.9,40.3)</td>
</tr>
<tr>
<td>CO</td>
<td>6.1</td>
</tr>
<tr>
<td></td>
<td>(-14.9,27.1)</td>
</tr>
</tbody>
</table>

Data are reported as percent increase and 95% confidence interval. PM$_{10}$ = particulate matter with aerodynamic diameter smaller than 10 µm. *P < 0.05 (likelihood-ratio test: full model vs reduced model).
Air pollutants increased asthma or bronchiolitis hospital admissions (Figure 3) and, although these effects were not statistically significant, except for NO₂, the size of the effects was almost the same as that observed for pneumonia or bronchopneumonia hospital admissions. The inclusion of PM₁₀ and SO₂ did not reduce the effect of NO₂. However, implausible protective effects appeared in both co-pollutant and multi-pollutant analyses (Table 5).

**Discussion**

In the present study, we found significant and positive associations between air pollution and respiratory morbidity in children. The effect of air pollution was detected in terms of emergency room visits and hospital admissions. In terms of respiratory admissions, when the cases were separated into diseases of the airways and diseases affecting pulmonary parenchyma, the effect was positive in both cases. The associations found occurred within a short period of time of the order of a few days. Although the daily air quality standards were exceeded on a few days during the study period, the association between air pollution and respiratory morbidity was robust enough to resist different options of statistical modeling, reinforcing the previous concept that air pollution may affect health even during non-episodic events of pollution (9-11).

It was difficult to ascribe the pathogenesis of the observed effect to a single pollutant. For instance, emergency room visits were significantly associated with all pollutants, probably because of the high degree of correlation among them, which is expected to occur in a pollution scenario generated by automotive emissions. The difficulty in isolating the most dangerous pollutant is a limitation of the ecological approach employed by us, and was present in our previous animal and epidemiological studies (7,25,32,33). In fact, it is not even possible to determine whether the observed effects were due to the measured pollutants or whether they represent a proxy variable of other compounds present in the atmosphere of São Paulo and that are not evaluated by the existing air pollution monitoring system. Anyway, as reported in the experimental studies by our group (32,33), the mixture of air pollutants present in São Paulo induces damage to the mucociliary epithelium leading to loss of cilia, increased mucus density, increased inflammatory responses, and increased frequency of inflammatory and infectious diseases.

When obstructive pathologies were evaluated the number of pollutants associated with morbidity decreased. Nitrogen dioxide was the pollutant most robustly associated with cases of severe asthma and bronchiolitis that required hospital admission. This is not the first time that this pollutant appears as a strong indicator of atmospheric risk in São Paulo. Our studies on children and on intrauterine mortality (6,24) also reported the same finding. We do not know whether this association is real, or if nitrogen oxides represent a good indicator of the complex mixture present in automotive emissions, as already mentioned. Moreover, in São Paulo nitrogen oxides represent one of the major sources of atmospheric oxidants generated by photochemical reactions. Thus, we cannot establish an explicit association between respiratory morbidity and nitrogen oxides per se.

The magnitude of the effect of air pollution on respiratory morbidity varies with the morbidity indicator (emergency room visits or hospital admissions) and with the pollutant considered. In the case of patients who required hospital admissions, this number was higher, reaching a peak for obstructive diseases and NO₂ levels (31.4%).

The three options of health outcomes and pollutants considered in the present study make the comparison of our results with the existing literature somewhat difficult. As a
general rule, the severity of the health effects associated with air pollution is greater for patients that require hospital admission than for patients looking for emergency care. Thus, in the present series, the estimated impact of air pollution varied with the severity of the health outcome. In addition, since several pollutants presented significant associations with respiratory morbidity, there was a variation of the estimated morbidity depending on the pollutant used as a predictor in the statistical modeling. These problems are aggravated by the relatively small number of studies focusing specifically on air pollution and children’s hospital admissions. For instance, our mean effect of pollution on emergency room visits is comparable to that reported by Lin et al. (25) and Braga et al. (26). Sunyer et al. (34) reported a relative risk of hospital admissions due to asthma (somewhat equivalent to our admissions due to asthma or bronchitis) of 1026 per 50 µg/m³ of NO₂, a value considerably lower than that reported in our study for an equivalent variation of this pollutant (1.16). In fact, the effects of air pollution on children’s health in São Paulo are somewhat greater than those reported for adults in other locations (35,36).

The consistency of our results reflects the usefulness of our teaching hospital as a sentinel for epidemiological studies. This is quite a convenient situation for locations like São Paulo and other Latin American cities where centralized morbidity data are not available. The large university hospitals play a major role in medical care in developing countries and probably report the most reliable morbidity data in these locations, permitting the collection of a large number of morbidity events that may be modeled on ecological studies such as ours. The Children’s Institute attends patients from most of São Paulo’s neighborhoods. In this specific case, its patients may be accepted as a representative sample of the population of São Paulo city. Thus, the validity of the use of morbidity data collected from a single large hospital in studies focusing the health impact of a widespread toxic agent such as air pollution should be further explored in other investigations at other locations.

This study reports a significant effect of air pollution on respiratory morbidity in São Paulo. Several pollutants were associated with increased respiratory events, making it difficult to isolate a single agent as the main atmospheric contaminant. The effects are strong enough to consider air pollution as a public health problem in our city, demanding policies of automobile emission control from our authorities.

References

Air pollution and respiratory morbidity


