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Effects of particulate matter on hospital admissions for respiratory diseases: an ecological study based on 12.5 years of time series data in Shanghai

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Abstract

Background: Previous epidemiological studies on the association between short-term exposure to particulate matter (PM) with hospital admission in major cities in China were limited to shorter study periods or a single hospital. The aim of this ecological study based on a 12.5-year time series was to investigate the association of short-term exposure to PM with aerodynamic diameter \leq 2.5 μ m (PM_{2.5}) and aerodynamic diameter \leq 10 μ m (PM₁₀) with hospital admissions for respiratory diseases.

Methods: Daily hospital admissions data were from the Shanghai Medical Insurance System for the period January 1, 2008 to July 31, 2020. We estimated the percentage change with its 95% confidence interval (CI) for each $10 \,\mu\text{g/m}^3$ increase in the level of $PM_{2.5}$ and PM_{10} after adjustment for calendar time, day of the week, public holidays, and meteorological factors applying a generalized additive model with a quasi-Poisson distribution.

Results: There were 1,960,361 hospital admissions for respiratory diseases in Shanghai during the study period. A $10\,\mu\text{g/m}^3$ increase in the level of each class of PM was associated with increased total respiratory diseases when the lag time was 0 day (PM_{2.5}: 0.755%; 95% CI: 0.422, 1.089%; PM₁₀: 0.250%; 95% CI: 0.042, 0.459%). The PM_{2.5} and PM₁₀ levels also had positive associations with admissions for COPD, asthma, and pneumonia. Stratified analyses demonstrated stronger effects in patients more than 45 years old and during the cold season. Total respiratory diseases increased linearly with PM concentration from 0 to $100\,\mu\text{g/m}^3$, and increased more slowly at higher PM concentrations.

Conclusions: This time-series study suggests that short-term exposure to PM increased the risk for hospital admission for respiratory diseases, even at low concentrations. These findings suggest that reducing atmospheric PM concentrations may reduce hospital admissions for respiratory diseases.

Keywords: Particulate matter, Respiratory diseases, Hospital admission, Time series

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Background

Respiratory diseases, including chronic obstructive pulmonary disease (COPD), asthma, interstitial lung disease, and pulmonary sarcoidosis, are major public health problems that impose a huge disease burden on society [1]. The disability-adjusted life years (DALYs) of respiratory diseases increased from 97.2 to 112.3 million a year from



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1990 to 2017 [2]. There are two well-known risk factors associated with respiratory diseases: behavioral factors (e.g. smoking) and environmental factors (e.g. air pollution) [2].

China is now faced with severe air pollution due to rapid urbanization and industrialization, especially in the more populous areas. According to the global burden of disease study 2017 [3], 1.24 million deaths were attributable to air pollution exposure, and air pollution resulted in 40.0 and 35.6% of DALYs for COPD and lower respiratory infections in China, respectively. To cope with the severe air pollution problem, the Chinese government has implemented several policies and regulations to improve the air quality and issued the Chinese National Ambient Air Quality Standard (GB3095–2012) in 2012, which limited the annual and 24-h mean concentrations of the major pollutants.

The most common pollutants are particulate matter (PM) with an aerodynamic diameter less than or equal to $2.5\,\mu m$ (PM $_{2.5}$) and with an aerodynamic diameter less than or equal to $10\,\mu m$ (PM $_{10}$). Longtime exposure to PM can cause multiple health problems that reduce human longevity [4]. Although numerous ecological studies have investigated the association between PM and outpatient or inpatient admissions of respiratory diseases, most of them were usually conducted for a relatively shorter study period [5–8] or a single hospital [7, 9] with a limited number of outcomes. Moreover, some studies [5, 8, 10] on the association of PM with respiratory diseases have produced inconsistent results.

Shanghai is the most populous megacity with a permanent population of 24.8 million in the seventh national census. It is also one of the heaviest polluted cities in China. The present study used time-series data for 12.5 years from all the medical insurance designated hospitals from Shanghai Healthcare Security Database for Urban Employees and Urban and Rural Residents. The database covers more than 95% of the adult population in Shanghai according to statistics released by the National Healthcare Security Administration, resulting in having a large number of hospitalizations and therefore providing good statistical power to examine the respiratory disease subcategories. We have investigated the association of short-term exposure to PM_{2.5} and PM₁₀ with total respiratory diseases and subcategories (COPD, asthma, and pneumonia) in Shanghai.

Methods

Respiratory disease data

Daily hospital admissions data were from the Shanghai Medical Insurance System for the period January 1, 2008 to July 31, 2020. The clinical diagnostic criteria for

respiratory diseases were from the International Classification of Diseases, 10th Revision as J00 to J99. This included COPD (J40–J44), asthma (J45), and pneumonia (J12–J18). Patients' basic information included gender and age (<45 years, 45–64 years, 65–74 years, and \geq 75 years). Prior to data collection, this study was approved by the Ethics Committee of the School of Public Health, Fudan University.

Air pollutants and meteorological data

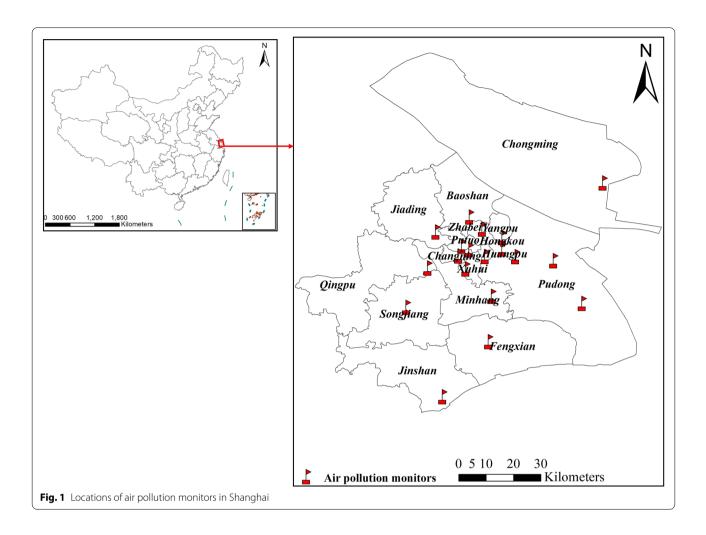
We collected six of the most common air pollutants from the Shanghai Municipal Bureau of Ecological Environment (https://sthj.sh.gov.cn/), namely PM_{2.5}, PM₁₀, nitrogen dioxide (NO₂), sulfur dioxide (SO₂), ozone (O₃), and carbon monoxide (CO). Of these, the daily concentrations of PM₁₀, NO₂, and SO₂ were available from January 1, 2008 to July 31, 2020. While, daily PM_{2.5}, O₃, and CO concentrations were only available after the establishment of the ground monitoring network after January 1, 2013. There are 19 environmental monitoring stations in Shanghai (Fig. 1). The daily concentration of air pollutants was simply an arithmetic mean measure across all the monitoring stations, as in most time-series studies. Meteorological data (mean temperature and relative humidity) in Shanghai were from the National Meteorological Information Center (http://data.cma.cn/).

Statistical analysis

A generalized additive model (GAM) [11] with a quasi-Poisson distribution was adopted to analyze the impact of $PM_{2.5}$ and PM_{10} on daily hospital admissions of respiratory diseases. The effect of different time lags was examined including six single-day lags: (i) lag 0, the present day; (ii) lag 1, the previous day; (iii) lag 2, the day before lag 1; (iv) lag 3, the day before lag 2; (v) lag 4, the day before lag 3; (vi) lag 5, the day before lag 4, and three moving average exposure lags: (i) lag 01, the 2-day moving average of the present and previous day; (ii) lag 02, the 3-day moving average of the present and previous 2 days; (iii) lag 03, the 4-day moving average of the present and previous 3 days.

Based on previous studies [12, 13], the following covariates were used: (*i*) a natural spline function of calendar time with 7 degrees of freedom per year (to exclude unmeasured time trends); (*ii*) a natural spline function with 6 degrees of freedom for present-day mean temperature and 3 degrees of freedom for present-day relative humidity (to control for the nonlinear confounding effects of weather conditions); and (*iii*) indicator variables for day of the week (DOW) and public holidays (PH). Therefore, the main model was:

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 $Log\left(E\left(Y_{t}\right)\right)=\alpha+\beta\times Z_{t}+ns\left(time,df=7/year\right)+ns\left(temperature,df=6\right)+ns\left(relative\ humidity,df=3\right)+as. factor(DOW)+as. factor(PH)$

where $E(Y_t)$ is the estimated daily hospital admissions for respiratory diseases, Z_t is the PM concentration on day t, β is the regression coefficient for Z_t , and α is the intercept. All results were presented as percentage change with a 95% confidence interval (CI) for each $10 \, \mu \text{g/m}^3$ increase of PM. Percentage change was calculated using the following formula [14]:

Percentage Change =
$$[\exp(\beta \times 10) - 1] \times 100$$

where β is the regression coefficient of PM from the GAM model.

Effect modification by individual characteristics was investigated using stratified analyses. The subgroup variables included gender (male and female), age (<45, 45-, 65- and 75- years), and season (cold: November, December, January, February, March and warm: April, May, June, July, August, September,

October). The subgroup differences were tested using a Z statistic [15]:

$$Z = (\beta_1 - \beta_2) / \sqrt{SE_1^2 - SE_2^2}$$

where β_1 and β_2 are the effect estimates of the two subgroups, SE_1 and SE_2 are their corresponding standard errors.

A cubic spline smoothing method was used to evaluate the exposure-response relationship between ${\rm PM}_{2.5}$ and ${\rm PM}_{10}$ concentrations and hospital admissions for respiratory diseases.

The attributable fraction (AF) and attributable number (AN) of hospital admissions due to PM exposure were estimated using the following formula based on a previous study [14]:

$$AF = 1 - \frac{1}{\exp\left[\beta \times (C - C_0)\right]}$$

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Table 1 Characteristics of hospital admissions for respiratory diseases in Shanghai, China from January 1, 2008 to July 31, 2020

Characteristics	n (%)		
Admissions of total respiratory diseases	1,960,361 (100.00)		
COPD	665,541 (33.95)		
Asthma	33,329 (1.70)		
Pneumonia	455,718 (23.25)		
Gender			
Male	1,091,645 (55.69)		
Female	868,714 (44.31)		
Missing	2 (0.00)		
Age groups, years			
<45	201,952 (10.30)		
45-	410,273 (20.93)		
65-	370,846 (18.92)		
75–	977,289 (49.85)		
Missing	1 (0.00)		
Season			
Cold	885,737 (45.18)		
Warm	1,074,624 (54.82)		

$$AN = AF \times N$$

where AF is the daily attributable fraction, AN is the daily attributable number of hospital admissions due to PM, N is the daily hospital admissions, β is the regression coefficient (from the main model above), C is the daily PM concentration, and C_0 is the reference PM concentration. The reference concentrations were from the air quality guidelines of the WHO: 24-h mean: $15\,\mu\text{g/m}^3$ for PM $_{2.5}$ and $45\,\mu\text{g/m}^3$ for PM $_{10}$. The total attributable number was estimated by summing daily AN.

We also performed three sensitivity analyses to determine the robustness of the results. First, a two-pollutant model ($PM_{2.5}$, PM_{10} with O_3 , SO_2 , NO_2 , CO)

was fitted. Second, we changed the degrees of freedom for the calendar time from 5 to 9 per year. Third, we excluded the data from 2020 due to the coronavirus disease 2019 pandemic.

All data processing and statistical analyses were conducted using R software (version 3.6.1).

Results

Hospital admissions for respiratory diseases and meteorological variables

There were 1,960,361 hospital admissions for total respiratory diseases (665,541 for COPD, 455,718 for pneumonia, and 33,329 for asthma) in Shanghai from January 1, 2008 to July 31, 2020 included in our analysis (Table 1). Males accounted for 55.69% of the patients and 49.85% of the patients were older than 75 years old.

Analysis of the air pollutants indicated the daily mean concentrations were $43.71\,\mu\text{g/m}^3$ for $PM_{2.5},~66.12\,\mu\text{g/m}^3$ for $PM_{10},~45.94\,\mu\text{g/m}^3$ for $NO_2,~21.08\,\mu\text{g/m}^3$ for $SO_2,~98.59\,\mu\text{g/m}^3$ for $O_3,~$ and $0.72\,\text{mg/m}^3$ for CO (Table 2). Our analysis of the annual average concentrations indicated decreasing trends for $PM_{2.5}$ and PM_{10} from 2013 to 2020 (Table S I).

Correlation between air pollutants and meteorological variables

We calculated Spearman correlation coefficients to examine the relationships of air pollution variables and meteorological factors (Table S II). The results indicated that daily $PM_{2.5}$ and PM_{10} concentrations had positive correlations with NO $_2$ (PM $_{2.5}$: $r_{\rm s}\!=\!0.721$, $P\!<\!0.001$; PM_{10} : $r_{\rm s}\!=\!0.685$, $P\!<\!0.001$), SO $_2$ (PM $_{2.5}$: $r_{\rm s}\!=\!0.729$, $P\!<\!0.001$; PM $_{10}$: $r_{\rm s}\!=\!0.655$, $P\!<\!0.001$), and CO (PM $_{2.5}$: $r_{\rm s}\!=\!0.879$, $P\!<\!0.001$; PM $_{10}$: $r_{\rm s}\!=\!0.759$,

Table 2 Summary statistics of air pollutants and meteorological variables in Shanghai, China from January 1, 2008 to July 31, 2020

Variables	$Mean \pm SD$	Minimum	P25	Median	P75	Maximum
Air pollutants						
PM _{2.5} , μg/m ^{3 *}	43.71 ± 31.18	5.00	22.00	36.00	56.00	255.00
PM_{10} , $\mu g/m^3$	66.12 ± 44.07	6.00	37.28	54.00	82.00	599.29
NO ₂ , μg/m ³	45.94 ± 20.31	5.00	31.00	43.00	57.00	141.65
SO ₂ , μg/m³	21.08 ± 17.39	3.00	9.00	15.00	26.83	146.78
O ₃ , μg/m ^{3 *}	98.59 ± 41.78	10.00	69.00	92.00	120.00	269.00
CO, mg/m ³ *	0.72 ± 0.26	0.30	0.52	0.68	0.80	2.28
Meteorological factors						
Temperature, °C	17.33 ± 8.86	-6.20	9.50	18.30	24.60	35.70
Relative humidity, %	71.50 ± 12.77	23.00	63.00	72.00	81.00	100.00

Note: * 2013.1.1–2020.7.31, P25 25th percentile, P75 75th percentile, SD standard deviation

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 $P\!<\!0.001).$ There were inverse correlations of $PM_{2.5}$ and PM_{10} concentrations with temperature ($PM_{2.5}$: $r_s\!=\!-0.311,\,P\!<\!0.001;\,PM_{10}$: $r_s\!=\!-0.229,\,P\!<\!0.001)$ and relative humidity ($PM_{2.5}$: $r_s\!=\!-0.142,\,P\!<\!0.001;\,PM_{10}$: $r_s\!=\!-0.365,\,P\!<\!0.001).$

Effect estimates between PM and respiratory diseases

We examined the effects of $PM_{2.5}$ and PM_{10} on hospital admissions for respiratory diseases using different lag times (Figs. 2 and 3). Overall, a $10\,\mu\text{g/m}^3$ increase of each PM was associated with increased total respiratory

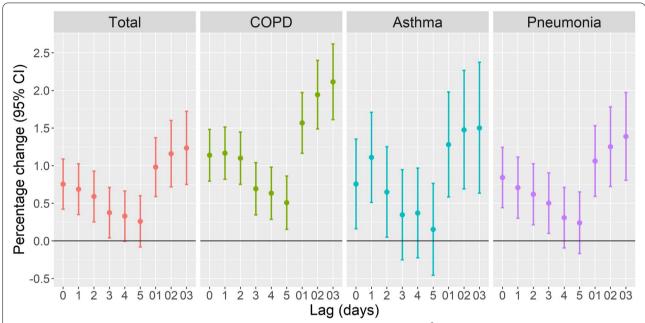


Fig. 2 Percentage change (95% CI) in hospital admissions for respiratory diseases for each $10 \,\mu\text{g/m}^3$ increase in the level of PM_{2.5} in Shanghai from 2013 to 2020 and effect of lag time. All models were adjusted for public holidays, day of the week, and calendar day

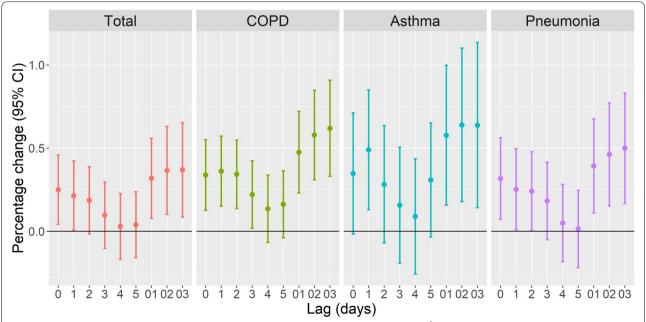


Fig. 3 Percentage change (95% CI) in hospital admissions for respiratory diseases for each $10 \,\mu\text{g/m}^3$ increase in the level of PM₁₀ in Shanghai from 2008 to 2020 and effect of lag time. All models were adjusted for public holidays, day of the week, and calendar day

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diseases at lag 0 (PM $_{2.5}$: 0.755%; 95% CI: 0.422, 1.089%; PM $_{10}$: 0.250%; 95% CI: 0.042, 0.459%). These associations decreased as the lag time increased to lag 5, but were increasingly greater for moving average exposure lags of lag01, lag02, and lag03. Analysis of each cause-specific respiratory disease indicated similar trends as total respiratory diseases. The percentage change seemed to be greater for COPD and asthma at lag 1. PM $_{2.5}$ had a greater effect than PM $_{10}$ for all lag times.

Stratified analyses by gender, age, and season

Stratification by gender indicated the associations of $PM_{2.5}$ and PM_{10} were similar in males and females (Table 3). Stratification by age indicated the associations of $PM_{2.5}$ with total respiratory diseases, asthma, and pneumonia were more pronounced in patients more than 45 years old. For patients younger than 45 years, there were no significant associations of PM with total respiratory diseases or with individual diseases, except for $PM_{2.5}$ and COPD. Stratification by season indicated the associations were more evident during the cold season, and that there were no significant associations of PM with total respiratory diseases or individual diseases during the warm season. We also analyzed these data using 9 different lag times (Figs. S I – Figs. S VI).

Exposure-response relationships

Exposure-response curves indicated that the percentage change of all types of respiratory diseases increased as PM concentrations increased (Figs. 4 and 5); these relationships were linear at low PM concentrations $(0 \sim 100 \, \mu g/m^3)$, but PM concentrations above $100 \, \mu g/m^3$ had less of an effect. Among the cause-specific respiratory diseases, $PM_{2.5}$ level had a nearly linear association with COPD up to $160 \, \mu g/m^3$, PM_{10} level had a nearly linear relationship with asthma up to $200 \, \mu g/m^3$, and asthma appeared to increase exponentially with $PM_{2.5}$ level up to $160 \, \mu g/m^3$.

Population attributable fraction (PAF)

We also assessed the excess number of respiratory diseases due to $PM_{2.5}$ and PM_{10} levels that exceeded the limits established by the WHO (Table S III - Table S XV). From 2013 to 2020, the overall total excess number of respiratory disease admissions was 21,678 (95% CI: 21,243, 22,112) due to $PM_{2.5}$ and 5865 (95% CI: 5587, 6142) due to PM_{10} . There were also trends of decreasing annual admissions for respiratory diseases from 2013 to 2020.

Sensitivity analyses

We initially performed a sensitivity analysis by fitting a two-pollutant model (Table S XVI). The results indicated

that the associations of $PM_{2.5}$ and PM_{10} with total respiratory diseases and individual diseases remained statistically significant after adjusting for other pollutants, although these effect estimates were not significant after adjusting for NO_2 . Our second sensitivity analysis indicated almost no change of the effect estimates after changing the degrees of freedom for calendar time (Table S XVII). When excluding the data from 2020, the effect estimates were similar to the main analysis (Table S XVIII).

Discussion

China is one of the most polluted countries in the world due to the rapid industrialization and urbanization [16]. In 2013, China experienced a severe air pollution event in most parts of the country, an event that was of great concern to the government. China's State Council subsequently issued the Air Pollution Prevention and Control Plan in 2013, and strengthening of environmental protections has greatly reduced pollution in China by PM_{2.5}. In response, the annual PM concentrations in Shanghai decreased markedly from 2013 to 2020. In 2019, the average $PM_{2.5}$ concentration (35.17 $\mu g/m^3$) was close to the national secondary standard limit of the Chinese National Ambient Air Quality Standard (35 µg/ m³). However, the concentrations were still significantly higher than the Air Quality Guidelines 2021 for PM_{2.5} $(5 \,\mu g/m^3)$. The health risks from exposure to PM should not be ignored.

Main findings and interpretation

The present ecological study found that short-term exposure to PM was significantly correlated with an increased risk of hospital admissions for total respiratory diseases and cause-specific respiratory diseases. Several previous studies also suggested an association of PM with respiratory diseases. For example, three studies [8, 17, 18] reported an association between PM and COPD and two other studies reported that short-term and longterm exposure to PM_{2.5} and PM₁₀ increased the risk of COPD [19, 20]. Three other studies [21-23] examined the influence of daily PM concentrations on the number of asthma-related admissions. Li et al. [24] found a significant reduction of asthma outpatient visits during the 2008 Olympic and Paralympic Games in Beijing, which they attributed to control measures that were implemented to improve air quality. A recent national time-series study in China from 2014 to 2017 showed a short-term positive association between ambient PM concentration and hospital admissions for pneumonia in Chinese adults [25].

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Table 3 Percentage change with 95% confidence interval for hospital admissions of respiratory diseases for a 10 μg/m3 increase in particulate matter, by gender, age groups, and season

Respiratory diseases	PM _{2.5}		PM ₁₀		
	Percentage change (95%CI)	P value	Percentage change (95%CI)	P value	
Total	0.755 (0.422,1.089)		0.250 (0.042,0.459)		
Gender					
Male	0.768 (0.439,1.098)	_	0.255 (0.052,0.457)	_	
Female	0.740 (0.390,1.092)	0.909	0.244 (0.018,0.470)	0.943	
Age groups, years					
<45	-0.019(-0.418,0.382)	_	0.006(-0.243,0.255)	_	
45-	0.701 (0.349,1.055)	0.008	0.338 (0.129,0.549)	0.046	
65-	0.865 (0.463,1.268)	0.002	0.193(-0.071,0.457)	0.312	
>=75	0.919 (0.567,1.273)	0.001	0.284 (0.058,0.510)	0.105	
Season					
Cold	0.652 (0.193,1.114)	_	0.296(-0.010,0.603)	_	
Warm	0.007(-0.711,0.731)	0.803	0.182(-0.173,0.539)	0.643	
COPD	1.167 (0.820,1.515)		0.361 (0.151,0.572)		
Gender					
Male	1.192 (0.840,1.544)	_	0.390 (0.182,0.599)	_	
Female	1.126 (0.747,1.507)	0.763	0.314 (0.079,0.550)	0.636	
Age groups, years					
<45	1.762 (0.769,2.765)	_	0.607(-0.038,1.257)	_	
45-	1.027 (0.546,1.510)	0.193	0.326 (0.052,0.601)	0.433	
65–	1.255 (0.827,1.685)	0.359	0.428 (0.174,0.683)	0.614	
>=75	1.157 (0.793, 1.523)	0.263	0.342 (0.120,0.565)	0.448	
Season	, , , , , , , , , , , , , , , , , , , ,		(,,		
Cold	1.170 (0.688,1.655)	_	0.387 (0.077, 0.699)	_	
Warm	0.336 (-0.407,1.084)	0.066	0.251 (— 0.101,0.604)	0.571	
Asthma	1.110 (0.513,1.710)		0.490 (0.131,0.850)		
Gender	, , , , , , , , , , , , , , , , , , , ,		,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,		
Male	0.915 (0.147,1.690)	_	0.356(-0.112,0.825)	_	
Female	1.278 (0.527,2.034)	0.509	0.607 (0.164,1.053)	0.452	
Age groups, years	, , , , , , , , , , , , , , , , , , , ,		,,,		
<45	-1.176(-2.857,0.534)	_	0.098(-0.851,1.056)	_	
45-	0.978 (0.165,1.799)	0.026	0.440(-0.019,0.901)	0.527	
65–	1.358 (0.348,2.378)	0.012	0.360(-0.330,1.055)	0.663	
>=75	2.161 (1.112,3.221)	0.001	0.935 (0.290,1.583)	0.155	
Season	,		,,		
Cold	1.186 (0.445,1.934)	_	0.534 (0.058,1.012)	_	
Warm	-0.271(-1.500,0.974)	0.049	-0.040(- 0.670,0.594)	0.179	
Pneumonia	0.842 (0.442,1.244)		0.317 (0.072,0.562)		
Gender	, , ,		(, ,		
Male	0.836 (0.432,1.243)	_	0.356 (0.113,0.602)	_	
Female	0.850 (0.417,1.286)	0.963	0.277 (0.006,0.549)	0.672	
Age groups, years	,,				
<45	-0.178(-0.832,0.480)		-0.036(-0.437,0.367)	_	
45-	0.966 (0.482,1.453)	0.006	0.477 (0.190,0.765)	0.042	
65-	1.129 (0.613,1.648)	0.002	0.296(-0.048,0.639)	0.219	
>=75	0.894 (0.482,1.308)	0.002	0.318 (0.058,0.580)	0.148	
Season				010	
Cold	0.945 (0.422,1.470)	_	0.547 (0.201,0.895)	_	
Warm	-0.312(-1.129,0.512)	0.012	-0.002(-0.410,0.409)	0.045	

 $\textit{Note:} \ Particulate \ matter \ concentration \ is \ lag\ 0 \ for\ total\ respiratory\ diseases\ and\ pneumonia; \ lag\ 1 \ for\ asthma\ and\ COPD$

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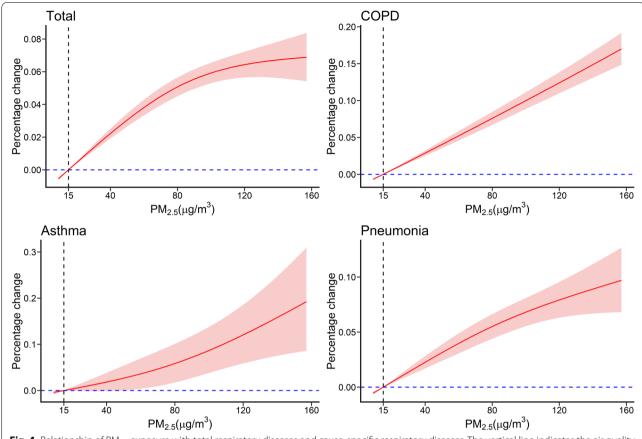


Fig. 4 Relationship of PM_{2.5} exposure with total respiratory diseases and cause-specific respiratory diseases. The vertical line indicates the air quality standard of the WHO for daily PM_{2.5} concentration (15 μ g/m³)

In line with previous studies [25, 26], we found that $PM_{2.5}$ had a greater impact on respiratory diseases than PM_{10} at all lag structures. Compared to PM_{10} , $PM_{2.5}$ adsorbs toxic substances and heavy metals more readily due to its larger relative surface area, it remains suspended in the atmosphere for longer periods, and it enters the terminal alveoli and even the bloodstream more easily [25].

Our age-stratified analysis found a significantly greater effect of PM on total respiratory diseases, asthma, and pneumonia in patients aged 45 years or more. The effect was especially strong in those aged 65 years or above, in accordance with previous studies [25–27]. Aged people may be more vulnerable to $PM_{2.5}$ due to their weaker immune responses [28, 29]. In addition, older people have a higher prevalence of pre-existing respiratory diseases [5].

We found a significant association of PM with respiratory diseases during the cold season, but not during the warm season, in line with previous studies of PM and respiratory diseases [30, 31]. There are several possible explanations for this finding. First, the $PM_{2.5}$ levels due

to heavy pollution were greater during the cold season than the warm season ($55.42\,\mu\text{g/m}^3$ vs. $35.41\,\mu\text{g/m}^3$). Second, low temperatures prolong the survival of respiratory viruses in the environment [32]. Third, inhalation of cold air can inhibit the mucociliary clearance and thereby promote the spread of viruses in the respiratory tract [33].

Our analysis of exposure-response relationships indicated a linear association between PM and total respiratory diseases at lower concentrations, but a progressively weaker effect of PM on respiratory diseases at concentrations above 100 µg/m³. These results are similar to those of a time-series study of 245,442 respiratory visits in Nanjing, China that examined the association of PM with respiratory system diseases [10]. Tian et al. [25] also reported a nonlinear relationship between PM_{2.5} level and pneumonia, with a weaker effect at higher concentrations. This nonlinear relationship may be because people avoid spending time outside or wear a dust mask when outside when the air is heavily polluted. The exposureresponse curves suggested no discernible thresholds of PM. Exposure to PM_{2.5} and PM₁₀ even at concentrations below the current regulatory limits is associated with

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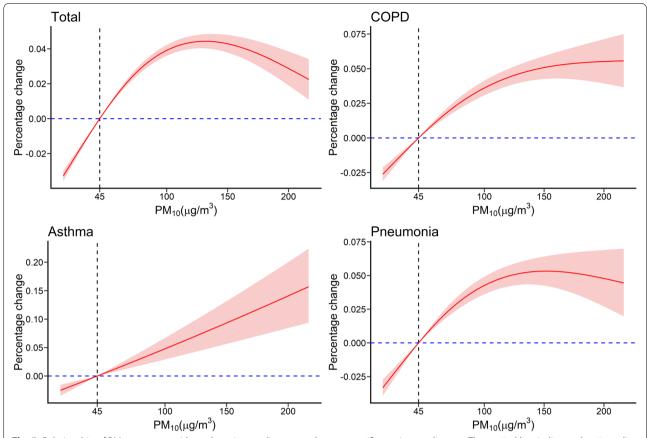


Fig. 5 Relationship of PM_{10} exposure with total respiratory diseases and cause-specific respiratory diseases. The vertical line indicates the air quality standard of the WHO for daily PM_{10} concentration (45 μ g/m³)

increases in daily admissions, suggesting that tightening the current air quality guidelines of PM for greater public health benefit is necessary.

Our two-pollutant model indicated the associations between PM and respiratory diseases remained positive, but not significant after adjusting for NO₂. Previous studies reported similar results. For example, Tian et al. [25] reported no significant association between PM and pneumonia after controlling for NO₂. Liu et al. [34] suggested that the effect of PM on respiratory admissions decreased dramatically and was no longer significant after controlling for NO₂. Tsai et al. [35] found that the adverse effect of PM_{2.5} was not statistically significant after adjusting for NO₂ or CO. It is possible that a high correlation among pollutants weakened the effect estimation. Therefore, the independent effect of PM on respiratory diseases should be examined.

Potential mechanism

The potential biological mechanism linking the exposure to PM and hospital admissions for respiratory diseases are not fully understood, but there are several plausible mechanisms. First, exposure to PM may lead to oxidative stress. For example, $PM_{2.5}$ can easily adsorb organic pollutants, such as polycyclic aromatic hydrocarbons and heavy metals, which promote the production of reactive oxygen species in lung cells, consume antioxidant substances, trigger oxidative stress reactions, and induce lung inflammation [36]. A meta-analysis [37] of epidemiology studies suggested that short-term exposure to $PM_{2.5}$ was associated with an increased level of malondialdehyde, a biomarker of oxidative stress. Second, $PM_{2.5}$ induction of a local or systematic inflammatory reaction is the major pathologic basis for the onset and progression of several related diseases [38]. Evidence from a meta-analysis [39] also showed that PM exposure increased the levels of fibrinogen and TNF-α.

Limitations

Compared with previous studies, we used a larger sample size and examined a longer time period, resulting in greater statistical power. However, this study had several limitations. First, our study was a one-city observational study, and multi-city studies are necessary to validate our Peng et al. Environmental Health (2022) 21:12 Page 10 of 12

conclusions. Second, due to unavailability of residential location where each patient lived, we used average concentrations of daily air pollutants from fixed environmental monitoring stations as an indicator of exposure. This might result in measurement error, which tends to be nondifferential and underestimate the effect of the PM effects [40]. Previous studies used satellite-derived PM_{2.5} concentrations based on modeling techniques to represent individual exposure levels and to examine the potential effect on several diseases, including asthma [41] and poor birth outcome [42]. Third, potential confounders, including education level, occupation, socioeconomic status (e.g., family income), and individual behaviors (e.g., smoking status), were not available from the medical insurance system, making stratified analysis impossible. Finally, this study was a time-series ecological investigation of the associations of PM with health outcomes, and cannot make definitive conclusions regarding causality. Our results also could have been affected by the ecological fallacy.

Conclusion

The main result of this study is that short-term exposure to PM in Shanghai was significantly associated with hospital admissions for total respiratory diseases and cause-specific respiratory diseases — COPD, asthma, and pneumonia. These associations were especially stronger for patients more than 45 years old and during the cold season. PM level had a nearly linear relationship with total respiratory diseases at low concentrations (0 ~ 100 $\mu g/m^3$), but had a weaker as PM concentrations increased. Reducing atmospheric PM concentrations may reduce hospital admissions for respiratory diseases. These findings have important implications for policy-makers to take concrete actions to reduce atmospheric PM concentrations. Further studies conducted on nation-wide regions are required to validate our results.

Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s12940-021-00828-6.

Additional file 1: Table S I. The annual average concentrations of PMs in Shanghai. Table S II. Spearman correlation coefficients among air pollution variables and meteorological factors. Table S III. The attributable number of respiratory diseases admissions due to exceeding PM₂₅ concentrations. Table S IV. The attributable number of respiratory diseases admissions due to exceeding ${\rm PM}_{10}$ concentrations. **Table S V.** The attributable number of respiratory diseases admissions due to exceeding PM₂₅ concentrations in males. **Table S VI.** The attributable number of respiratory diseases admissions due to exceeding PM_{2.5} concentrations in females. Table S VII. The attributable number of respiratory diseases admissions due to exceeding PM₁₀ concentrations in males. Table S VIII. The attributable number of respiratory diseases admissions due to exceeding PM₁₀ concentrations in females. Table S III. The attributable number of respiratory diseases admissions due to exceeding $PM_{2.5}$ concentrations in < 45 years. Table S IX. The attributable number of respiratory diseases admissions due to exceeding PM_{2.5} concentrations in

 $45 \sim 64$ years. **Table S X.** The attributable number of respiratory diseases admissions due to exceeding $PM_{2.5}$ concentrations in 65 ~ 74 years. **Table S XI.** The attributable number of respiratory diseases admissions due to exceeding PM_{2.5} concentrations in \geq 75 years. **Table S XII.** The attributable number of respiratory diseases admissions due to exceeding PM₁₀ concentrations in < 45 years. Table S XIII. The attributable number of respiratory diseases admissions due to exceeding PM₁₀ concentrations in 45~64 years. **Table S XIV.** The attributable number of respiratory diseases admissions due to exceeding PM_{10} concentrations in 65 ~ 74 years. **Table S XV.** The attributable number of respiratory diseases admissions due to exceeding PM₁₀ concentrations in >75 years. **Table S XVI.** Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 µg/m³ increase in concentrations of PM_{2.5} and PM₁₀ using two-pollutant models. Table S XVII. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases associated with a 10 µg/m³ increase in concentrations of PM₂₅ and PM₁₀ through changing degrees of freedom for the calendar time. Table S XVIII. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases associated with a 10 µg/m3 increase in concentrations of PM₂₅ and PM₁₀ through excluding the data from 2020 due to the coronavirus disease 2019 pandemic. Fig. S I. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 µg/m³ increase in concentrations of PM₂₅ stratified by genders. All the models were adjusted by public holidays, DOW, and calendar day. Fig. S II. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 µg/m³ increase in concentrations of PM₁₀ stratified by genders. All the models were adjusted with public holidays, DOW, and calendar day. Fig. S III. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 μg/m³ increase in concentrations of PM_{2.5} stratified by age groups. All the models were adjusted with public holidays, DOW, and calendar day. Fig. S IV. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 µg/m³ increase in concentrations of PM₁₀ stratified by age groups. All the models were adjusted with public holidays, DOW, and calendar day. Fig. S V. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per $10 \,\mu g/m^3$ increase in concentrations of PM_{2.5} stratified by season. All the models were adjusted with public holidays, DOW, and calendar day. Fig. S VI. Percentage change with 95% confidence interval in hospital admissions for respiratory diseases per 10 µg/m³ increase in concentrations of $\ensuremath{\mathsf{PM}_{10}}$ stratified by season. All the models were adjusted with public holidays, DOW, and calendar day.

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Authors' contributions

Wenjia Peng: Conceptualization, Software, Formal analysis, Writing - Original Draft. Hao Li: Conceptualization, Data Curation, Software, formal analysis. Li Peng: Data Curation, Resources. Ying Wang: Conceptualization, Writing - Review & Editing, supervision. Weibing Wang: Conceptualization, Resources, Methodology, Writing - Review & Editing, Project administration, Funding acquisition. The author(s) read and approved the final manuscript.

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Availability of data and materials

The data that support the findings of this study are available from the corresponding author upon reasonable request.

Declarations

Ethics approval and consent to participate

Not applicable.

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Consent for publication

All authors have approved the manuscript for submission.

Competing interests

The authors declare that they have no competing interests.

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