

Research Article

A Longitudinal Analysis of the Association Between Long-Term Exposure to Air Pollution and Cognitive Function Among Adults Aged 45 and Older in China

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Abstract

Objectives: Evidence suggests long-term exposure to fine particulate matter air pollution (PM_{2.5}) is associated with a higher risk of cognitive impairment, especially among older adults. This study examines the relationship between PM_{2.5} exposure and cognitive function in China's aging population.

Methods: We used longitudinal data from the China Health and Retirement Longitudinal Study (2011–2015) linked with historical PM_{2.5} concentrations (2000–2015) from remotely sensed satellite data. Growth curve models were applied to estimate associations between PM_{2.5} exposure (measured in intensity, duration, and a joint variable of intensity with duration for cumulative exposure) and cognitive function.

Results: Relative to the lowest exposure group, exposure in the second group of PM_{2.5} intensity (35–50 µg/m³) is associated with poorer cognitive function, but higher levels of PM_{2.5} appear to be associated with better cognitive function, indicating a U-shaped association. Similar patterns are seen for fully adjusted models of PM_{2.5} duration: the second group (13–60 months) is associated with worse cognitive function than the first group (0–12 months), but coefficients are nonsignificant in longer duration groups. Joint analysis of PM_{2.5} intensity with duration suggests that duration may play a more detrimental role in cognitive function than intensity. However, we do not find a statistically significant association between PM_{2.5} exposure and the rate of cognitive decline.

Discussion: Our findings are mixed and suggest that some categories of higher and longer exposure to PM_{2.5} are associated with poorer cognitive function, while that exposures do not hasten cognitive decline. However, more work is necessary to disentangle PM_{2.5} exposure from individuals' background characteristics, particularly those jointly associated with cognitive function and urban living.

Keywords: Air pollution, Cognitive function, Cumulative exposure, Health disparities, PM_{2.5}

Cognitive impairment reduces healthy life expectancy, affects the quality of later life, and contributes substantially to health care burdens for individuals and societies (Clifford et al., 2016; Yao et al., 2022). Globally, 40%

of dementia cases are related to 12 main modifiable risk factors, which include ambient air pollution (Calderón-Garcidueas et al., 2012; Fu & Yung, 2020; Livingston et al., 2020; Peters et al., 2019; Wang et al., 2020). There

are emerging population-based studies of the association between fine particulate matter smaller than 2.5 micrometers (μm) aerodynamic diameter ($\text{PM}_{2.5}$) in outdoor air pollution and cognitive impairment and other neurological diseases (Ailshire & Clarke, 2015; Russ et al., 2019; Wang et al., 2020). $\text{PM}_{2.5}$ consists of complex and varying mixtures of particles suspended in the air. Once inhaled, these particles can lead to systemic inflammation and oxidative stress across the blood–brain barrier (Block & Calderón-Garcidueñas, 2009). As a result, $\text{PM}_{2.5}$ is a risk factor for declining brain function (Guxens & Sunyer, 2012).

China—one of the fastest-aging countries—experiences severe air pollution, where more than 50% of cities recorded $\text{PM}_{2.5}$ levels far exceeding the World Health Organization safe levels in 2013 (Lancet, 2014). Air pollution in China is a result of the rapid industrial expansion occurring since 1979, which caused increases in coal consumption, motor, and industrial dust (Xu et al., 2013). The lives of those born before or during the 1970s have coincided with a period of rapid economic development, so they have spent large parts of their adult life course exposed to hazardous air pollution. For urban dwellers aged 45 and older, the long-term damage of air pollution is likely to be substantial (Lu et al., 2020). The effects of air pollution on health outcomes may vary by the age of individuals at exposure, because both children and older individuals are especially susceptible to deleterious air pollution (Peled, 2011; Shier et al., 2019). Research shows that exposure to $\text{PM}_{2.5}$ is significantly associated with cognitive function in older adults (Ailshire & Clarke, 2015; Ailshire et al., 2017; Wang et al., 2020). Therefore, the aging and air pollution context in China is conducive to understanding the association between air pollution and cognitive function in more detail.

Although we have some knowledge about the association between exposure to severe air pollution and cognitive function, there remain some uncertainties. First, some studies, which have found that cognitive function is associated with air pollution exposure (Chen et al., 2017; Wang et al., 2020), have used average concentrations of air pollutants (mainly measured by the annual average) that do not completely capture effects of intensity and duration. Second, other studies use data that do not cover a long-term exposure window (Power et al., 2011; Sun & Gu, 2008). Thus, using multiple measures for exposure to air pollution over a long-term period (e.g., more than 10 years) is necessary to explore the association between time-integrated exposures and health risks.

In this study, we analyzed survey data from a large, prospective, nationally-representative cohort of Chinese adults ranging from 45 to 105 years of age, linked with 15 years of historical satellite data on $\text{PM}_{2.5}$ exposure. We used growth curve models (GCM) to study the associations between $\text{PM}_{2.5}$ exposure and cognitive function, comparing different ways of measuring $\text{PM}_{2.5}$ exposure: intensity, duration, and a measure that takes into consideration both exposure intensity and its duration, which we called “cumulative exposure.”

Method

Study Population

Data were from three waves of the China Health and Retirement Longitudinal Study (CHARLS 2011–2015), which is a nationally representative longitudinal survey of the middle-aged and older population of China, consisting of persons 45 years of age or older, as well as their spouses when possible. The CHARLS used computer-assisted in-person interviews to obtain samples through four-stage stratified sampling, with an overall response rate of 80.5% at the baseline. From June 2011 to March 2012, the CHARLS conducted a baseline survey that included assessments of the social, economic, and health circumstances of 17,705 respondents from 28 provinces, 150 cities/counties/districts, and 10,257 households (Zhao et al., 2014). Following this baseline survey, two follow-up surveys were conducted in 2013 and 2015.

In 2011, the baseline CHARLS sample size was 17,705. Between 2011 and 2013, 3,130 respondents were lost due to death ($n = 441$) or nonspecified reasons ($n = 2,509$). In 2013, the CHARLS, to maintain age representation, added a refreshment sample of individuals who entered age eligibility of being 45 years and older between waves 1 and 2. If the baseline respondent shared a household with someone aged 40 and 44, he or she was reserved for a refreshment sample for future survey rounds. In wave 2, respondents who were aged 43–44 in wave 1 (plus their spouses) were added from the refreshment sample, the same for wave 3 in 2015, out of those aged 41–42 in wave 1. In wave 2 of the CHARLS 2013 ($n = 18,064$), 4,029 refreshment respondents were selected from household members of the original wave 1 respondent. Using the same strategy, the third wave of CHARLS in 2015 included 3,275 new individuals. Between 2013 and 2015, 689 respondents died, and by 2015, 809 respondents, interviewed in 2011 but missing in 2013, returned ($n = 21,100$).

Figure 1 describes how we selected the analytical samples. First, we excluded 1,719 observations that were under the age of 45. The CHARLS did not collect detailed residential histories, so if respondents moved, it was difficult to reconstruct their air pollution exposure history. However, respondents self-reported the date they moved to their current residence, allowing us to exclude those who reported moving residence from 2000 to baseline or during the panel. Only 246 respondents (0.45% of the total samples) changed their residence in 2011, and 427 (0.79% of the total) and 913 (1.7% of the total) respondents moved in 2013 and 2015. We used listwise deletion for missingness on cognitive function (7,430 deleted) and other predictors (6,270 deleted). In order to make the exposure window consistent (see Supplementary Table S12), we have restricted the analytical sample to those individuals who entered the survey in 2011 (2013 and 2015 entrants were removed). Finally, our statistical analysis contains 29,484 observations (12,481 respondents) from 125 cities.

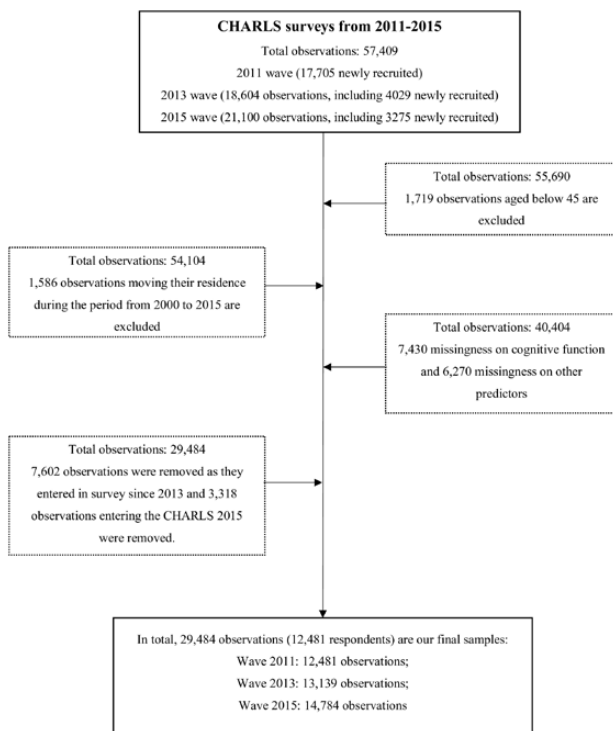


Figure 1. Flowchart of study inclusion criteria. *Notes:* CHARLS = China Health and Retirement Longitudinal Study.

As a robustness check, we employed multiple imputation (MI) using chained equations, including all predictors that were in the analytical model (Carpenter & Kenward, 2012; van Ginkel et al., 2020; White et al., 2011; full details in Supplementary Material and Supplementary Tables S1 and S2). After creating multiple imputed data sets, the analysis was rerun, and the resulting models were combined using Rubin’s rules (Rubin, 1976), which take into account variation both within and between data sets.

Outcome: Cognitive Function

Cognitive function was measured in the CHARLS by the modified Telephone Interview for Cognitive Status Survey, which includes orientation (recalling the date [year, month day], day of the week, and season of the year, 0–5 points), numeric ability (the serial subtraction of seven from 100, up to five times, 0–5 points), word recall (immediately repeating in any order 10 Chinese nouns, 0–10 points), and recalling the same list of words 4 min later, 0–10 points), and visuospatial ability (drawing a pentagon, 0–1 points). Following previous studies (Crimmins et al., 2011; Huang & Maurer, 2019; Kesavayuth et al., 2018), we used only variables reflective of fluid cognitive function (immediate and delayed word recall and serial 7s), which are considered more indicative of neurophysiological health rather than highly related to education and other sociocultural factors (Ghisletta et al., 2012). The range of fluid cognitive function in this study is from 0 to 25, with higher scores indicating better cognitive function.

Air Pollution: PM_{2.5}

We used a data source derived from satellite aerosol remote sensing data products such as the Moderate Resolution Imaging Spectroradiometer (MODIS) Collection six level 2 aerosol products at 10 km resolution from Aqua and Terra satellites (<http://adsweb.nascom.nasa.gov/>) as well as other information. This PM_{2.5} data set was generated by applying machine learning algorithms to predict historical PM_{2.5} concentrations with satellite-retrieved aerosol optical depth from MODIS, gridded meteorological parameters, as well as land use information in China as predictors (Xiao et al., 2018). Compared with ground monitoring data, satellite data with a broad spatial coverage (all of China), long-term records (from March 2000 to December 2015), and high spatial resolution (10 km) support the assessment of historical air pollution levels in developing regions (Xiao et al., 2018). To validate PM_{2.5} estimates from satellite data, the data set creators conducted the 10-fold cross validation (CV) to evaluate the prediction performance, with a CV R² of 0.79, which was significantly higher than previous methods (Xiao et al., 2018). Note that although these PM_{2.5} data were detailed in each (10 km × 10 km) grid cell every month starting in March 2000, the CHARLS did not provide the exact residence address for each respondent due to concerns about identifiability. Thus, we aggregated PM_{2.5} concentrations from satellite data at the city level because the city information from the primary sampling units (PSU) of CHARLS is the smallest spatial unit. In the CHARLS, there were 125 cities selected as the PSU, which were distributed across 28 of 34 provinces, covering 95% of the population in China.

In our study, the period of exposure to PM_{2.5} is from March of 2000 to the month preceding cognitive assessment in each wave of the CHARLS. Supplementary Figure S1 shows the period of exposure for the CHARLS samples in each wave. Considering the evidence of the nonlinear association between air pollution and cognitive function (Ailshire & Crimmins, 2014; Power et al., 2011), we analyzed PM_{2.5} exposure as a categorical variable. Following previous studies related to exposure to risk factors (De Voucht et al., 2015; Pope et al., 2011), there were three subindices for exposure to PM_{2.5} in this study.

The first one was the average concentrations of PM_{2.5} as the “PM_{2.5} intensity.” As the National Ambient Air Quality Standard for annual mean PM_{2.5} is 35 µg/m³ (level 1) and 75 µg/m³ (level 2) in China (Cao et al., 2013) and the median of the average PM_{2.5} concentration during the period between 2000 and 2015 was 50 µg/m³, we categorized PM_{2.5} intensity into four groups using the following cut-points: 1 (0–35 µg/m³), 2 (36–50 µg/m³), 3 (51–75 µg/m³), and 4 (76+ µg/m³). The second measure was the duration of exposure, measured as months over a fixed PM_{2.5} concentration threshold. Evidence on the effects of PM_{2.5} in China suggests that exposure to concentration over 50 µg/m³ has an adverse impact on cognitive function (Wang et al., 2020). Thus, this study used 50 µg/m³ (which is also the median of

PM_{2.5} intensity) as the threshold to establish the duration of exposure. Following previous studies (Ranft et al., 2009; Tallon et al., 2017; Zeng et al., 2010), “PM_{2.5} duration” was categorized into four groups: 1 (0–12 months), 2 (13–60 months), 3 (61–120 months), and 4 (121+ months). For the third measure, “cumulative exposure,” we interacted intensity with duration, establishing a joint variable with 16 categories as the cumulative PM_{2.5} indicator. This takes into consideration that regions with higher levels of pollution intensity were likely to have had those higher levels for longer durations, which would have caused a problem of multicollinearity if assessed simultaneously in the same model (details shown in [Supplementary Table S11](#)).

Covariates: Demographic, Socioeconomic Status (SES), and Regional Factors

In this study, we categorized covariates into three groups. The first is demographic information, including sex, age, and partnership status. Women may have a higher risk of cognitive decline associated with increased PM_{2.5} exposure than men (Kim et al., 2019). To account for the well-established curvilinear association between cognitive function and age (Ballesteros et al., 2009), we included both age and a quadratic function for age in our analysis. Living with a partner might have a protective effect against cognitive impairment, especially in later life (Håkansson et al., 2009), so we include a time variant control for partnership status: single (separate, divorced, widowed, or never married) or partnered (married or living with a partner).

The second set of covariates measures socioeconomic status (SES): education at baseline, primary occupational attainment (time-invariant), and time-variant household expenditure. Educational attainment was the most substantial predictor of cognitive function in China as elsewhere (Cagney & Lauderdale, 2002; Huang & Zhou, 2013). Occupational attainment is also associated with cognition. For example, civil servants and managers have a better cognitive function, net of potential confounders (Myung et al., 2016). In this study, occupational attainment was measured by the main job reported during respondents' occupational history and includes three categories: agricultural, nonagricultural, and managerial occupations. Wealth has been found to be associated with lower cognitive function, so we controlled household expenditure (logged) as an operationalization of wealth (Cagney & Lauderdale, 2002).

HuKou is a household registration system in China that has two categories: rural and urban. People usually remain as the same HuKou as their parents, as once HuKou is registered, it is difficult to change even if people move (Hou et al., 2019). Particularly before China enacted the reform and opening-up policy in 1978, HuKou stipulated whether people could work in industrial sectors, have more educational opportunities, and even have better access to medical insurance (Walder, 1995). As such, HuKou is strongly

related to individual SES in addition to residential locations (Hou et al., 2019).

Rapid urbanization and industrialization are associated with improvements in population health (e.g., improved health care system and more health facilities), alongside high levels of air pollution (Gong et al., 2012). In this study, we included annual regional gross domestic product (GDP) at the city level (logged) to reflect the urbanization and industrialization of cities to adjust for this potential confounding factor (Sun & Gu, 2008).

Analysis Strategies

This study analyzed whether changes in individual cognitive function during the CHARLS 2011–2015 are related to cumulative PM_{2.5} exposure. We employed GCM to examine the relationship between exposure to PM_{2.5} between 2000–2015 and the trajectory of cognitive decline between 2011–2015. An important advantage of GCM is the ability to model the trajectories of individuals over time and distinguish within-individual from between-individual heterogeneity in estimating cognitive changes shaped by other variables. GCM is a special case of random-coefficient models that can take a variety of shapes of growth trajectories, and it is the time coefficient (here, age) that varies randomly between participants (Rabe-Hesketh & Skrondal, 2012). In this study, we used three waves of longitudinal data across 4 years of data collection.

We estimated three separate sets of models, each with different operationalisations of air pollution exposure. The first set of models includes intensity, as the average concentration of PM_{2.5} during the period of exposure. The second set includes duration measured as the number of months where PM_{2.5} concentration is over the threshold of 50 µg/m³ during the period of exposure. We tested different thresholds for PM_{2.5} duration as robustness checks ([Supplementary Material](#)). The third set includes a joint variable of intensity with duration. To investigate the rate of cognitive decline, we used the interaction term between age, age squared, and PM_{2.5} exposure to examine the association between PM_{2.5} exposure and cognitive trajectories. This is to determine whether higher exposure leads to a faster rate of cognitive decline or just overall lower cognitive function. To examine heterogeneity in the associations of PM_{2.5} exposure among different groups, we also stratified the associations between PM_{2.5} exposure and cognitive function by HuKou status.

We conducted various robustness checks. First, we ran the same models using the imputed data (through MI) to check consistency and verify that our results were not biased by missing data (shown in [Supplementary Tables S3–S5](#)). Second, we set up alternative specifications to check the sensitivity of the threshold value in PM_{2.5} exposure variable. For PM_{2.5} intensity, there were six categories using 10 µg/m³ as the interval from 35 to 75 µg/m³; for PM_{2.5} duration, taking 45 µg/m³ as another threshold, and then the

joint intensity-duration variable expanded to 24 categories. All of these analyses can be found in [Supplementary Tables S6–S8](#). Third, although the main analysis with PM_{2.5} categories establishes the curvilinear relationship between PM_{2.5} exposure and cognitive function trajectories, categorizing PM_{2.5} records sacrifices some details compared with using continuous PM_{2.5} concentrations. Therefore, we also estimated a model with a quadratic term of continuous exposure (in both intensity and duration) to examine the curvilinear dose–response curve ([Supplementary Tables S9 and S10](#)). Fourth, we have added a robustness check using the data that includes only individuals who participated in all three waves of CHARLS (there are 6,589 respondents, including 19,767 observations). This ensures all respondents have the same possibility of exposure ranging from 0 to 328 months (the total observation period) over PM_{2.5} intensity of 50 µg/m³ ([Supplementary Figure S6](#)).

Results

Our study population includes 12,481 respondents (29,484 observations) from three waves of the CHARLS 2011, 2013, and 2015. The average age of 2011 entrants was 59 years old, with some variations by PM_{2.5} exposure

groups; 49% were men and 76% of the population attained primary or higher education ([Table 1](#)). 76% of the study population had rural HuKou, 72% of the population worked in agricultural jobs, and 88% were partnered. The average cognitive function score was 10. Respondents with higher education (e.g., primary or higher), higher household expenditure (8.30), and in areas with higher GDP (10.55) had higher exposure to PM_{2.5} (especially exposed to PM_{2.5} over 76 µg/m³).

[Table 2](#) shows the results from the GCMs. Model 1 is the base model with PM_{2.5} intensity, age, age squared, and gender. The coefficients in Model 1 show that the associations between PM_{2.5} intensity and cognitive function are positive (higher PM_{2.5} is associated with better cognitive function). However, after education was included in Model 2 as a confounding variable, the association between PM_{2.5} intensity and cognitive function takes a U-shaped pattern, with both lower and higher intensity being associated with higher cognitive function, a finding to which we return below. In Model 3, which controls additionally for occupation, household expenditure, and partnership status, we found that compared with people exposed to the lowest PM_{2.5} intensity (0–35 µg/m³), those in the next group of higher exposure (36–50 µg/m³) had lower cognitive function

Table 1. Characteristics of 2011 Entrants in the CHARLS by PM_{2.5} Intensity

	Total	PM _{2.5} (µg/m ³) mean average monthly exposure between 2000 and baseline (mean [SD] or n [%])			
		1 (0–35 µg/m ³)	2 (36–50 µg/m ³)	3 (51–75 µg/m ³)	4 (76+ µg/m ³)
Cognitive score	9.97 (4.48)	9.75 (4.44)	9.88 (4.52)	9.92 (4.56)	10.10 (4.39)
Age	58.99 (9.47)	58.14 (9.49)	59.09 (9.48)	59.46 (9.58)	57.55 (8.43)
Gender (%)					
Men	6,069 (48.63)	955 (48.22)	1,697 (48.22)	2,918 (49.02)	499 (47.34)
Women	6,421 (51.37)	1,000 (51.15)	1,822 (51.78)	3,035 (50.98)	555 (52.66)
Education (%)					
No schooling	3,199 (25.63)	642 (32.84)	860 (24.44)	1,537 (25.82)	160 (15.18)
Primary	5,011 (40.15)	808 (41.33)	1,388 (39.44)	2,453 (41.21)	362 (34.35)
Middle	4,271 (34.22)	505 (25.83)	1,271 (36.12)	1,963 (32.97)	532 (50.47)
HuKou (%)					
Rural	9,705 (77.76)	1,694 (86.61)	2,596 (73.77)	4,582 (76.97)	833 (79.03)
Urban	2,776 (22.24)	261 (13.39)	923 (26.23)	1,371 (23.03)	221 (20.97)
Occupation (%)					
Agricultural	9,038 (72.41)	1,506 (77.03)	2,389 (67.89)	4,405 (74.00)	738 (70.02)
Nonagricultural	2,789 (22.35)	358 (18.31)	923 (26.23)	1,263 (21.22)	245 (23.24)
Managerial	654 (5.24)	91 (4.65)	207 (5.88)	285 (4.79)	71 (6.74)
Partnership status (%)					
Partnered	11,014 (88.25)	1,684 (86.14)	3,054 (86.79)	5,330 (89.53)	946 (89.75)
Single	1,467 (11.75)	271 (13.86)	465 (13.21)	623 (10.47)	108 (10.25)
Log household expenditure	8.16 (1.67)	8.18 (1.66)	8.15 (1.82)	8.14 (1.57)	8.30 (1.74)
Log GDP	10.29 (0.55)	10.19 (0.51)	10.21 (0.63)	10.33 (0.50)	10.55 (0.43)
Number of respondents (%)	12,481	1,955 (15.66)	3,519 (28.19)	5,953 (47.71)	1,054 (8.44)

Notes: All statistics are calculated after list wise deletion (see details in the “Method” section). The characteristics of a cognitive score, age, log household expenditure and log GDP are shown using mean (SD), and others are in N (%). CHARLS = China Health and Retirement Longitudinal Study; SD = standard deviation; GDP = gross domestic product.

Table 2. Associations Between PM_{2.5} Intensity and Cognitive Function

	Model 1: Base	Model 2: Model 1 + Education	Model 3: Model 2 + SES + Partnership	Model 4: Model 3 + GDP
PM _{2.5} intensity (ref: 0–35 µg/m ³)				
2 (36–50)	0.374*** (0.173 to 0.575)	–0.141 (–0.320 to 0.0373)	–0.252** (–0.428 to –0.0769)	–0.233** (–0.408 to –0.0577)
3 (51–75)	0.500*** (0.314 to 0.685)	0.0363 (–0.128 to 0.200)	–0.0291 (–0.190 to 0.132)	–0.0443 (–0.205 to 0.116)
4 (76+)	1.087*** (0.835 to 1.339)	0.199# (–0.0271 to 0.425)	0.197# (–0.0248 to 0.419)	0.118 (–0.105 to 0.340)
Age	0.127*** (0.0560 to 0.197)	0.249*** (0.185 to 0.313)	0.236*** (0.172 to 0.299)	0.234*** (0.171 to 0.297)
Age squared	–0.00231*** (–0.0029 to –0.0018)	–0.00275*** (–0.0033 to –0.0023)	–0.00269*** (–0.0032 to –0.0022)	–0.00269*** (–0.0032 to –0.0022)
Gender (ref: men)				
Women	–1.286*** (–1.415 to –1.157)	–0.0665 (–0.187 to 0.0540)	–0.116# (–0.235 to 0.00329)	–0.143* (–0.262 to –0.0235)
Education (ref: no-schooling)				
Primary		2.626*** (2.476 to 2.776)	2.414*** (2.266 to 2.563)	2.356*** (2.207 to 2.505)
Middle		5.053*** (4.887 to 5.219)	4.320*** (4.144 to 4.496)	4.247*** (4.071 to 4.424)
HuKou (ref: rural)				
Urban			1.165*** (1.010 to 1.320)	1.130*** (0.976 to 1.285)
Occupation (ref: agricultural)				
Nonagricultural			0.475*** (0.327 to 0.623)	0.440*** (0.292 to 0.588)
Managerial			0.294* (0.0619 to 0.525)	0.273* (0.0410 to 0.504)
Log household expenditure			0.0912*** (0.0668 to 0.116)	0.0876*** (0.0632 to 0.112)
Partnership (ref: partnered)				
Single			–0.349*** (–0.516 to –0.183)	–0.340*** (–0.507 to –0.174)
Log GDP				0.332*** (0.239 to 0.426)
Constant	10.98*** (8.785 to 13.17)	2.265* (0.262 to 4.267)	2.180* (0.181 to 4.179)	–1.060 (–3.254 to 1.135)
Random effects				
Within individual				
Change rate (age)	0.003***	0.005***	0.004***	0.004***
Intercept	2.931***	2.797***	2.660***	2.663***
Covariance	0.014	–0.008	–0.008	–0.009
Between individual				
Residuals	3.290***	3.051***	3.056***	3.056***
Log likelihood	–82,019.908	–80,455.387	–80,218.955	–80,194.873
Observations	29,484	29,484	29,484	29,484
Number of IDs	12,481	12,481	12,481	12,481

Notes: Cognitive function includes three components: immediate recall, delayed recall, and serial 7s, 0–25 points (see details in “Methods”). SES = socioeconomic status; GDP = gross domestic product.

*** $p < .001$, ** $p < .01$, * $p < .05$, # $p < .1$.

($\beta = -0.252$, $p < .01$); however, the fourth group (those with the highest levels of exposure) had a better cognitive function—an unexpected result. This could be caused by SES confounding (details in marginal plots in [Supplementary](#)

[Figure S2](#)). Model 4 (including logged GDP), indeed, supports the U-shaped results of Models 2 and 3.

The associations between cognitive function and other covariates in Model 4 were as expected. First, women have

poorer cognitive function than men. Age and age squared show the expected curvilinear association with cognitive function. The log-likelihood test for age and age squared in [Supplementary Table S14](#) also shows the necessity of the quadratic age term. Higher SES across all three indicators (urban HuKou, nonagricultural occupation, and higher household expenditure) is associated with higher cognitive function. Unpartnered individuals have lower cognitive scores compared with their partnered counterparts. Model 4 also shows a positive association between GDP and cognitive function.

In Model 4 (of [Table 2](#)), the random intercept standard deviation is 2.7 ($p < .001$), reflecting the significant variation in average cognitive function scores between individuals. The small standard deviation for the age random slope (0.003, $p < .01$) reflects the fact that cognitive decline follows a relatively predictable, downward trajectory with age. Similar findings were shown in previous studies, including ones that assessed other important factors that might predict the rate of cognitive decline, such as race/ethnicity ([Hale, 2017](#)), education ([Seblova et al., 2020](#)), and neighborhood environment ([Luo et al., 2019](#)).

[Table 3](#) presents results from the models that analyze the duration of exposure. Model 1 shows positive associations between $PM_{2.5}$ duration and cognitive function score, but after controlling for education, increased duration of exposure to higher $PM_{2.5}$ was negatively associated with cognitive function only for those in the second group of $PM_{2.5}$ duration (13–60 months). The association was nonsignificant for the two longer-duration groups (61–120 and 121+ months). In Model 3, SES variables explained a part, but not all, of the associations. In Model 4, when all covariates were included, the associations of $PM_{2.5}$ duration were similar to Models 2 and 3, suggesting people in the second group of exposure duration were more likely to have the poorer cognitive function, while the third and fourth groups were not statistically different than the first. The strong association for duration group 2 compared with the other groups is addressed at length later.

As well as considering $PM_{2.5}$ intensity and duration separately, we were also interested in their joint or cumulative association with cognitive function. At baseline, the individual distribution of cumulative $PM_{2.5}$ exposure (intensity \times duration) from March of 2000 to the survey date was shown in [Supplementary Table S11](#). Due to small numbers in the “3–2” group (only 169), we merged that group into the “3–3” group. [Table 4](#) shows the associations between cumulative exposure (measured by a joint variable of intensity with duration) and cognitive function. In Model 1, compared with the “1–1” group (lowest intensity and shortest duration), higher levels of intensity-duration were associated with better cognitive function, except for the group of “1–2,” which suggests people in that group had lower cognitive function than

those in the “1–1” group. In Model 2, when education was included, most coefficients of $PM_{2.5}$ exposure were reversed to negative, suggesting that education was a confounder of the association between $PM_{2.5}$ exposure and cognitive function. When more SES covariates and partnership status were added in Model 3, the negative patterns between cumulative exposure to $PM_{2.5}$ and cognitive function persisted for most groups, though the most groups were nonsignificant. These results were also robust in Model 4 when logged GDP was controlled. We found the effect size of the “1–2” group was significantly larger than other groups, which is worth exploring in future analyses. Comparing the coefficients of “1–2” (longer duration) and “2–1” (higher intensity) group suggests that duration might be more significantly associated with cognitive function than intensity.

To test whether $PM_{2.5}$ exposure is associated with accelerated cognitive decline, based on Model 4 in [Table 4](#), Model 5 adds interaction terms between cumulative exposure, age, and age squared (full results in [Supplementary Table S13](#)). [Figure 2](#) plots predicted cognitive function by cumulative $PM_{2.5}$ across age, visually depicting that level of cumulative $PM_{2.5}$ exposure is not statistically significantly associated with different trajectories (details in [Supplementary Table S13](#)). We found no statistically significant associations between higher exposure for longer duration groups (e.g., “3–3” or “4–4” groups) and faster rate of cognitive decline.

Considering the significant difference in education and air pollution between urban and rural areas, we also stratified the analysis by HuKou status. [Figure 3](#) displays the coefficients for $PM_{2.5}$, which suggest that the harmful associations of cumulative air pollution exposure were more significant among respondents with rural HuKou than those with urban HuKou. [Supplementary Figures S4 and S5](#) show the coefficients of $PM_{2.5}$ intensity and duration.

We also conducted robustness checks. First, we ran all of the above models using the MI data sets (see [Supplementary Tables S3–S5](#)). Comparing all results in [Tables 2–4](#) with [Supplementary Tables S3–S5](#), we found that there were no meaningful differences. Second, when we divided $PM_{2.5}$ intensity into six categories ([Supplementary Table S6](#)) for the data with MI, the findings were similar to [Table 2](#), but with reduced effect size and not always significant. Results for the duration and the cumulative measure ([Supplementary Tables S7 and S8](#)) were also consistent with the main findings ([Tables 3 and 4](#)), suggesting an association of more intensive cumulative $PM_{2.5}$ exposure with lower cognitive function scores. Third, [Supplementary Tables S9 and S10](#) (using continuous variable for $PM_{2.5}$ exposure) suggest that this association between $PM_{2.5}$ exposure and cognitive function may not be a quadratic curve but a more complex curvilinear relationship. Fourth, the robustness check using the fully balanced data, with a smaller sample size

(e.g., no attritors; [Supplementary Figure S6](#)) confirms that cumulative exposure remains associated with poor cognitive function, but these associations diminish for PM_{2.5}

exposures of high intensity and high duration. This is possibly because those who attrited are more likely to have poor health.

Table 3. Associations Between PM_{2.5} Duration (at a Threshold of 50 µg/m³) and Cognitive Function

	Model 1: Base	Model 2: Model 1 + Education	Model 3: Model 2 + SES + Partnership	Model 4: Model 3 + GDP
PM _{2.5} duration (ref: 1 [0–12 months])				
2 (5 years: 13–60 months)	0.0319 (–0.150 to 0.214)	–0.437*** (–0.604 to –0.270)	–0.541*** (–0.706 to –0.377)	–0.495*** (–0.661 to –0.330)
3 (10 years: 61–120 months)	0.306*** (0.139 to 0.472)	–0.0785 (–0.229 to 0.0715)	–0.128# (–0.275 to 0.0198)	–0.125# (–0.272 to 0.0228)
4 (10+ years: 121 months+)	0.509*** (0.329 to 0.689)	–0.0734 (–0.236 to 0.0895)	–0.0981 (–0.258 to 0.0622)	–0.155# (–0.316 to 0.00597)
Age	0.125*** (0.0542 to 0.196)	0.252*** (0.188 to 0.316)	0.239*** (0.175 to 0.302)	0.238*** (0.175 to 0.301)
Age square	–0.00231*** (–0.0029 to –0.0018)	–0.00277*** (–0.0033 to –0.0023)	–0.00270*** (–0.0032 to –0.0022)	–0.00271*** (–0.0032 to –0.0022)
Gender (ref: men)				
Women	–1.286*** (–1.415 to –1.157)	–0.0564 (–0.177 to 0.0641)	–0.105# (–0.224 to 0.0138)	–0.131* (–0.250 to –0.0123)
Education (ref: no-schooling)				
Primary		2.642*** (2.492 to 2.792)	2.430*** (2.281 to 2.578)	2.372*** (2.223 to 2.520)
Middle		5.095*** (4.929 to 5.260)	4.359*** (4.183 to 4.535)	4.287*** (4.111 to 4.464)
HuKou (ref: rural)				
Urban			1.178*** (1.023 to 1.332)	1.143*** (0.988 to 1.298)
Occupation (ref: agricultural)				
Nonagricultural			0.475*** (0.326 to 0.623)	0.440*** (0.291 to 0.588)
Managerial			0.313** (0.0819 to 0.545)	0.291* (0.0597 to 0.523)
Log household expenditure			0.0907*** (0.0664 to 0.115)	0.0873*** (0.0629 to 0.112)
Partnership (ref: partnered)				
Single			–0.349*** (–0.516 to –0.183)	–0.342*** (–0.508 to –0.176)
Log GDP				0.324*** (0.229 to 0.418)
Constant	11.27*** (9.075 to 13.47)	2.264* (0.260 to 4.268)	2.165* (0.166 to 4.165)	–1.035 (–3.240 to 1.170)
Random effects				
Within individual				
Change rate (age)	0.002***	0.005***	0.004***	0.004***
Intercept	2.946***	2.801***	2.663***	2.664***
Covariance	0.139	–0.008	–0.008	–0.009
Between individual				
Residuals	3.053***	3.049***	3.054***	3.054***
Log likelihood	–82,035.322	–80,444.162	–80,204.329	–80,181.936
Observations	29,484	29,484	29,484	29,484
Number of IDs	12,481	12,481	12,481	12,481

Notes: Cognitive function includes three components: immediate recall, delayed recall, and serial 7s, 0–25 points (see details in “Methods”). SES = socioeconomic status; GDP = gross domestic product.

*** $p < .001$, ** $p < .01$, * $p < .05$, # $p < .1$.

Table 4. Associations Between Cumulative PM_{2.5} Exposure (Intensity-Duration) and Cognitive Function

	Model 1: Base	Model 2: Model 1 + Education	Model 3: Model 2 + SES + Partnership	Model 4: Model 3 + GDP
Cumulative PM _{2.5} (ref: 1-1)				
1-2	-0.612* (-1.098 to -0.125)	-0.775*** (-1.214 to -0.337)	-0.691** (-1.123 to -0.260)	-0.684** (-1.115 to -0.253)
2-1	0.394** (0.0969 to 0.691)	0.125 (-0.144 to 0.393)	0.102 (-0.163 to 0.366)	0.0850 (-0.179 to 0.349)
2-2	0.202# (-0.0192 to 0.423)	-0.399*** (-0.597 to -0.202)	-0.523*** (-0.718 to -0.329)	-0.482*** (-0.676 to -0.287)
2-3	0.522*** (0.246 to 0.797)	-0.0327 (-0.287 to 0.222)	-0.146 (-0.398 to 0.105)	-0.156 (-0.408 to 0.0950)
3-3	0.419*** (0.222 to 0.617)	-0.0292 (-0.205 to 0.146)	-0.0886 (-0.261 to 0.0836)	-0.0869 (-0.259 to 0.0849)
3-4	0.486*** (0.269 to 0.702)	-0.0895 (-0.285 to 0.106)	-0.143 (-0.335 to 0.0487)	-0.195* (-0.388 to -0.00315)
4-4	1.027*** (0.768 to 1.285)	0.107 (-0.124 to 0.339)	0.115 (-0.112 to 0.342)	0.0359 (-0.192 to 0.264)
Age	0.124*** (0.0527 to 0.194)	0.251*** (0.187 to 0.315)	0.239*** (0.175 to 0.302)	0.238*** (0.175 to 0.302)
Age squared	-0.00229*** (-0.0029 to -0.0017)	-0.00276*** (-0.0033 to -0.0023)	-0.00270*** (-0.0032 to -0.0022)	-0.00271*** (-0.0032 to -0.0022)
Gender (ref: men)				
Women	-1.288*** (-1.417 to -1.159)	-0.0608 (-0.181 to 0.0597)	-0.109# (-0.228 to 0.0101)	-0.134* (-0.253 to -0.0154)
Education (ref: no-schooling)				
Primary		2.635*** (2.485 to 2.785)	2.423*** (2.275 to 2.571)	2.366*** (2.217 to 2.515)
Middle		5.078*** (4.912 to 5.244)	4.342*** (4.166 to 4.518)	4.273*** (4.096 to 4.450)
HuKou (ref: rural)				
Urban			1.177*** (1.023 to 1.332)	1.143*** (0.988 to 1.297)
Occupation (ref: agricultural)				
Nonagricultural			0.477*** (0.329 to 0.625)	0.442*** (0.294 to 0.590)
Managerial			0.313** (0.0810 to 0.544)	0.291* (0.0593 to 0.522)
Log household expenditure			0.0901*** (0.0657 to 0.114)	0.0866*** (0.0623 to 0.111)
Partnership (ref: partnered)				
Single			-0.350*** (-0.517 to -0.184)	-0.343*** (-0.509 to -0.176)
Log GDP				0.319*** (0.225 to 0.414)
Constant	11.15*** (8.949 to 13.35)	2.238* (0.231 to 4.245)	2.132* (0.129 to 4.136)	-1.022 (-3.230 to 1.186)
Random effects				
Between individual				
Change rate (age)	0.003***	0.005***	0.004***	0.004***
Intercept	2.930***	2.792***	2.653***	2.655***
Covariance	0.007***	-0.009	-0.010	-0.010
Within individual				
Residuals	3.052***	3.048***	3.054***	3.054***
Log likelihood	-82,012.150	-80,437.023	-80,197.855	-80,176.084
Observations	29,484	29,484	29,484	29,484
Number of IDs	12,481	12,481	12,481	12,481

Notes: Cognitive function includes three components: immediate recall, delayed recall, and serial 7s, 0-25 points (see details in "Methods"). The first number in the cumulative PM_{2.5} is intensity (1: 0-35 µg/m³; 2: 36-50 µg/m³; 3: 51-75 µg/m³; 4: 76+ µg/m³), and the second represents duration (1: 0-12 months; 2: 13-60 months; 3: 61-120 months; 4: 121+ months). SES = socioeconomic status; GDP = gross domestic product.
***p < .001, **p < .01, *p < .05, #p < .1.

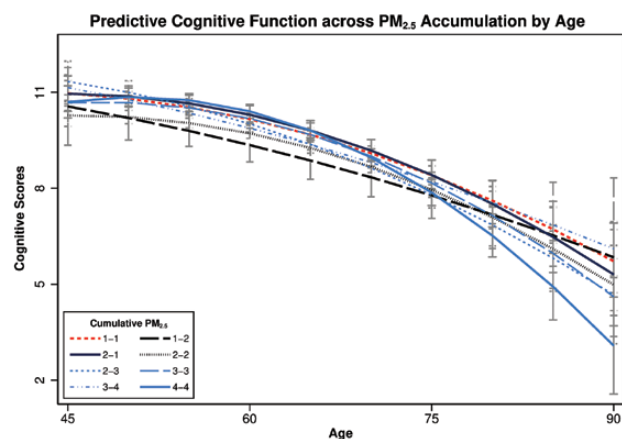


Figure 2. Trajectories of cognitive function associated with cumulative $PM_{2.5}$ (intensity-duration) across age with 95% confidence intervals. Adjusted covariates were from Model 5 in [Supplementary Table S13](#). The first number in the cumulative $PM_{2.5}$ is intensity (1: 0–35 $\mu g/m^3$; 2: 36–50 $\mu g/m^3$; 3: 51–75 $\mu g/m^3$; 4: 76+ $\mu g/m^3$), and the second represents duration (1: 0–12 months; 2: 13–60 months; 3: 61–120 months; 4: 121+ months).

Discussion

Using the CHARLS, a nationally representative data set, linked to historic $PM_{2.5}$ records derived from remotely sensed satellite data, we investigated the relationship between $PM_{2.5}$ exposure at the city level and cognitive function via a sequence of GCMs. Our findings were mixed but suggested that some categories of higher and longer exposure to $PM_{2.5}$ are associated with poorer cognitive function among Chinese adults aged 45 and older.

First, we used the intensity (average $PM_{2.5}$ exposure) as the measure for exposure over a long-term period of 15 years. Our findings show that respondents in the second group of $PM_{2.5}$ intensity have poorer cognitive function than those in the first group, but their associations retained only weak levels of significance once adjusted for education and other SES factors. Moreover, our findings in [Table 2](#) and [Supplementary Table S3](#) show that people living in areas of much higher $PM_{2.5}$ intensity (e.g., categories three and four) have a better cognitive function.

Although some studies show no significant associations between $PM_{2.5}$ intensity and cognitive impairment ([Gatto et al., 2014](#); [Loop et al., 2013](#)), our findings are consistent with most of the previous research, which indicates higher mean concentrations of $PM_{2.5}$ were significantly associated with lower cognitive function ([Ailshire & Clarke, 2015](#); [Ailshire et al., 2017](#); [Tallon et al., 2017](#); [Wang et al., 2020](#)). We have controlled for many individual and city-level confounders, but as in most observational studies, we cannot rule out residual confounding factors related to air pollution or cognitive function, or both, which might result in high between-person residuals and unexpected patterns of associations between air pollution and cognitive function.

Although our study adjusts for HuKou status, this may not capture every important element of rural–urban

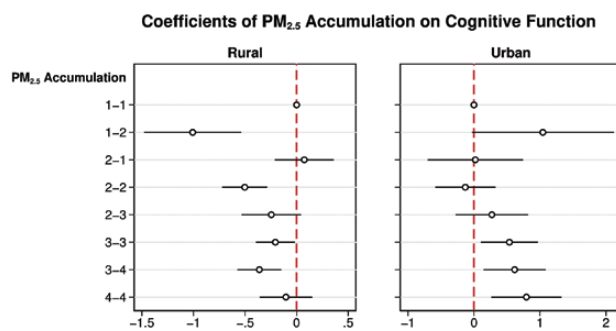


Figure 3. Associations between cumulative $PM_{2.5}$ exposure (intensity-duration) and cognitive function stratified by HuKou status. Adjusted covariates include age, age squared, gender, education, occupation, household expenditure (logged), and annual GDP at the city level (logged). GDP = gross domestic product.

differences. For example, economic development and urbanization correlate with high pollution and high rates of rural–urban migration ([Chai et al., 2014](#); [Liu et al., 2007](#); [Zhang & Kanbur, 2005](#)). Therefore, the negative association between higher air pollution (e.g., the highest category of $PM_{2.5}$ intensity or duration) and poorer cognition might be offset by the advantageous characteristics of urban dwellers. This might explain why the analyses stratified by HuKou show that rural respondents had stronger associations between $PM_{2.5}$ exposure and poor cognitive function than urban ones. We additionally included measures of educational attainment and other SES covariates (household expenditure and occupation), but these also may not capture all rural–urban differences (e.g., medical insurance, housing, and community services). Furthermore, the remaining high between-person residuals might be due to measuring air pollution exposure at the city-level resolution. In China, the city region may consist of rural and urban areas where air pollution exposures are remarkably different. This should be explored in future research with the different data source(s) that allow the inclusion of more potential confounding factors and more precise spatial resolution.

Second, our results suggest that exposure to 13–60 months of $PM_{2.5}$ duration was associated with worse cognitive function than the first-group duration (0–12 months), consistent with previous studies regarding the effects of exposure durations ([Liu et al., 2017](#); [Tan et al., 2018](#)). However, coefficients for durations longer than 60 months were nonsignificant, and likely for similar reasons as observed with intensity: the stronger association between the second duration group and cognitive function could be caused by some unobserved individual factors (e.g., people living in the regions of the second duration group might have less protection against air pollution) or urbanization factors. Compared with the effect sizes and significance of $PM_{2.5}$ intensity, our findings reflect that $PM_{2.5}$ duration might have more detrimental influences on cognitive function.

Third, the results using a joint measure of intensity with duration for cumulative exposure indicate that the exposure to a longer duration at a lower intensity may be more harmful than a shorter duration at a higher intensity. This suggests that duration of exposure may play a more important role in cognitive impairment than the intensity of exposure, especially in a 15-year period or longer. On the other hand, there might be an alternative explanation for the lower cognitive level of the “1–2” group. After further investigation, we found that more than 80% of the individuals in the “1–2” group were from one particular city (Maoming of Guangdong Province), where urbanization dramatically increased the level of $PM_{2.5}$ concentration during the last 15 years (Maoming Municipal Government, 2020). Supplementary Figure S3 shows that there were only 16 months with more than $50 \mu\text{g}/\text{m}^3$ at $PM_{2.5}$ concentration, which indicates most samples in Maoming are exposed to the “1–1” and “1–2” groups. This also partly explains why the coefficients of the third and fourth groups of duration were nonsignificant in Table 4. Therefore, these findings suggest that the measure of air pollution should include intensity and duration simultaneously.

It should be noted that our findings related to duration were substantiated only when we defined that the harmful threshold is more than $50 \mu\text{g}/\text{m}^3$ at $PM_{2.5}$ concentration. Evidence shows that poor cognitive performance is associated with a lower level of $PM_{2.5}$ concentration (e.g., $15 \mu\text{g}/\text{m}^3$) among American older adults (Ailshire & Clarke, 2015). In China, although the national standard for annual $PM_{2.5}$ concentration is $35 \mu\text{g}/\text{m}^3$ (level 1) and $75 \mu\text{g}/\text{m}^3$ (level 2; Cao et al., 2013), we found that 85% of respondents in the CHARLS were living in cities where the annual level of $PM_{2.5}$ exposure is over $35 \mu\text{g}/\text{m}^3$. Thus, if we used a lower threshold to measure the duration, the variation of duration would be too small to accurately reflect the relationship between $PM_{2.5}$ exposure and cognitive function. In addition, China was experiencing rapid urbanization, which to some extent leads to a positive correlation between human health and air pollution because the increased GDP and improved infrastructure are likely to be the main drivers of beneficial health outcomes; this might have downwardly biased our estimates of the association between air pollution and cognitive function (Hou et al., 2019).

Several limitations should be noted. First, because we did not have access to respondents' address data, we matched individuals to $PM_{2.5}$ exposure data based on the city where they lived; hence, we cannot compare respondents within the same city. Future studies that can match air pollution data at the individual level (not possible with our data) could produce more robust results. This study can only indicate the associations between $PM_{2.5}$ exposure and cognitive function, but cannot examine the effects of other air pollutants, such as NO_2 , PM_{10} , and ozone, which are also associated with cognitive impairment (Kulick et al., 2020; Shin et al., 2018). Thus, we cannot rule out that our findings might result from other pollutants that are highly

correlated with $PM_{2.5}$. Finally, although we use long-term $PM_{2.5}$ exposure data measured over a 15-year period, we have no measure of the earlier life course exposure.

Nevertheless, our study makes two important contributions. First, we extend the measure of air pollution exposure beyond exposure intensity (average $PM_{2.5}$ concentrations) to exposure duration (the number of months with high levels of $PM_{2.5}$ concentrations) and cumulative exposure (a joint variable of intensity with duration), providing a more comprehensive assessment of the associations between cumulative $PM_{2.5}$ exposure and cognitive function. Second, compared with monitoring station data in China, the satellite-based $PM_{2.5}$ data used in this study have many methodological advantages, including a broad spatial coverage (all of China) and long-term records (more than 15 years). Third, the linkage between survey data and satellite data is established using exact interview dates and locations over a period of 15 years, enabling us to accurately observe the temporal trends of $PM_{2.5}$ exposure, even if interview dates varied between respondents. Using a long-term exposure period is important when studying cognitive impairment, because the initial deposition of brain diseases (e.g., dementia and Alzheimer's disease) may begin at least 10–15 years before clinically detectable symptoms associated with cognitive impairment (Fortea et al., 2021; Tarawneh & Holtzman, 2012).

Conclusions

This study contributes to the accumulating literature linking cumulative exposure to $PM_{2.5}$ and cognitive function. We find some evidence that higher and longer $PM_{2.5}$ exposure is associated with worse cognitive function. This suggests that studies of the association between $PM_{2.5}$ exposure and cognitive function should consider both intensity and duration simultaneously. On the other hand, we consistently find that those with the lowest exposure to $PM_{2.5}$ have the lowest levels of cognitive function. The relationship is likely complicated by the association between SES, residence, and migration patterns. Future studies should try to unpick the influence of these factors, and to better understand the causal mechanisms underlying the association between cumulative exposure to $PM_{2.5}$ and cognitive decline.

Supplementary Material

Supplementary data are available at *The Journals of Gerontology, Series B: Psychological Sciences and Social Sciences* online.

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Conflict of Interest

The authors declare that they have no competing interests.

References

- Ailshire, J., & Clarke, P. (2015). Fine particulate matter air pollution and cognitive function among U.S. older adults. *The Journals of Gerontology, Series B: Psychological Sciences and Social Sciences*, 70(2), 322–328. doi:10.1093/geronb/gbu064
- Ailshire, J., & Crimmins, E. (2014). Fine particulate matter air pollution and cognitive function among older US adults. *American Journal of Epidemiology*, 180(4), 359–366. doi:10.1093/aje/kwu155
- Ailshire, J., Karraker, A., & Clarke, P. (2017). Neighborhood social stressors, fine particulate matter air pollution, and cognitive function among older U.S. adults. *Social Science and Medicine*, 172, 56–63. doi:10.1016/j.socscimed.2016.11.019
- Ballesteros, S., Nilsson, L. -G., & Lemaire, P. (2009). Ageing, cognition, and neuroscience: An introduction. *European Journal of Cognitive Psychology*, 21(2–3), 161–175. doi:10.1080/09541440802598339
- Block, M. L., & Calderón-Garcidueñas, L. (2009). Air pollution: Mechanisms of neuroinflammation and CNS disease. *Trends in Neurosciences*, 32(9), 506–516. doi:10.1016/j.tins.2009.05.009. Elsevier Current Trends
- Cagney, K. A., & Lauderdale, D. S. (2002). Education, wealth, and cognitive function in later life. *The Journals of Gerontology, Series B: Psychological Sciences and Social Sciences*, 57(2), 163–172. doi:10.1093/geronb/57.2.P163
- Calderón-Garcidueñas, L., Kavanaugh, M., Block, M., D'Angiulli, A., Delgado-Chávez, R., Torres-Jardón, R., González-Maciél, A., Reynoso-Robles, R., Osnaya, N., Villarreal-Calderon, R., Guo, R., Hua, Z., Zhu, H., Perry, G., & Diaz, P. (2012). Neuroinflammation, hyperphosphorylated tau, diffuse amyloid plaques, and down-regulation of the cellular prion protein in air pollution exposed children and young adults. *Journal of Alzheimer's Disease*, 28(1), 93–107. doi:10.3233/JAD-2011-110722
- Cao, J., Chow, J. C., Lee, F. S. C., & Watson, J. G. (2013). Evolution of PM_{2.5} measurements and standards in the U.S. And future perspectives for China. *Aerosol and Air Quality Research*, 13(4), 1197–1211. doi:10.4209/aaq.2012.11.0302
- Carpenter, J. R., & Kenward, M. G. (2012). *Multiple imputation and its application*. John Wiley & Sons, Ltd. doi:10.1002/9781119942283
- Chai, F., Gao, J., Chen, Z., Wang, S., Zhang, Y., Zhang, J., Zhang, H., Yun, Y., & Ren, C. (2014). Spatial and temporal variation of particulate matter and gaseous pollutants in 26 cities in China. *Journal of Environmental Sciences*, 26(1), 75–82. doi:10.1016/s1001-0742(13)60383-6
- Chen, H., Kwong, J. C., Copes, R., Hystad, P., van Donkelaar, A., Tu, K., Brook, J. R., Goldberg, M. S., Martin, R. V., Murray, B. J., Wilton, A. S., Kopp, A., & Burnett, R. T. (2017). Exposure to ambient air pollution and the incidence of dementia: A population-based cohort study. *Environment International*, 108, 271–277. doi:10.1016/j.envint.2017.08.020
- Clifford, A., Lang, L., Chen, R., Anstey, K. J., & Seaton, A. (2016). Exposure to air pollution and cognitive functioning across the life course—A systematic literature review. In *Environmental research* (Vol. 147, pp. 383–398). doi:10.1016/j.envres.2016.01.018
- Crimmins, E. M., Kim, J. K., Langa, K. M., & Weir, D. R. (2011). Assessment of cognition using surveys and neuropsychological assessment: The Health and Retirement Study and the Aging, Demographics, and Memory Study. *The Journals of Gerontology, Series B: Psychological Sciences and Social Sciences*, 66B(Suppl. 1), i162–i171. doi:10.1093/GERONB/GBR048
- De Vocht, F., Burstyn, I., & Sanguanchaiyakrit, N. (2015). Rethinking cumulative exposure in epidemiology, again. *Journal of Exposure Science and Environmental Epidemiology*, 25(5), 467–473. doi:10.1038/jes.2014.58
- Fortea, J., Zaman, S. H., Hartley, S., Rafii, M. S., Head, E., & Carmona-Iragui, M. (2021). Alzheimer's disease associated with Down syndrome: A genetic form of dementia. *Lancet Neurology*, 20(11), 930–942. doi:10.1016/S1474-4422(21)00245-3
- Fu, P., & Yung, K. K. L. (2020). Air pollution and Alzheimer's disease: A systematic review and meta-analysis. *Journal of Alzheimer's Disease*, 77(2), 701–714. doi:10.3233/JAD-200483
- Gatto, N. M., Henderson, V. W., Hodis, H. N., St. John, J. A., Lurmann, F., Chen, J. -C., & Mack, W. J. (2014). Components of air pollution and cognitive function in middle-aged and older adults in Los Angeles. *NeuroToxicology*, 40, 1–7. doi:10.1016/J.NEURO.2013.09.004
- Ghisletta, P., Rabbitt, P., Lunn, M., & Lindenberger, U. (2012). Two thirds of the age-based changes in fluid and crystallized intelligence, perceptual speed, and memory in adulthood are shared. *Intelligence*, 40(3), 260–268. doi:10.1016/j.intell.2012.02.008
- Gong, P., Liang, S., Carlton, E. J., Jiang, Q., Wu, J., Wang, L., & Remais, J. V. (2012). Urbanisation and health in China. *Lancet*, 379(9818), 843–852. doi:10.1016/S0140-6736(11)61878-3. Elsevier
- Guxens, M., & Sunyer, J. (2012). A review of epidemiological studies on neuropsychological effects of air pollution. *Swiss Medical Weekly*, 142(1), 1–7. doi:10.4414/smw.2011.13322. EMH Swiss Medical Publishers Ltd
- Häkansson, K., Rovio, S., Helkala, E. L., Vilks, A. R., Winblad, B., Soininen, H., Nissinen, A., Mohammed, A. H., & Kivipelto, M. (2009). Association between mid-life marital status and cognitive function in later life: Population based cohort study. *BMJ (Online)*, 339(7712), 99. doi:10.1136/bmj.b2462
- Hale, J. M. (2017). Cognitive disparities: The impact of the great depression and cumulative inequality on later-life cognitive function. *Demography*, 54(6), 2125–2158. doi:10.1007/s13524-017-0629-4
- Hou, B., Nazroo, J., Banks, J., & Marshall, A. (2019). Are cities good for health? A study of the impacts of planned urbanization

- in China. *International Journal of Epidemiology*, 48(4), 1083–1090. doi:10.1093/ije/dyz031
- Huang, W., & Zhou, Y. (2013). Effects of education on cognition at older ages: Evidence from China's Great Famine. *Social Science and Medicine*, 98, 54–62. doi:10.1016/j.socscimed.2013.08.021
- Huang, Z., & Maurer, J. (2019). Validity of self-rated memory among middle-aged and older Chinese adults: Results from the China Health and Retirement Longitudinal Study (CHARLS). *Assessment*, 26(8), 1582–1593. doi:10.1177/1073191117741188
- Kesavayuth, D., Liang, Y., & Zikos, V. (2018). An active lifestyle and cognitive function: Evidence from China. *Journal of the Economics of Ageing*, 12, 183–191. doi:10.1016/j.jeo.2018.05.001
- Kim, H., Noh, J., Noh, Y., Oh, S. S., Koh, S. B., & Kim, C. (2019). Gender difference in the effects of outdoor air pollution on cognitive function among elderly in Korea. *Frontiers in Public Health*, 7, 1–10. doi:10.3389/fpubh.2019.00375
- Kulick, E. R., Wellenius, G. A., Boehme, A. K., Joyce, N. R., Schupf, N., Kaufman, J. D., Mayeux, R., Sacco, R. L., Manly, J. J., & Elkind, M. S. V. (2020). Long-term exposure to air pollution and trajectories of cognitive decline among older adults. *Neurology*, 94(17), E1782–E1792. doi:10.1212/WNL.00000000000009314
- Lancet, T. (2014). (Barely) living in smog: China and air pollution. *Lancet*, 383(9920), 845. doi:10.1016/S0140-6736(14)60427-X. Lancet Publishing Group
- Liu, M., Zhang, Q., Lu, M., Kwon, C. S., & Quan, H. (2007). Rural and urban disparity in health services utilization in China. *Medical Care*, 45(8), 767–774. doi:10.1097/MLR.0b013e3180618b9a
- Liu, S., Yan, Z., Liu, Y., Yin, Q., & Kuang, L. (2017). Association between air pollution and chronic diseases among the elderly in China. *Natural Hazards*, 89(1), 79–91. doi:10.1007/s11069-017-2955-7
- Livingston, G., Huntley, J., Sommerlad, A., Ames, D., Ballard, C., Banerjee, S., Brayne, C., Burns, A., Cohen-Mansfield, J., Cooper, C., Costafreda, S. G., Dias, A., Fox, N., Gitlin, L. N., Howard, R., Kales, H. C., Kivimäki, M., Larson, E. B., Ogunniyi, A., & Mukadam, N. (2020). Dementia prevention, intervention, and care: 2020 report of the Lancet Commission. *Lancet*, 396(10248), 413–446. doi:10.1016/S0140-6736(20)30367-6
- Loop, M. S., Kent, S. T., Al-Hamdan, M. Z., Crosson, W. L., Estes, S. M., Estes, M. G., Quattrocchi, D. A., Hemmings, S. N., Wadley, V. G., & McClure, L. A. (2013). Fine particulate matter and incident cognitive impairment in the REasons for Geographic and Racial Differences in Stroke (REGARDS) cohort. *PLoS One*, 8(9), e75001. doi:10.1371/journal.pone.0075001
- Lu, X., Zhang, S., Xing, J., Wang, Y., Chen, W., Ding, D., Wu, Y., Wang, S., Duan, L., & Hao, J. (2020). Progress of air pollution control in China and its challenges and opportunities in the ecological civilization era. *Engineering*, 6(12), 1423–1431. doi:10.1016/J.ENG.2020.03.014
- Luo, Y., Zhang, L., & Pan, X. (2019). Neighborhood environments and cognitive decline among middle-aged and older people in China. *The Journals of Gerontology, Series B: Psychological Sciences and Social Sciences*, 74(7), e60–e71. doi:10.1093/geronb/gbz016
- Maoming Municipal Government. (2020). *Yearbooks of Maoming city (in Chinese)*. <http://www.maoming.gov.cn/zwgk/sjfb/tjnj/>
- Myung, W., Lee, C., Park, J. H., Woo, S., Kim, S., Kim, S., Chung, J. W., Kang, H. S., Lim, S. -W., Choi, J., Na, D. L., Kim, S. Y., Lee, J. -H., Han, S. -H., Choi, S. H., Kim, S. Y., Carroll, B. J., & Kim, D. K. (2016). Occupational attainment as risk factor for progression from mild cognitive impairment to Alzheimer's disease: A CREDOS study. *Journal of Alzheimer's Disease*, 55(1), 283–292. doi:10.3233/JAD-160257
- Peled, R. (2011). Air pollution exposure: Who is at high risk? *Atmospheric Environment*, 45(10), 1781–1785. doi:10.1016/J.ATMOSNV.2011.01.001
- Peters, R., Ee, N., Peters, J., Booth, A., Mudway, I., & Anstey, K. J. (2019). Air pollution and dementia: A systematic review. *Journal of Alzheimer's Disease*, 70(s1), S145–S163. doi:10.3233/JAD-180631. IOS Press
- Pope, C. A., Brook, R. D., Burnett, R. T., & Dockery, D. W. (2011). How is cardiovascular disease mortality risk affected by duration and intensity of fine particulate matter exposure? An integration of the epidemiologic evidence. *Air Quality, Atmosphere and Health*, 4(1), 5–14. doi:10.1007/s11869-010-0082-7
- Power, M. C., Weisskopf, M. G., Alexeeff, S. E., Coull, B. A., Avron, S., & Schwartz, J. (2011). Traffic-related air pollution and cognitive function in a cohort of older men. *Environmental Health Perspectives*, 119(5), 682–687. doi:10.1289/ehp.1002767
- Rabe-Hesketh, S., & Skrondal, A. (2012). *Multilevel and longitudinal modeling using Stata* (3rd ed., Vol. 1). Stata Press.
- Ranft, U., Schikowski, T., Sugiri, D., Krutmann, J., & Krämer, U. (2009). Long-term exposure to traffic-related particulate matter impairs cognitive function in the elderly. *Environmental Research*, 109(8), 1004–1011. doi:10.1016/j.envres.2009.08.003
- Rubin, D. B. (1976). Inference and missing data. *Biometrika*, 63(3), 581. doi:10.2307/2335739
- Russ, T. C., Reis, S., & Van Tongeren, M. (2019). Air pollution and brain health: Defining the research agenda. *Current Opinion in Psychiatry*, 32(2), 97–104. doi:10.1097/YCO.0000000000000480
- Seblova, D., Berggren, R., & Lövdén, M. (2020). Education and age-related decline in cognitive performance: Systematic review and meta-analysis of longitudinal cohort studies. *Ageing Research Reviews*, 58, 101005. doi:10.1016/J.ARR.2019.101005
- Shier, V., Nicosia, N., Shih, R., & Datar, A. (2019). Ambient air pollution and children's cognitive outcomes. *Population and Environment*, 40(3), 347–367. doi:10.1007/s11111-019-0313-2
- Shin, S., Burnett, R. T., Kwong, J. C., Hystad, P., van Donkelaar, A., Brook, J. R., Copes, R., Tu, K., Goldberg, M. S., Villeneuve, P. J., Martin, R. V., Murray, B. J., Wilton, A. S., Kopp, A., & Chen, H. (2018). Effects of ambient air pollution on incident Parkinson's disease in Ontario, 2001 to 2013: A population-based cohort study. *International Journal of Epidemiology*, 47(6), 2038–2048. doi:10.1093/ije/dyy172
- Sun, R., & Gu, D. (2008). Air pollution, economic development of communities, and health status among the elderly in urban China. *American Journal of Epidemiology*, 168(11), 1311–1318. doi:10.1093/aje/kwn260
- Tallon, L. A., Manjourides, J., Pun, V. C., Salhi, C., & Suh, H. (2017). Cognitive impacts of ambient air pollution in the National Social Health and Aging Project (NSHAP) cohort.

- Environment International*, 104, 102–109. doi:10.1016/j.envint.2017.03.019
- Tan, C., Wang, Y., Lin, M., Wang, Z., He, L., Li, Z., Li, Y., & Xu, K. (2018). Long-term high air pollution exposure induced metabolic adaptations in traffic policemen. *Environmental Toxicology and Pharmacology*, 58, 156–162. doi:10.1016/j.etap.2018.01.002
- Tarawneh, R., & Holtzman, D. M. (2012). The clinical problem of symptomatic Alzheimer disease and mild cognitive impairment. *Cold Spring Harbor Perspectives in Medicine*, 2(5), a006148. doi:10.1101/cshperspect.a006148
- van Ginkel, J. R., Linting, M., Rippe, R. C. A., & van der Voort, A. (2020). Rebutting existing misconceptions about multiple imputation as a method for handling missing data. *Journal of Personality Assessment*, 102(3), 297–308. doi:10.1080/00223891.2018.1530680
- Walder, A. G. (1995). Career mobility and the communist political order. *American Sociological Review*, 60(3), 309. doi:10.2307/2096416
- Wang, J., Li, T., Lv, Y., Kraus, V. B., Zhang, Y., Mao, C., Yin, Z., Shi, W., Zhou, J., Zheng, T., Kinney, P. L., Ji, J., Tang, S., & Shi, X. (2020). Fine particulate matter and poor cognitive function among Chinese older adults: Evidence from a community-based, 12-year prospective cohort study. *Environmental Health Perspectives*, 128(6), 1–9. doi:10.1289/EHP5304
- White, I. R., Royston, P., & Wood, A. M. (2011). Multiple imputation using chained equations: Issues and guidance for practice. *Statistics in Medicine*, 30(4), 377–399. doi:10.1002/sim.4067
- Xiao, Q., Chang, H. H., Geng, G., & Liu, Y. (2018). An ensemble machine-learning model to predict historical PM2.5 concentrations in China from satellite data. *Environmental Science and Technology*, 52(22), 13260–13269. doi:10.1021/acs.est.8b02917
- Xu, P., Chen, Y., & Ye, X. (2013). Haze, air pollution, and health in China. *Lancet*, 382(9910), 2067. doi:10.1016/S0140-6736(13)62693-8. Lancet Publishing Group
- Yao, Y., Lv, X., Qiu, C., Li, J., Wu, X., Zhang, H., Yue, D., Liu, K., Eshak, E. S., Lorenz, T., Anstey, K. J., Livingston, G., Xue, T., Zhang, J., Wang, H., & Zeng, Y. (2022). The effect of China's Clean Air Act on cognitive function in older adults: A population-based, quasi-experimental study. *Lancet Healthy Longevity*, 3(2), e98–e108. doi:10.1016/S2666-7568(22)00004-6
- Zeng, Y., Gu, D., Purser, J., Hoenig, H., & Christakis, N. (2010). Associations of environmental factors with elderly health and mortality in china. *American Journal of Public Health*, 100(2), 298–305. doi:10.2105/AJPH.2008.154971
- Zhang, X., & Kanbur, R. (2005). Spatial inequality in education and health care in China. *China Economic Review*, 16(2), 189–204. doi:10.1016/j.chieco.2005.02.002
- Zhao, Y., Hu, Y., Smith, J. P., Strauss, J., & Yang, G. (2014). Cohort profile: The China health and retirement longitudinal study (CHARLS). *International Journal of Epidemiology*, 43(1), 61–68. doi:10.1093/ije/dys203