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Association of individual-level concentrations and human respiratory tract deposited doses of fine particulate matter with alternation in blood pressure*



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ABSTRACT

Fine particulate matter (PM2.5) contributes to the risk of cardiovascular events, partially owing to its deposition in the human respiratory tract. To investigate short-term effects of ambient $PM_{2.5}$ exposure on alternation of blood pressure (BP), this study was conducted during the winter-summer period between 2014 and 2015. The study included 106 community residents in Wuhan city, China. We repeatedly monitored the household and outdoor PM_{2.5} concentrations as well as individual-level PM_{2.5} in each season, and then assessed personal PM_{2.5} exposure (including deposited doses of PM_{2.5} in the human respiratory tract) by using different methodology (such as using a dosimetry model). All participants took part in the physical examination, including the inflammatory indicators, BP and lung function parameters measurements. Subsequently, we assessed the health damage of exposure to PM2.5 using generalized additive models. We observed increased BP at 2-day lag for an interquartile range increase in ambient fixed-site, households, individual-level PM2.5 exposure and the corresponding lung deposited doses of each exposure concentration (p < 0.05), decreased BP at 3-day lag for an interquartile range increase in ambient fixed-site, households PM_{2.5} and the corresponding lung deposited doses of each exposure concentration (p < 0.05). The estimated deposited doses of PM_{2.5} by the deposition fractions in this study and the referenced deposition fractions by previous reported method were equivalent associated with alternation in BP. In conclusion, lung deposited dose of PM_{2.5} as a quantitative indicator may be used to assess adverse cardiovascular effects following the systemic inflammation. However, we require careful assessment of acute adverse cardiovascular effects using ambient fixed-site PM_{2.5} after short-term PM_{2.5} exposure.

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1. Introduction

Exposure to fine particulate matter (\leq 2.5 µm in aerodynamic diameter, PM_{2.5}) was associated with increased risk of hypertension (Cai et al., 2016; Wu et al., 2016). Chinese women with high individual-level PM_{2.5} (\geq 58 µg/m³) had an increase of 4.6 mmHg in systolic blood pressure (SBP) and 4.1 mmHg in pulse pressure, compared with those with low individual-level PM_{2.5} (<58 µg/m³) (Shan et al., 2014). A cohort study indicated that American women aged 35–76 years (n = 43,629) had an increase of 1.4 mmHg in SBP for each 10-µg/m³ increase in PM_{2.5} (Chan et al., 2015). Noteworthyly, the results remain controversial. A recent study reported

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Abbreviation: BMI, body mass index; BF, breathing frequency; BP, blood pressure; CRP, C-reactive protein; CI, confidence interval; DF, deposition fraction; ET, extrathoracic/head region; FRC, functional residual capacity; FeNO, fractional exhaled nitric oxide; HRT, human respiratory tract; IQR, interquartile range; MPV, mean platelet volume; MPPD, multiple path particle dosimetry model; PM, particulate matter; PM_{2.5}, particulates with an aerodynamic diameter \leq 2.5 mm; PUL, pulmonary/alveolar region; PLT, platelets; PDW, platelet distribution width; SD, standard deviation; TV, tidal volume; TB, pulmonary/tracheobronchial region; URT, upper respiratory tract; WBC, white blood count.

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that a $1-\mu g/m^3$ increment in PM_{2.5} elemental carbon was associated with 1.7% (95% confidence interval (CI): -2.81, -0.58) decreases in SBP (Mirowsky et al., 2015).

The discrepancy between studies may partially be explained by differences in exposure assessment methods. At present, there are two main kinds of methods to assess personal exposure level of PM. The direct measurement method was based on the measurement of personal exposure to PM_{2.5} using a PM personal sampler, while the indirect measurement method was to simulate personal exposure by a modeling approach based on outdoor PM_{2.5} concentrations, human time-activity patterns (Ni et al., 2016; Nieuwenhuijsen et al., 2006). However, the characteristics of PM_{2.5} (such as lung deposition, surface area and solubility) were omitted in most studies. Thus the multiple path particle dosimetry model (MPPD, Chemical Industry Institute of Toxicology, Research Triangle Park, NC) has been used in quantitative estimation of deposited PM_{2.5} doses in the human respiratory tract (HRT) (Hussein et al., 2013).

Mechanisms underlying the association between exposure to PM_{2.5} and alternation in BP remain unknown. Several experimental studies indicated that biological mechanisms (e.g.), inflammatory response, oxidative stress and platelet activation) were involved in PM_{2.5}-induced cardiovascular events (Huang et al., 2012b; Ying et al., 2014). Inhalable PM_{2.5} induced releases of proinflammatory mediators (such as cytokines, nitric oxide and platelets (PLT) from the lung cells, and then mediators spilled over into circulation system and stimulated generations of acute phase proteins from liver hepatocytes (such as C-reactive protein (CRP)), finally led to cardiovascular toxicity (Brook et al., 2010). A panel study showed that oxidative stress and inflammatory mediators (including leukocytes, PLT and CRP) modified effect of exposure to PM_{2.5} on the nighttime standard deviation of normal-to-normal intervals reductions in American adults (Lee et al., 2014). Additionally, the associations of mean platelet volume (MPV) and platelet distribution width (PDW) with inflammation and thrombosis were found (Yuri Gasparyan et al., 2011).

The study assessed individual-level PM exposure using different technological methods, including measurements of PM_{2.5} concentrations in the ambient fixed-site, residential outdoor environment, households, personal exposure level and estimation of deposited PM_{2.5} dose in the HRT, and investigated health effects of personal short-term exposure level of PM_{2.5} based on the environmental PM_{2.5} concentrations and deposited doses of PM_{2.5} in the HRT on alterations in fractional exhaled nitric oxide (FeNO), PLT, CRP and BP.

2. Material and methods

2.1. Study population and design

A total of 1240 participants (including three age groups: under 18 years, 18–60 years, over 60 years) were recruited from the Jinqiao community of Jiangan district and Qingchuan community of Hanyang district in Wuhan city, China using stratified random cluster sampling method. They had continuously lived in the communities for at least two years and did not have to think about moving in the next year. They took part in a basic physical examination and questionnaire survey covered the winter (from December 12, 2014 to February 5, 2015) and summer (from May 27, 2015 to July 30, 2015).

To assess the individual-level $PM_{2.5}$ exposure and its association with acute adverse health effect of $PM_{2.5}$ exposure, a panel study with repeated measurements of 106 participants was conducted after controlling major factors (such as gender, age, education, monthly per capita income, smoking status, drinking alcohol, physical activity, self-cooking and body mass index (BMI)) in

relation to the clinic outcomes. They were free for pulmonary diseases (including tuberculosis, emphysema and asthma), cardiovascular disease (including myocardial infarction, peripheral vascular diseases, angina and heart failure) and cancer. The participants in this study were asked to take part in measurements on three consecutive days in each season (including 24 h PM_{2.5} concentrations in their households and personal PM_{2.5} level) and provide the blood samples on the day of the physical examination and urine samples for three consecutive days within the winter and summer seasons. We only found a significant difference in the distribution of participants with self-cooking meals between the panel study individuals and the baseline ones, but no significant differences in the other variables (including gender, age, education, monthly per capita income, active smoking, passive smoking, drinking alcohol, physical activity, FeNO, BMI, SBP, and diastolic BP (DBP)) (see Supplementary Materials, Table S1).

A written informed consent was obtained from all participants prior to this study. The study was approved by the Medical Research Ethics Committee of Tongji Medical College, Huazhong University of Science and Technology.

2.2. Measurements of PM_{2.5} concentrations

We collected data on the PM_{2.5} concentrations reported by the China National Environmental Monitoring Center (marked as ambient fixed-site) nearby the communities (one ambient fixedsite is located approximately 2.0 km away from the Jingiao community of liangan district, the other is located approximately 1.2 km away from the Qingchuan community of Hanyang district) (see Supplementary Materials, Fig. S1). Simultaneously, in each season we monitored outdoor PM_{2.5} concentration in these two communities for 15 consecutive days, household concentrations of PM_{2.5} and individual-level PM_{2.5} using personal environmental monitor for three consecutive days with daily simultaneous measurements among 106 participants. To monitor the residential outdoor PM_{2.5} concentrations, the air samplers were set up on the roof of the school building at an altitude of about 18 m in the Jingiao community of Jiangan district and on the roof of the hospital building at an altitude of about 15 m in the Qingchuan community of Hanyang district, respectively. The straight-line distance between the school building or community hospital building and the main traffic road was corresponding about 330 m or 260 m. No obvious air pollution sources were found in 300 m surrounding area of the school building and in 200 m surrounding one of the community hospital building. The 24 h PM_{2,5} samples were collected on quartz microfiber filters with 90-mm diameter (Whatman Inc., Maidstone, UK) by air samplers at a flow rate of 100 L/min (TH-150C type, Wuhan Tianhong Instruments Co., Ltd., Wuhan, China). The filter was replaced at 10:00 a.m. Beijing time daily during the sampling periods.

Meanwhile, we detected PM_{2.5} concentrations in the households and individual-level PM_{2.5} using personal environmental monitor (model 200, MSP Corporation, Minnesota, USA) for three consecutive days with daily simultaneous measurements. The household PM_{2.5} sampling site was located at the living room and away from the opening windows. Air sampling pump (Gilian 5000, Gilian Instruments Corp., Sensidyne Inc., Clearwater, Florida, USA) was placed at about 1.5 m above the floor. Meanwhile, each participant was required to wear a backpack with a Gilian 5000 pump. The personal environmental monitor was kept in the breathing zone of each participant during the daytime, but was done near the participant's bed during the night sleeping period. PM_{2.5} sample was collected on quartz microfiber filters with 37-mm diameter (Munktell Inc., SE-79110 Falun, Sweden) at a flow rate of 2.0 L/min during 24-h continuous sampling. The sampler

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