

## Preplanned Studies

## Declines in Blood Lead Levels Among General Population — China, 2000–2018

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### Summary

#### What is already known about this topic?

Environmental and occupational lead exposure has generally declined in the past two decades. However, there is no large-scale monitoring of blood lead levels (BLLs) in the Chinese general population.

#### What is added by this report?

This nationally representative study showed declines of BLLs in all ages of participants; for children aged 3–5 years, down from 78.1 µg/L to 16.9 µg/L, corresponding to 78.4% decrease in the past two decades (2000–2018).

#### What are the implications for public health practice?

Recommendations for elevated BLLs on screening children at high risk now need to be revisited and updated from 100 µg/L to 50 µg/L in guidelines to conform with the substantial declines in China.

High exposure to lead is associated with numerous adverse health outcomes, particularly impairment of neural development in children (1). Large-scale biomonitoring is necessary to quantify risk assessments in public health. The United States carries out the National Health and Nutrition Examination Surveys (NHANES) to monitor changes in environmental chemicals. Monitoring of blood lead levels (BLLs) in children led to action on the reduction of exposure from for example leaded gasoline and leaded paint (2). Following the recent BLL decline, the US CDC updated the blood lead reference value (BLRV) to 35 µg/L based on the latest two rounds of the NHANES and reported in *Morbidity and Mortality Weekly Report* (3). In China, the latest national recommendations for BLLs in children, issued in 2006, established a BLRV of 100 µg/L for hyperleademia (4). Recent studies indicated that BLLs in Chinese children have been declining over the past two decades (5).

However, China does not have a suitable definition of elevated BLLs. To this end, the China CDC initiated the China National Human Biomonitoring (CNHBM) in 2017.

The distribution and temporal trends of BLLs during the past two decades in the Chinese population were from the data of a nationally representative study and two previous regional-scale surveys. The 2017/18 survey based on CNHBM, a nationally representative study for tracing the dynamic change of environmental chemicals, included 21,746 samples recruited from 152 counties in 31 provincial-level administrative divisions (PLADs) of China (Supplementary Figure S1, available in <http://weekly.chinacdc.cn>). Due to 45 participants without available BLLs information, 21,701 participants aged 3–79 years were included in this study. The CNHBM used an advanced computer-assisted personal interviewing (CAPI) online system to conduct a household interview and EpiData (version 3.2, EpiData Association, Odense, Denmark) for data entry and management, collecting the data of age, gender, residence, districts, and so on. Details of covariate definition are shown in *Supplementary Methods* (available in <http://weekly.chinacdc.cn>). Details about the study design have been previously reported (6). Approval was obtained from National Institute of Environmental Health ethics committee, Chinese Center for Disease Control and Prevention (No.201701). All participants signed written informed consent. The 2000 survey included 6,085 children aged 3–5 years randomly selected from 19 cities (7). The 2009/10 survey included 11,090 participants of the general population aged 6–59 years (8). In 2000, 2009/10, and 2017/18 surveys, venous blood specimens were collected in the morning after an overnight fast for all persons in the local community health centers, frozen, and shipped to China CDC (Beijing) for analysis. Inductively coupled plasma mass spectrometry (ICP-MS) was used to measure lead

concentrations in blood samples in the three surveys (Supplementary Methods). The lead measurement in the 2017/18 survey was finished in 2019. The BLLs in each of the three surveys were calibrated using standards prepared from lead nitrate Standard Reference Material obtained from the National Institute of Standards and Technology.

Characteristics of participants in the 2017/18 survey were stratified by quartiles of BLLs. Geometric means and distributions of BLLs and the number of participants with elevated BLLs were calculated by incorporating the weights for the entire Chinese population. The spatial distribution of BLLs in the Chinese population was mapped. The 97.5th percentile of the BLLs in participants of all age groups from CNHBM was chosen as the BLRV (3). Description of the method adopted to estimate the disease burden for children aged 3–5 years and participants aged 6–59 years are presented in Supplementary Methods, respectively. Analyses were done with SAS (version 9.4, SAS Institute Inc., Cary, USA), R (version 3.4.1, R Development Core Team, Vienna, Austria), and ArcGIS (version 3.6.1, Environmental Systems Research Institute Inc., California, USA). All *P* values were two-sided, and a *P* value of less than 0.05 was considered significant.

The geometric mean of BLLs of all 21,701 participants aged 3–79 years in the 2017/18 survey was 20.66 µg/L, and the proportions of BLLs over 50 µg/L was 6.0% (Table 1). The spatial distribution of BLLs in the Chinese population in 2017/18 is shown in Figure 1. During the past two decades (2000–2018), the BLLs of children aged 3–5 years were down from 78.1 µg/L to 16.9 µg/L, dropping by 78.4%. For participants aged 6–59 years, the BLLs were 20.41 µg/L in 2017/18, down from 36.9 µg/L in 2009/10, dropping by 44.7%. The lead exposure-related burden of disease measured in DALYs showed clear downward trends over time (Table 2). Based on the BLLs in 2017/18, the BLRV for participants aged 3–5 years was updated to 50 µg/L considering the 97.5th percentile of the BLLs in this population (Table 1). Characteristics of participants in surveys are shown in Supplementary Tables S1–S2 (available in <http://weekly.chinacdc.cn>). Supplementary Figure S2 (available in <http://weekly.chinacdc.cn>) compares the proportion of BLLs over 50 µg/L in participants from 2000 to 2018. More details of disease burden are shown in Supplementary Tables S3–S4 (available in <http://weekly.chinacdc.cn>).

## DISCUSSION

Our study found that both the BLLs among the Chinese population aged 3–5 years and 6–59 years exhibited decreases of varying degrees in the past two decades. Our study demonstrated that the decline in BLLs in the Chinese population was akin to the decline in the USA. After the implementation to eliminate leaded gasoline, which preceded the observed decline in BLLs. Leaded paint, while a major source of indoor exposure in the United States, was not widely used in the Chinese housing stock, and most residents reside in dwellings built after the banning of leaded paint. Therefore, the BLLs in Chinese residents were closely related to the levels of environmental lead exposure and the formulation of policies related to lead. However, compared with the latest monitoring results of developed countries such as the United States and Canada, the internal exposure to lead in Chinese population was still at a relatively higher level, indicating a higher disease burden and limiting intellectual quotient attainment of children on a population scale (9–10). It has been observed that BLLs in Tibet are higher than in other regions, which may be due to this population's unique dietary and living habits.

The BLRV is 100 µg/L in China at present, higher than the American standard (35 µg/L) (3) and the World Health Organization (WHO) standard (50 µg/L) (11), and therefore needs updates based on our measurements in the decline of BLLs over the past two decades. However, setting standards is complicated by various literature using inconsistent criteria for BLRV definition. In this study, we established reference values based on blood lead concentrations stratified according to age. Although a lower BLRV increases the burden on the medical system, especially in pediatrics, we cannot deny that a more stringent BLRV would identify at-risk populations as much as possible and give caregivers, communities, and officials more opportunities to act earlier. Measures to improve lead exposure monitoring and prevention interventions for high-risk populations are priorities for future work. Government agencies, employers, and worker-affiliated organizations are responsible for carrying out education in the workplace and community in accordance with the latest guidelines and recommendations.

The strengths of this study include the CNHBM data obtained by the official organization, which firstly permits the characterization of BLLs in a nationally representative sample of the general Chinese

TABLE 1. Blood lead levels (BLLs) in different age, gender, residence and districts among Chinese population in 2017/18.

Sub group	<LOD* (%)	Geometric mean (95% CI), µg/L	Median (95% CI), µg/L	P <sub>90</sub> (95% CI), µg/L	P <sub>95</sub> (95% CI), µg/L	P <sub>97.5</sub> (95% CI), µg/L	Participants with BLLs ≥35 µg/L, n (%), million <sup>†</sup>	Participants with BLLs ≥50 µg/L, n (%), million <sup>‡</sup>	Participants with BLLs ≥100 µg/L, n (%), million <sup>§</sup>
Total	0.27	20.66 (19.90–21.46)	20.72 (20.03–21.42)	42.82 (40.73–44.91)	52.72 (50.03–55.41)	65.80 (60.93–70.67)	222.4 (16.7)	79.9 (6.0)	9.0 (0.7)
Age (years)									
3–5	0.36	16.87 (16.06–17.72)	17.01 (16.33–17.68)	33.87 (31.87–35.87)	41.67 (38.32–45.02)	50.95 (44.86–57.03)	11.8 (8.7)	3.7 (2.7)	0.7 (0.5)
6–11	0.24	16.84 (16.12–17.59)	17.01 (16.34–17.67)	31.27 (29.49–33.04)	38.65 (35.91–41.40)	45.87 (42.46–49.29)	11.6 (7.1)	3.0 (1.8)	0.1 (0.1)
12–18	0.36	15.21 (14.53–15.93)	15.27 (14.71–15.83)	29.30 (27.18–31.42)	37.66 (33.73–41.59)	46.28 (38.59–53.97)	9.7 (6.1)	3.2 (2.0)	0.6 (0.3)
19–39	0.36	21.32 (20.33–22.36)	21.65 (20.50–22.79)	41.79 (39.12–44.45)	51.15 (47.70–54.61)	62.64 (55.65–69.63)	47.9 (16.2)	15.2 (5.2)	1.2 (0.4)
40–59	0.30	24.78 (23.70–25.91)	25.02 (23.90–26.14)	49.47 (47.09–51.85)	58.68 (54.54–62.81)	71.33 (64.98–77.68)	84.6 (25.2)	32.2 (9.6)	3.3 (1.0)
60–79	0.06	24.40 (23.32–25.52)	23.70 (22.75–24.64)	49.37 (45.49–53.26)	63.92 (57.30–70.54)	85.72 (78.31–93.14)	56.8 (23.8)	22.6 (9.5)	3.1 (1.3)
Sex									
Men	0.24	24.41 (23.37–25.50)	24.43 (23.39–25.48)	47.34 (44.72–49.96)	57.34 (53.26–61.41)	71.38 (65.29–77.47)	154.1 (23.3)	54.5 (3.8)	5.2 (0.8)
Women	0.28	17.51 (16.86–18.18)	17.39 (16.90–17.89)	35.37 (33.31–37.44)	45.54 (42.46–48.62)	57.44 (52.24–62.64)	68.3 (10.2)	25.4 (8.2)	3.8 (0.6)
Residence									
Urban	0.26	20.50 (19.46–21.59)	20.70 (19.83–21.57)	42.61 (39.87–45.35)	52.25 (48.98–55.52)	63.10 (55.88–70.33)	132.8 (16.5)	46.6 (5.8)	5.0 (0.6)
Rural	0.29	20.92 (19.78–22.11)	20.81 (19.60–22.02)	43.09 (39.84–46.34)	53.35 (48.27–58.42)	67.48 (61.25–73.71)	89.6 (17.1)	33.3 (6.4)	4.0 (0.8)
Districts									
North China	0.28	19.34 (18.21–20.54)	19.45 (18.28–20.62)	38.57 (34.24–42.90)	49.10 (42.41–55.80)	55.93 (46.99–64.88)	25.8 (13.5)	8.3 (4.3)	1.0 (0.5)
Northeast China	0.11	19.39 (17.40–21.61)	18.93 (16.70–21.16)	44.14 (36.97–51.30)	56.15 (47.37–64.94)	72.30 (48.78–95.83)	18.2 (16.4)	7.3 (7.1)	0.7 (0.6)
East China	0.18	19.39 (17.90–21.00)	20.03 (18.94–21.12)	38.11 (35.36–40.86)	45.53 (42.22–48.84)	54.52 (49.59–59.45)	55.2 (13.4)	13.8 (3.3)	1.4 (0.4)
South-Central China	0.09	22.22 (20.34–24.28)	21.63 (19.36–23.89)	46.97 (41.79–52.14)	56.06 (48.65–63.47)	67.70 (56.18–79.22)	65.0 (20.2)	26.3 (8.1)	2.5 (0.8)
Southwest China	0.66	23.95 (21.95–26.14)	23.69 (21.91–25.48)	51.88 (45.02–58.74)	70.76 (62.21–79.32)	89.66 (78.16–101.15)	40.7 (24.6)	18.8 (11.2)	3.1 (1.8)
Northwest China	0.36	20.36 (18.70–22.15)	20.30 (18.85–21.75)	38.13 (32.54–43.72)	47.78 (36.45–59.11)	60.89 (41.25–80.53)	17.5 (13.9)	5.4 (4.3)	0.3 (0.3)

Abbreviation: LOD=limit of detection.

\* The LOD of blood lead was 0.035 µg/L; <LOD (%) was the proportion below LOD.

† Participants with BLLs ≥35 µg/L, n (%) were calculated incorporating the sample weights.

‡ Participants with BLLs ≥50 µg/L, n (%) were calculated incorporating the sample weights.

§ Participants with BLLs ≥100 µg/L, n (%) were calculated incorporating the sample weights.

¶ Participants with BLLs ≥100 µg/L, n (%) were calculated incorporating the sample weights.

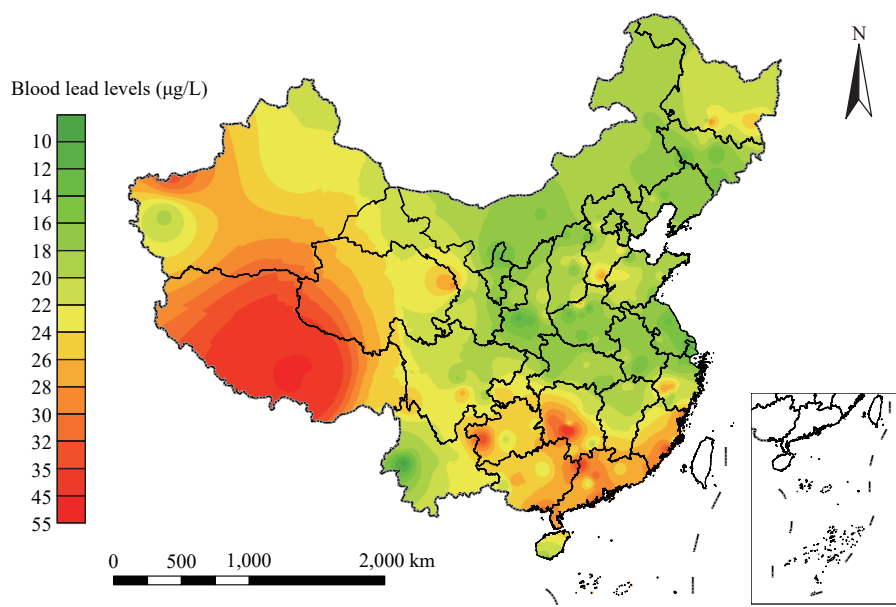


FIGURE 1. The spatial distribution of blood lead levels in Chinese population aged 3 to 79 years in 2017–2018 (N=21,701).

TABLE 2. Declines in blood lead levels (BLLs) for Chinese population aged 3–59 years during the past two decades.

Sub-population	Year	Geometric mean (95% CI), µg/L	BLLs $\geq 50$ µg/L, %	Rate of decline, %	Disease burden, DALYs (million person-years)
Aged 3–5 years					
Total				78.4	
	2000	78.1 (77.0–79.0)	84.8		16.66
	2017–2018	16.9 (16.1–17.7)	2.7		0.41
Sex					
Men				77.7	
	2000	80.0 (78.5–81.5)	88.3		8.83
	2017–2018	17.8 (16.4–19.4)	2.6		0.21
Women				78.1	
	2000	75.9 (74.3–77.5)	81.2		7.83
	2017–2018	16.6 (15.8–17.5)	2.7		0.20
Aged 6–59 years					
Total				44.7	
	2009–2010	36.9 (32.1–44.6)	29.8		9.33
	2017–2018	20.4 (20.0–20.8)	4.9		6.46
Sex					
Men				43.1	
	2009–2010	44.3 (37.8–50.8)	34.6		6.65
	2017–2018	24.2 (23.2–25.3)	7.7		4.95
Women				44.6	
	2009–2010	30.3 (24.6–36.0)	19.5		2.70
	2017–2018	16.8 (16.1–17.4)	3.2		1.50

TABLE 2. (Continued)

Sub-population	Year	Geometric mean (95% CI), µg/L	BLLs ≥50 µg/L, %	Rate of decline, %	Disease burden, DALYs (million person-years)
Age (years)					
6–11	2009–2010	35.3 (30.0–41.6)	23.7	52.4	0.21
	2017–2018	16.8 (16.1–17.6)	1.8		0.12
12–18	2009–2010	33.3 (26.1–42.6)	22.5	54.4	0.47
	2017–2018	15.2 (14.5–15.9)	2.0		0.25
19–39	2009–2010	35.8 (28.4–45.2)	28.9	40.5	1.90
	2017–2018	21.3 (20.3–22.4)	5.2		1.23
40–59	2009–2010	39.6 (33.1–47.6)	32.5	37.5	6.74
	2017–2018	24.8 (23.7–25.9)	9.6		4.71

Abbreviation: DALYs=disability-adjusted life years.

population aged 3–79 years. We updated the BLRV based on the current blood lead levels from the results of the nationally representative study with the largest sample size and scope so far. Our study conducted a longitudinal comparison of BLLs among Chinese population through three surveys, focusing on the general population rather than specific groups such as those with high occupational exposure.

This study was subject to some limitations. First, surveys in 2000 and 2009/10 were only regional-scale, though being the largest study of BLLs monitoring in China at the time, and the study samples only covered part of the age group, limiting the comparison of the changing trend of BLLs. Second, our study involved three different cross-sectional studies, with the result that different people were sampled with iterations of the study, so that determination of within-individual temporal trends in BLLs cannot be obtained.

The study demonstrated a substantial decline in BLLs and the disease burden of the entire Chinese population in the past two decades, which provided evidence that China has made significant achievements in controlling lead pollution. Furthermore, there is an urgent need to update a lower BLRV based on the significant reductions in current blood lead levels that are more sensitive to monitor and screen people with high BLLs with significant public health value for prevention of hazards of lead exposure. The study added needed evidence on behalf of populous developing countries. There is no known safe blood lead concentration, and even as BLL declines,

governments and public health institutions need to protect future generations if continued efforts to preemptively control or even eliminate lead sources and provide timely interventions for persons at the highest risk for exposure.

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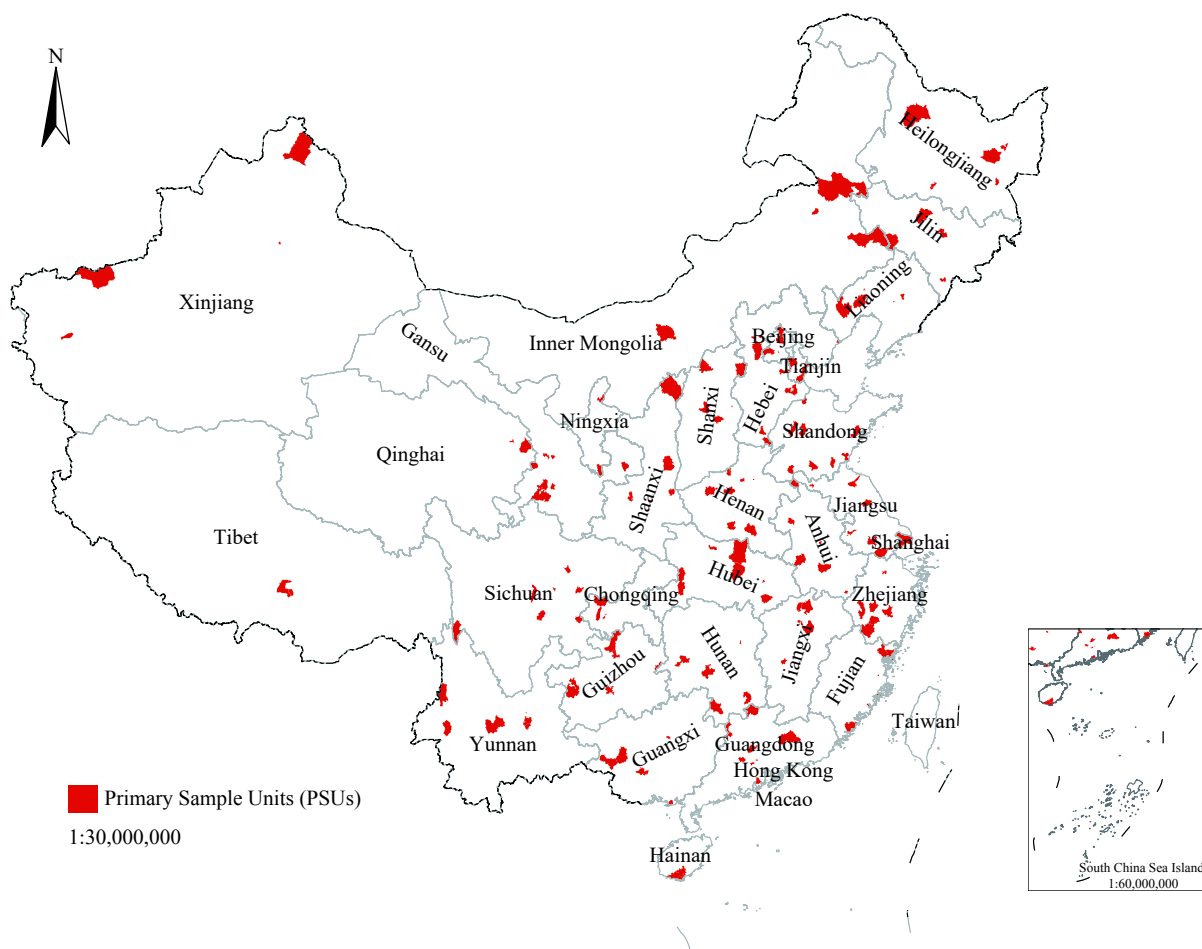


## SUPPLEMENTARY MATERIALS

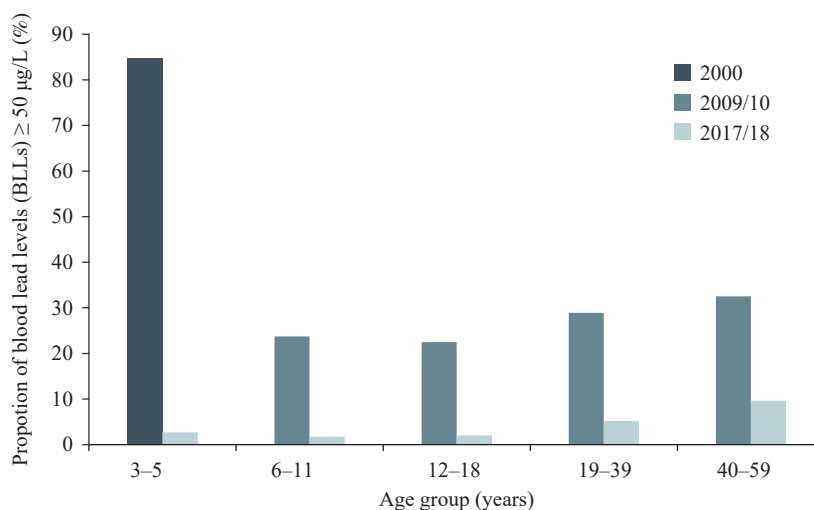
### Supplementary Methods

#### Variable Definition

Residences were categorized as rural and urban. Districts were defined as northern, northeastern, eastern, central south, southwest, and northeast. Occupation was defined as the first, secondary and tertiary industries. House type was categorized as cottage, simple building, villa, and other types. Lifestyle variables were cigarette smoking, tea drinking, and alcohol drinking. Cigarette smoking was defined as never, <70 cigarettes/week, 71–105 cigarettes/week, 106–140 cigarettes/week,  $\geq 141$  cigarettes/week. Alcohol drinking was defined as never, <2 times/month, 2–4 times/month, 2–3 times/week, and  $\geq 4$  times/week. Tea drinking was defined as <once/month, once/month–6 times/week, 7–20 times/week,  $\geq 21$  times/week. Dietary habits were constructed as described previously based on food groups including sea fish (never, <once/month, 1–7 times/month, 2 times/week, 3 times/week,  $\geq 4$  times/week, missing), sea shell (once/month, 1–7 times/month, 2 times/week,  $\geq 3$  times/week, missing), freshwater fish (once/month, 1–3 times/month, once/week, 2 times/week,  $\geq 3$  times/week, missing), rice (<4 times/week, 4–6 times/week, 7–13 times/week, 14–20 times/week,  $\geq 21$  times/week, missing), meat (<once/month, 1–2 times/month, 3–6 times/week, 7–13 times/week, 14–20 times/week,  $\geq 21$  times/week, missing), milk (<once/month, 1–3 times/month, 1–6 times/week,  $\geq 7$  times/week, missing), fruits (<once/month, 1–3 times/month, 1–6 times/week, 7–13 times/week,  $\geq 14$  times/week, missing), flour (<once/month, 1–3 times/month, 1–6 times/week, 7–13 times/week,  $\geq 14$  times/week, missing), tubers (<once/month, 1–3 times/month, 1–2



SUPPLEMENTARY FIGURE S1. Spatial locations of the monitoring units in China National Human Biomonitoring.



SUPPLEMENTARY FIGURE S2. Comparison of proportion of blood lead levels (BLLs) over 50 µg/L in 2000–2018.

times/week, ≥3 times/week, missing), beans (<once/month, 1–3 times/month, once/week, 2 times/week, ≥3 times/week, missing), nuts (<once/month, 1–3 times/month, 1–2 times/week, 3–6 times/week, ≥7 times/week, missing).

### Blood Lead Measurements

The methods for determining lead in blood (1 µg/L = 0.00483 µmol/L), including quality control and assurance procedures, have been described for each survey. Comparability has been established for the method used in five laboratories. The limits of determination (LOD) for blood lead were 0.8, 0.28, and 0.03 µg/L in the 2000, 2009/10, and 2017/18 surveys, respectively. 2.3% of samples were less than LOD in the 2017/18 surveys. For the calculation of mean lead levels, the LOD/2 was imputed for samples below the LOD. Although there is no safe blood lead level (BLL) for the population, using dichotomous BLL thresholds is advantageous for assessing trends over time. These cut points may be more easily understood than statistically derived cut points such as quartiles. An elevated BLL was defined a priori as 50 µg/L at or above selected values chosen in part because of their prior or potential use in public health policy. Given the multiple hazards of exposure to lead, during a May 2021 meeting of the Lead Exposure and Prevention Advisory Committee (LEPAC), the workgroup recommended that the blood lead reference value (BLRV) be updated from 50 µg/L to 35 µg/L using data derived from the two most recent National Health and Nutrition Examination Surveys (NHANES) cycles (2015–2016 and 2017–2018).

### Quality Control in the Laboratory of Blood Lead

We used high-purity reagents for standard solutions and sample preparation. Before analysis and determination, the instrument was tuned with a mass spectrometry tuning solution, and the experiment was carried out after the main technical indicators (low, medium, and high-quality element sensitivity, oxide ratio, and double charge ratio) reached the standard. The nebulizer and sample cone were periodically soaked and cleaned with diluted nitric acid. Calibration curves, certified standard reference materials (external quality control), parallel samples, field blanks, and laboratory reagent blanks were determined for each batch of experiments. The linear correlation coefficient  $r$  of the calibration curve was required to be greater than 0.995. The certified reference material was measured once every 30 samples, and the measurement result was within the allowable error range of the reference value; one parallel sample (randomly selected) was measured for every 20 samples. The relative error of the parallel sample was less than 10%. The laboratory reagent blank determination was less than the method's detection limit. When the measured concentration of the sample exceeded the measurement range of the calibration curve, we reduced the sampling amount or increased the dilution factor and then re-measured.



## Methods of Estimating the Burden of Disease Associated with Lead Exposure in Participants Aged 6–59 years

The burden of disease was assessed using the disability-adjusted life years (DALYs) developed by the Global Burden of Disease Study. This study assessed *YLLs* in the 6–59 age group.

*YLLs* calculation method: Firstly, based on the literature to find the dose-response relationship between lead and the risk of all-cause mortality, the  $HR(C_i)$  values under the exposure level  $C_i$  of different age groups and sex groups of Chinese residents were calculated by the hazard ratio ( $HR$ ). Mortality  $M$  in different age and gender groups was collected and calculated according to formula (1) the mortality  $M'$  in the region if exposure levels were reduced to the theoretical minimum risk value across the region. According to formula (2), the number of premature deaths due to exposure to lead ( $M_i$ ) can be calculated. This study estimated *YLLs* according to Equation (3). The  $HR(C_i)$  values involved in the formula were estimated by finding the  $HR$  values in the dose-response relationship between internal exposure and the risk of all-cause death in the relevant literature, combined with the actual exposure level of Chinese residents. The mortality data of each age group when calculating *YLLs* comes from the “2016 China Cause of Death Surveillance Dataset”.

$$M' = M/HR(C_i) \quad (1)$$

$$M_i = P_i \times M' \times (HR(C_i) - 1) \quad (2)$$

$$YLLs = M_i \times (\text{Average life expectancy} - \text{The average life expectancy of the age group}) \quad (3)$$

## Methods of Estimating the Burden of Disease Associated with Lead Exposure in Children Aged 3–5 Years

The burden of disease was calculated based on the method published in 2003 by World Health Organization (WHO). The method indicates that the loss of intelligence quotient (IQ) points can be calculated ideally based on the linear dose-response relationship between BLLs and IQ loss. When the BLL is between 50 and 200  $\mu\text{g/L}$ , the IQ decreases by 1.3 points for every 50  $\mu\text{g/L}$  increase; for BLLs over 200  $\mu\text{g/L}$ , a loss of 3.5 IQ points is assumed. When IQ levels are above 50 points, intelligence in human populations generally approximates a normal distribution. According to the distribution of BLL of Chinese residents obtained from our investigation and the assessment method provided by WHO, the proportion of children aged 3–5 years with different BLLs and the proportion of children with IQ loss caused by lead exposure were calculated respectively on the premise that the IQ distribution in the population was a normal distribution.

The WHO guidelines use the incidence of mild mental retardation (MMR) as a quantified indicator of the health effects caused by lead exposure and assess the burden of MMR disease. MMR occurs when the IQ is below 70 points but above 50 points. Even if IQ points decrease due to various diseases or exposures, they can also remain above 70 because of the higher original IQ. Consequently, assessing the disease burden of MMR requires counting the number of people with an IQ slightly above the threshold of 70 who enter the MMR range through loss of IQ points due to lead exposure. Some infectious and parasitic diseases can also cause MMR, therefore, according to the prevalence of MMR caused by known non-congenital causes in different regions, WHO calculates the adjustment ratio ( $AR$ ) of MMR. According to the method, the incidence of MMR in each age group was calculated by multiplying the proportion of children with the defined IQ point losses by the respective percentage of the population in that range of BLLs, based on the distributions of BLLs and IQ levels of children. The formula is as follows:

$$I_{MMR} = AR \times [(E1 \times P1) + (E2 \times P2) + (E3 \times P3) + (E4 \times P4)] \quad (4)$$

In formula (4),  $E1$ ,  $E2$ ,  $E3$ , and  $E4$  represent the proportion of children with BLLs of 50–100  $\mu\text{g/L}$ , 100–150  $\mu\text{g/L}$ , 150–200  $\mu\text{g/L}$ , and >200  $\mu\text{g/L}$ , respectively;  $P1$ ,  $P2$ ,  $P3$ , and  $P4$  represent the proportion of children with IQ points of 70–70.65, 70–71.95, 70–73.25, and 70–73.50, respectively;  $AR$  is the adjustment ratio and the value of China (WHO Western Pacific Region) is 3.03.3.

Disability adjusted life years (DALYs) were used to quantitatively measure the burden of MMR in children caused by lead exposure. The DALYs are the sum of years of life lost due to premature death [years of life lost (*YLLs*)] and years lived with disability including permanent disability and temporary incapacity [years lived with disability (*YLDs*)]. Because MMR due to lead exposure in children rarely leads to death, only *YLDs* were estimated. So we

performed the calculation based on the simplified formula for DALYs recommended in the approach of The Global Burden of Diseases, Injuries, and Risk Factors enterprise (GBD 2010):

$$YLD = I \times DW \times L \quad (5)$$

In formula (2),  $I$  is the number of prevalent cases, it can be obtained by multiplying IMMR by the number of children in that age group;  $DW$  is the disability weight;  $L$  is the duration of disability. It is generally assumed that intellectual impairment exists early in life and remains throughout life, life expectancy is therefore calculated as the duration of disability. The incidence of MMR, the number of children in 3–5 age group in the sixth national census of China, the average life expectancy data (77.0 years) in the 2018 Statistical Bulletin of China's Health Development and the disability weight of MMR (0.361) were substituted into the formula for calculation.

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SUPPLEMENTARY TABLE S1. The blood lead levels of Chinese children aged 3–5 years in 2000.

Sub group	Frequency	Geometric mean (95% CI), µg/L	P <sub>10</sub> (95% CI), µg/L	P <sub>25</sub> (95% CI), µg/L	Median (95% CI), µg/L	P <sub>75</sub> (95% CI), µg/L	P <sub>90</sub> (95% CI), µg/L	P <sub>95</sub> (95% CI), µg/L	P <sub>99</sub> (95% CI), µg/L
Total	6,085	78.1 (77.0–79.2)	38.5 (36.5–40.4)	62.0 (60.8–63.0)	81.4 (80.5–82.1)	110.9 (109.0–112.5)	150.5 (147.0–153.0)	175.0 (171.0–178.5)	228.0 (219.5–241.5)
Gender									
Male	3,260	80.0 (78.5–81.5)	41.5 (38.5–43.5)	63.0 (61.6–64.0)	82.5 (81.5–83.6)	114.5 (111.0–117.5)	152.4 (149.5–156.0)	176.9 (170.7–183.0)	226.6 (216.0–238.5)
Female	2,825	75.9 (74.3–77.5)	35.5 (33.0–38.0)	60.7 (59.1–62.2)	80.2 (79.0–81.3)	106.5 (103.8–109.2)	146.2 (143.0–152.1)	173.0 (167.0–178.2)	234.0 (218.0–260.0)
PLADs									
Anhui	837	70.5 (68.9–72.1)	52.0 (49.2–53.6)	61.0 (59.5–62.5)	70.0 (68.9–70.9)	82.0 (79.8–85.0)	102.5 (98.5–110.1)	124.5 (114.5–142.0)	175.2 (168.3–204.5)
Gansu	248	116.0 (109.6–122.7)	65.5 (56.5–74.0)	85.6 (79.9–93.8)	118.3 (107.6–125.0)	156.4 (147.5–176.5)	211.1 (197.8–229.0)	241.5 (226.6–266.0)	288.0 (266.0–299.0)
Hainan	526	103.7 (99.6–107.9)	52.5 (45.1–60.4)	78.61 (72.7–82.8)	115.0 (105.8–123.0)	151.9 (147.0–156.8)	176.4 (169.1–180.0)	185.5 (182.7–189.8)	197.7 (193.0–210.9)
Henan	543	128.6 (124.9–132.3)	83.3 (78.6–88.0)	107.9 (102.5–111.0)	134.0 (129.6–139.5)	160.0 (155.5–163.0)	190.0 (183.0–202.0)	215.0 (203.6–222.9)	259.50 (241.0–283.0)
Heilongjiang	1,105	76.9 (75.0–78.8)	52.2 (48.4–55.7)	70.7 (67.9–72.7)	82.0 (81.0–82.6)	90.7 (88.8–92.1)	113.5 (108.6–120.0)	136.0 (128.0–143.6)	190.30 (176.0–214.5)
Hubei	495	55.3 (52.0–58.9)	17.5 (15.0–20.0)	40.5 (32.5–49.5)	66.5 (64.5–68.5)	86.0 (80.0–90.0)	116.0 (107.5–125.5)	135.0 (126.0–149.5)	228.0 (167.0–328.5)
Hunan	443	37.3 (35.1–39.6)	17.5 (15.5–19.0)	24.0 (23.0–26.0)	35.0 (32.5–37.0)	52.0 (47.5–56.5)	99.0 (83.0–124.5)	134.0 (124.5–146.0)	199.5 (163.0–319.0)
Jilin	893	93.7 (91.0–96.6)	62.0 (58.2–64.4)	76.5 (74.6–78.2)	93.5 (90.5–96.0)	121.0 (117.5–125.0)	155.8 (146.7–165.4)	182.0 (172.0–195.5)	270.0 (234.0–443.4)
Ningxia	995	71.9 (70.0–73.8)	41.0 (38.5–43.5)	55.5 (53.5–57.5)	71.5 (69.5–74.1)	97.0 (93.4–100.0)	121.5 (118.0–128.0)	141.5 (134.5–150.5)	194.0 (170.5–231.0)

Abbreviation: PLADs=provincial level administrative divisions.

SUPPLEMENTARY TABLE S2. The blood lead levels of Chinese population in 2009/10.

Sub group	Frequency	Geometric mean (95% CI), µg/L	P <sub>10</sub> (95% CI), µg/L	P <sub>25</sub> (95% CI), µg/L	Median (95% CI), µg/L	P <sub>75</sub> (95% CI), µg/L	P <sub>90</sub> (95% CI), µg/L	P <sub>95</sub> (95% CI), µg/L	P <sub>99</sub> (95% CI), µg/L
Total	11,090	36.9 (30.3–44.9)	17.6 (14.7–20.5)	25.7 (22.3–29.1)	37.3 (31.0–43.5)	55.9 (40.0–71.9)	83.9 (38.6–129.2)	109.7 (46.4–173.0)	211.6 (–)
Age, years									
6–11	1,525	35.4 (30.0–41.6)	19.96 (17.1–22.8)	26.8 (23.0–30.6)	35.6 (30.6–40.6)	49.1 (34.8–63.4)	67.6 (36.8–98.5)	81.96 (–)	112.6 (91.0–134.3)
12–18	3,573	33.3 (26.1–42.6)	16.8 (13.0–20.6)	23.6 (19.2–28.0)	33.8 (26.2–41.4)	47.8 (26.8–68.8)	70.7 (–)	89.6 (–)	151.5 (–)
19–39	3,533	35.9 (28.5–45.2)	17.2 (13.2–21.2)	25.3 (21.3–29.3)	36.9 (29.9–44.0)	55.0 (36.6–73.5)	84.1 (34.2–134.1)	109.2 (67.0–151.3)	176.3 (–)
40–60	2,459	39.7 (33.1–47.6)	17.8 (15.2–20.4)	26.7 (23.8–29.5)	39.7 (34.1–45.3)	60.4 (45.2–75.7)	92.0 (32.2–151.7)	126.2 (–) <sup>a</sup>	262.4 (–)
Gender									
Female	5,616	29.9 (23.6–38.0)	14.6 (11.4–17.8)	21.5 (18.1–25.0)	30.3 (24.6–36.0)	45.1 (28.5–61.6)	69.1 (–) <sup>a</sup>	88.8 (–)	166.8 (–)
Male	5,474	45.0 (38.0–53.3)	22.7 (18.9–26.5)	31.5 (28.0–35.0)	44.3 (37.8–50.8)	65.5 (47.1–83.8)	96.3 (58.8–133.9)	124.2 (70.0–178.5)	225.9 (–)
Region									
Eastern China	8,010	31.9 (26.6–38.3)	21.0 (–)	27.7 (13.6–41.8)	43.1 (20.7–65.5)	65.0 (19.4–110.7)	92.2 (16.6–167.9)	114.1 (64.4–163.8)	208.5 (–)
Central China	1,376	43.4 (29.2–64.3)	15.7 (12.5–19.0)	23.6 (20.4–26.9)	33.6 (29.4–37.7)	47.0 (40.3–53.7)	65.95 (44.6–87.4)	83.2 (–)	200.1 (–)
Western China	1,704	63.8 (38.5–105.9)	30.3 (–)	43.8 (–)	66.3 (20.2–112.3)	93.0 (55.3–130.7)	125.4 (94.9–155.9)	147.8 (112.0–183.6)	227.0 (–)
City									
Beijing	568	33.9 (32.1–35.9)	18.4 (16.2–20.6)	26.1 (24.5–27.6)	36.2 (34.7–37.8)	49.8 (47.4–52.2)	66.3 (61.2–71.5)	75.9 (70.2–81.7)	100.2 (81.1–119.2)
Chaozhou	699	47.5 (45.5–49.6)	26.8 (24.7–28.8)	35.3 (33.5–37.2)	45.97 (44.1–47.8)	63.8 (60.0–67.6)	83.9 (78.2–89.5)	100.3 (85.97–114.7)	269.7 (165.0–374.5)
Chengde	554	29.9 (28.4–31.5)	15.8 (14.4–17.3)	21.1 (19.9–22.3)	30.4 (27.7–33.1)	44.3 (41.1–47.5)	54.7 (50.7–58.7)	60.1 (51.4–68.8)	81.8 (–)
Dandong	492	33.0 (31.3–34.8)	19.4 (18.1–20.7)	25.8 (24.4–27.2)	33.8 (32.5–35.2)	44.0 (40.7–47.2)	56.2 (50.2–62.2)	66.1 (56.5–75.6)	83.9 (–)
Haidong	560	44.7 (42.3–47.3)	22.4 (20.7–24.0)	30.8 (28.6–33.0)	42.9 (40.4–45.4)	62.4 (56.9–67.9)	95.5 (77.0–114.0)	130.4 (114.5–146.3)	177.9 (–)
Heze	313	9.8 (8.2–11.8)	2.3 (0.9–3.6)	5.9 (4.3–7.6)	13.9 (11.9–15.8)	23.5 (20.1–26.8)	33.1 (28.2–38.0)	41.3 (29.6–53.0)	66.4 (–)

Sub group	Frequency	Geometric mean		Median	P <sub>75</sub>		P <sub>90</sub>		P <sub>95</sub>		P <sub>99</sub>	
		(95% CI), µg/L	(95% CI), µg/L		(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L	(95% CI), µg/L
Jiaozuo	463	60.3 (56.9–64.1)	32.8 (30.1–35.5)	57.5 (53.2–61.8)	82.5 (76.8–88.3)	108.3 (98.9–117.7)	135.7 (93.3–178.1)	311.1 (–)				
Jinan	465	22.7 (20.9–24.7)	8.6 (6.5–10.7)	25.8 (23.6–27.9)	36.4 (34.4–38.4)	47.0 (43.4–50.5)	56.8 (50.2–63.3)	71.0 (30.9–111.2)				
Jinzhou	566	31.3 (29.9–32.8)	17.8 (16.7–19.0)	30.1 (28.7–31.5)	42.2 (39.3–45.2)	55.3 (51.3–59.4)	63.9 (55.2–72.7)	81.9 (–)				
Langfang	549	36.8 (34.6–39.2)	19.9 (17.9–22.0)	37.6 (35.3–40.0)	51.8 (47.7–55.9)	68.1 (64.2–72.0)	75.9 (70.6–81.2)	110.9 (–)				
Lianyungang	455	34.5 (33.1–36.0)	22.0 (20.7–23.3)	34.9 (33.5–36.3)	43.95 (40.9–47.0)	55.4 (52.5–58.3)	60.5 (55.9–65.0)	76.6 (–)				
Pingdingshan	467	32.4 (30.5–34.4)	18.4 (17.0–19.8)	30.8 (28.4–33.1)	44.5 (41.4–47.6)	54.1 (48.1–60.1)	72.7 (43.9–101.5)	151.9 (–)				
Puyang	446	39.96 (37.4–42.7)	19.4 (18.3–20.5)	39.1 (34.4–43.8)	62.8 (57.3–68.2)	83.6 (75.1–92.1)	109.1 (92.7–25.5)	144.8 (–)				
Qingdao	313	20.4 (18.1–23.0)	7.1 (5.3–9.0)	21.9 (18.4–25.4)	35.4 (29.2–41.6)	56.1 (50.1–62.1)	61.3 (52.7–69.9)	97.0 (–)				
Qingyuan	530	83.1 (76.3–90.6)	34.7 (32.1–37.3)	76.9 (67.8–86.0)	136.7 (113.5–159.8)	227.6 (188.1–267.0)	298.6 (161.3–435.8)	604.7 (–)				
Shenyang	247	25.9 (24.3–27.5)	14.9 (13.4–16.4)	27.4 (25.0–29.7)	33.7 (30.5–36.9)	44.1 (41.3–46.9)	47.8 (43.8–51.2)	66.1 (–)				
Shijiazhuang	543	32.1 (30.8–33.3)	19.4 (18.0–20.7)	31.3 (29.5–33.1)	41.6 (39.8–43.4)	52.9 (48.5–57.2)	58.9 (54.3–63.6)	80.1 (–)				
Suzhou	522	33.3 (31.7–35.1)	18.9 (17.1–20.6)	33.5 (31.7–35.4)	45.4 (43.5–47.2)	57.1 (50.8–63.3)	68.8 (61.8–75.8)	85.2 (68.1–102.3)				
Taizhou	549	35.7 (34.4–37.0)	23.2 (21.7–24.8)	35.6 (34.1–37.0)	44.2 (42.5–46.0)	55.1 (51.4–58.8)	62.7 (56.6–68.9)	84.3 (–)				
Xining	1,144	76.0 (73.4–78.6)	41.6 (38.8–44.4)	76.3 (73.5–79.1)	100.4 (96.4–104.4)	131.7 (122.8–140.6)	157.1 (139.7–174.6)	261.2 (182.8–339.7)				
Zhanjiang	645	22.9 (21.1–24.9)	10.6 (9.0–12.3)	26.1 (24.8–27.5)	36.9 (34.4–39.4)	48.8 (42.7–55.0)	60.7 (54.9–66.5)	91.5 (32.5–150.5)				

\*The frequency is too low for bootstrap to resample.

SUPPLEMENTARY TABLE S3. Burden of disease from lead exposure in Chinese population aged 6–59 years measured in 2000 and 2017/18.

Age group, years	HR	Weighted mortality, %	Adjusted weighted mortality, %	Population, million	Population of premature deaths	DALYs
6–11						
2009–2010						
Male	1.23	0.024	0.019	45.83	2,028	139,530
Female	1.23	0.015	0.012	38.71	1,097	75,453
Total	–	–	–	–	–	214,984
2017–2018						
Male	1.12	0.024	0.021	45.83	1,131	77,794
Female	1.11	0.015	0.014	38.71	564	38,785
Total	–	–	–	–	–	116,579
12–18						
2009–2010						
Male	1.22	0.036	0.030	65.74	4,257	265,231
Female	1.22	0.019	0.015	99.89	3,306	205,934
Total	–	–	–	–	–	471,166
2017–2018						
Male	1.11	0.036	0.033	65.74	2,436	151,745
Female	1.09	0.019	0.017	99.89	1,581	98,491
Total	–	–	–	–	–	250,236
19–39						
2009–2010						
Male	1.23	0.093	0.075	136.50	23,913	1,155,014
Female	1.23	0.038	0.030	218.83	15,492	748,246
Total	–	–	–	–	–	1,903,260
2017–2018						
Male	1.16	0.093	0.080	136.50	17,318	836,466
Female	1.11	0.038	0.034	218.83	8,196	395,878
Total	–	–	–	–	–	1,232,344
40–59						
2009–2010						
Male	1.26	0.450	0.358	198.83	183,229	5,093,771
Female	1.26	0.151	0.120	191.58	59,334	1,649,472
Total	–	–	–	–	–	6,743,243
2017–2018						
Male	1.18	0.450	0.381	198.83	136,697	3,800,183
Female	1.13	0.151	0.134	191.58	32,919	915,159
Total	–	–	–	–	–	4,715,342
60–79						
2009–2010						
Male	–	–	–	–	–	–
Female	–	–	–	–	–	–
Total	–	–	–	–	–	–



TABLE S3. (Continued)

Age group, years	HR	Weighted mortality, %	Adjusted weighted mortality, %	Population, million	Population of premature deaths	DALYs
2017–2018						
Male	1.17	2.929	2.493	78.27	341,216	2,661,485
Female	1.14	1.408	1.233	78.34	136,747	1,066,628
Total	–	–	–	–	–	3,728,114
Total						
2009–2010						
Male	–	–	–	–	–	6,653,547
Female	–	–	–	–	–	2,679,105
Total	–	–	–	–	–	9,332,652
2017–2018						
Male	–	–	–	–	–	7,615,225
Female	–	–	–	–	–	2,568,477
Total	–	–	–	–	–	10,183,702

Abbreviation: HR=hazard ratio; DALYs=disability-adjusted life years.

SUPPLEMENTARY TABLE S4. Burden of disease from lead exposure in Chinese population aged 3–5 years old measured in 2000 and 2017/18.

Year	No. of participants	Blood lead geometric mean, µg/L	Incidence of mild mental retardation (‰)	DALYs (Person-year/1,000 children)	DALYs (Person-year)
2000	46,612,176	78.1	13.87	357.4	16,659,509
2017–2018	45,202,983	16.9	0.33	9.0	408,780

Abbreviation: DALYs=disability-adjusted life years.