Association of education levels with the risk of hypertension and hypertension control: a nationwide cohort study in Chinese adults

Kan Sun,1 Diaoazhu Lin,2 Mian Li,3 Yiming Mu,4 Jiajun Zhao,5 Chao Liu,6 Yufang Bi,3 Lulu Chen,7 Linxian Shi,8 Qiang Li,9 Tao Yang,10 Qin Wan,11 Shengli Wu,12 Guixia Wang,13 Zuojie Luo,14 Yingfen Qin,14 Xulei Tang,15 Gang Chen,16 Yanan Huo,17 Zhengnan Gao,18 Qing Su,19 Zhen Ye,20 Ruying Hu,21 Youmin Wang,22 Guijun Qin,23 Huacong Deng,24 Xuefeng Yu,25 Feixia Shen,26 Li Chen,27 Weiqing Wang,28 Guang Ning,28 Li Yan29

ABSTRACT
Background Education attainment can improve life expectancy and guide healthy behaviours throughout an entire lifetime. A nationwide longitudinal study of the association of education status with the risk of hypertension and its control in China is lacking.

Methods The China Cardiometabolic Disease and Cancer Cohort Study is a multicentre, population-based, prospective cohort study. We performed the baseline survey from 2011 to 2012. A follow-up visit was conducted during 2014–2016. 101 959 subjects were included in the final data analyses. Cox proportional hazards regression was used to examine the associations of education levels with the risk of hypertension and uncontrolled hypertension.

Results During follow-up, 11 189 (19.9%) participants had developed hypertension among subjects without hypertension at baseline. Among the participants with hypertension at baseline, only 40.6% had controlled hypertension. Compared with the participants’ education level at elementary school and below, the multivariable-adjusted HR for incident hypertension was 0.76 (95% CI, 0.72 to 0.80) in those with a middle school education level and 0.67 (95% CI, 0.63 to 0.70) in those with a high school degree or above. Correspondingly, multivariable-adjusted HRs associated with uncontrolled hypertension were 0.90 (95% CI, 0.87 to 0.92) in participants with a middle school education level and 0.85 (95% CI, 0.82 to 0.88) in participants with a high school degree or above level.

Conclusion Participants with education attainment at elementary school and below exhibited excess risks of newly diagnosed hypertension and worse blood pressure control compared with individuals with education attainment at middle school or above.

INTRODUCTION
As the leading cause of adverse cardiovascular outcomes, hypertension (HTN) remains a worldwide burden in most economically developing and developed countries.1 The social determinants of health (SDOH) partly explain the high burden of HTN and related diseases in developing countries.2-4 Compared with other determinants, such as income level or social status, education has taken centre stage among SDOH, affecting people throughout their entire lifetimes by improving life expectancy and guiding healthy behaviours.5

As recently reported in the Prospective Urban Rural Epidemiologic study, low-income individuals with low educational levels have a lower prevalence of HTN than those with higher education attainment.6 Inconsistent with this finding, in the same study, high-income individuals with high levels of education are more likely to be health-conscious and therefore have a markedly reduced prevalence of HTN. Differences in the relationship between educational groups and HTN prevalence can be explained by markedly poorer healthcare according to the level of education and income.7 8 Additionally, individuals in low-income countries and with lower levels of education have received poorer secondary prevention and poorer therapy for HTN than other groups.9

The effect of fast-paced economic growth on SDOH changes has resulted in a rapid increase in the incidence of HTN in China.10 With the resumption of college entrance examination in 1977 and the reform and opening-up policy in 1978, the education level and lifestyle of Chinese individuals aged 40 years and older have been significantly different. The prevalence of HTN in individuals aged 40 years and older has increased significantly in China. The link between education and HTN treatment has been examined in previous studies.11-13 The influence of education is wide, not only in the lifestyle of the subjects but also in their medication compliance and frequency of blood pressure self-monitoring after a diagnosis of HTN. According to previous studies, a low education level may directly lead to the increased incidence of HTN and may also directly affect the blood pressure control of patients with previous HTN. However, although a proportion of the previous studies comprised longitudinal studies, they only included some specific ages and some regions and the sample sizes were not sufficiently large. According to previous studies, we considered that the education level is related to the incidence and control of HTN, but this finding must be confirmed in larger samples of the Chinese.
population, particularly in those aged 40 years and older and including both high-income and low-income or middle-income communities. Therefore, we performed the present cohort study in a large nationwide Chinese population to evaluate the effects of education levels on the incidence of HTN and blood pressure control in subjects with previously diagnosed HTN.

METHODS

Study population and study design

The study participants were obtained from the China Cardiometabolic Disease and Cancer Cohort (4C) Study, and details of this cohort have been published previously. From 2011 to 2012, 259,657 individuals aged ≥40 years were recruited from 25 communities throughout various regions of China. However, in 2014, because of limited funds, only 193,846 participants from 20 communities from various geographical regions in China, selected to represent the general population, were invited to participate. From 2014 to 2016, 170,240 participants completed the follow-up examination visit, and the follow-up rate of the study was 87.8%. Participants who failed to provide information on education and blood pressure were excluded from the analyses. To evaluate the effects of education levels on the incidence of HTN and blood pressure control in subjects with previously diagnosed HTN, subjects with previously diagnosed HTN at baseline were also enrolled. Finally, 101,959 subjects were included in the final data analyses. The detailed processes of the study population are presented in a flow diagram figure 1. Before data collection, each participant had provided written informed consent.

Questionnaire and biochemical investigation

Information on sociodemographic characteristics, lifestyle factors and family history were collected using a standardised questionnaire at baseline and the follow-up visit. Data collection and anthropometric measurements were conducted by clinical staff members in local community clinics who were trained and certified. Education attainment was self-reported and categorised as follows: (1) elementary school and those below elementary school, (2) middle school and (3) high school degree or above. Smoking regularly in the past 6 months was classified as current smokers, while other participants were classified as non-current smokers. Drinking regularly in the past 6 months was classified as current drinkers, while other participants were classified as non-current drinkers. Physical activity was estimated by the International Physical Activity Questionnaire, according to questions on the frequency and duration of moderate or vigorous activities and walking. Ideal physical activity was defined as ≥7.5 min/week of vigorous intensity or ≥150 min/week of moderate and vigorous intensity or ≥150 min/week of moderate intensity.

Body weight and height were, respectively, measured to the nearest 0.1 kg and 0.1 cm. Body mass index (BMI) was calculated as follows: weight in kilograms/(height in metres). Obesity was defined as BMI ≥30 kg/m², and overweight was defined as BMI ≥25 kg/m² and <30 kg/m². Venous blood samples were collected after fasting for at least 10 hours. Fasting plasma glucose (FPG), fasting insulin, creatinine, γ-glutamyltransferase (γ-GGT), total cholesterol (TC), triglycerides (TG), high-density lipoprotein cholesterol (HDL-C) and low-density lipoprotein cholesterol (LDL-C) were evaluated using the autoanalyser (Beckman CX-7 Biochemical Autoanalyser; Brea, California, USA). Homeostasis model assessment of insulin resistance (HOMA-IR), which measures the Insulin Resistance Index, was calculated as follows: fasting insulin (mIU/mL)×fasting glucose (mmol/L)/22.5. To calculate the estimated glomerular filtration rate (eGFR), we used the following abbreviated Modification of Diet in Renal Disease formula recalculated for the Chinese population: eGFR (mL/min per 1.73 m²)=175 × (serum creatinine (μmol/L)×0.011)−1.234 × (age)−0.179 × (0.79 if woman). The Three consecutive blood pressure measurements within a 5-min interval were obtained from each participant using an automated electronic device (OMRON; Omron Company, China). The average of three measurements of systolic blood pressure (SBP) and diastolic blood pressure (DBP) were used for analysis. Newly diagnosed HTN was defined as SBP ≥140 mmHg and/or DBP ≥90 mmHg and/or currently taking antihypertension medication during the follow-up visit in subjects without HTN at baseline. Uncontrolled HTN was defined as SBP ≥140 mmHg and/or DBP ≥90 mmHg during the follow-up visit in subjects with HTN at baseline.

Definition of newly diagnosed HTN and uncontrolled HTN

Continuous variables were presented as means±SD, while skewed variables were presented as medians (IQRs). Categorical variables are expressed as numbers (proportions). Differences in clinical characteristics and laboratory variables among groups were tested by one-way analysis of variance. X² test was used to compare categorical variables. Trends across different education group levels were evaluated using linear regression analyses. We used Cox proportional hazards regression models to assess the risk of newly diagnosed HTN events and uncontrolled HTN events according to individual education levels. HRs and the corresponding 95% CIs were calculated. The covariates in fully adjusted Cox proportional hazards regression models were generated according to previous publications and potential
intermediate factors related to the development and progression of HTN. Model 1 was unadjusted. Model 2 was adjusted for age. Model 3 was further adjusted for BMI and sex. Model 4 was further adjusted for current drinking, current smoking, a family history of diabetes, creatinine, GGT, FPG, insulin, LDL-C and TG. Model 5 was further adjusted for baseline SBP and DBP. In subgroup analyses, the associations of education levels with newly diagnosed HTN and uncontrolled HTN were stratified by median age (≥56.5/<56.5 years), sex (men/women), BMI (normal/overweight/obesity), diabetes status (yes/no), current drinking status (yes/no) and current smoking status (yes/no). The interaction tests were performed by simultaneously including the interaction terms (strata variable multiplied by HTN status), interaction tests were performed by simultaneously including the interaction terms (strata variable multiplied by HTN status), each strata factor and HTN status in the Cox regression analyses.

SAS V9.3 (SAS Institute) was used to perform statistical analysis. All statistical tests were two-sided, and a p value < 0.05 was indicated statistical significance.

RESULTS

Clinical characteristics of the cohort
During a 3.8-year follow-up period, 11189 (19.9%) subjects developed HTN among the 56151 subjects without HTN at baseline. Among participants with HTN at baseline, 27221 (59.4%) participants had uncontrolled HTN during the follow-up visit. The participants’ characteristics according to the HTN status are reported in table 1. Compared with participants without HTN at baseline and follow-up stage (non-HTN group), those with HTN at baseline or follow-up were older, had a higher BMI, waist circumference, FPG, fasting insulin, HOMA-IR, TC, TG, LDL-C, alanine aminotransferase, aspartate aminotransferase and γ-GGT, and were more likely to be a current drinker (all p < 0.0001). In addition to the difference in investigation and biochemical measurements among groups, a larger proportion of subjects in the non-HTN group had attained higher education levels than in the other two groups (p < 0.0001). Compared with participants with an elementary school education level and below, those with higher education levels exhibited a significantly lower HTN prevalence and decreased SBP and DBP at baseline (table 2; all p for trend < 0.0001).

Associations of the education level with newly diagnosed HTN and uncontrolled HTN
The incidences of newly diagnosed HTN in the elementary school and below group, middle school group and high school

Table 1  Baseline characteristics of the study population at follow-up according to hypertension (HTN) status

|                    | Non-HTN       | HTN at follow-up | HTN at baseline | P value |
|--------------------|---------------|------------------|-----------------|---------|
| n (%)              | 44962 (44.10) | 11189 (10.97)    | 45808 (44.93)   | --      |
| Age (years)        | 53.8±8.06     | 56.8±8.57 *†     | 59.4±8.46 *     | <0.0001 |
| Male (n (%))       | 13223 (29.41) | 4082 (36.48) *†  | 17504 (38.21) * | <0.0001 |
| BMI (kg/m²)        | 23.7±3.35     | 24.7±3.54 *†     | 25.5±3.55 *     | <0.0001 |
| WC (cm)            | 81.2±9.19     | 84.5±9.59 *†     | 86.3±9.73 *     | <0.0001 |
| SBP (mmHg)         | 118.2±11.25   | 127.1±8.95 *†    | 149.6±17.67 *   | <0.0001 |
| DBP (mmHg)         | 72.4±7.75     | 76.7±7.56 *†     | 85.1±10.95 *    | <0.0001 |
| Current smoking (n (%)) | 6434 (14.31) | 1849 (16.53) *†  | 6460 (14.10) *  | <0.0001 |
| Current drinking (n (%)) | 3863 (8.59) | 1283 (11.47) *†  | 5550 (12.12) *  | <0.0001 |
| TG (mmol/L)        | 1.1±0.86–1.71 | 1.3±0.96–1.95 *† | 1.4±1.04–2.13 * | <0.0001 |
| TC (mmol/L)        | 4.8±1.13      | 4.9±1.13 *†      | 5.1±1.12 *      | <0.0001 |
| HDL-C (mmol/L)     | 1.3±0.37      | 1.3±0.36 *       | 1.3±0.36 *      | <0.0001 |
| LDL-C (mmol/L)     | 2.8±0.86      | 2.8±0.86 *†      | 2.9±0.88 *      | <0.0001 |
| FPG (mmol/L)       | 5.7±1.38      | 6.0±1.74 *       | 6.2±1.74 *      | <0.0001 |
| Fasting insulin (μIU/ml) | 6.2 (4.40–8.50) | 6.8 (4.80–9.60) *† | 7.5 (5.20–10.60) * | <0.0001 |
| ALT (U/L)          | 14.0 (11.00–20.00) | 15.4 (11.00–22.00) *† | 16.0 (12.00–22.80) * | <0.0001 |
| AST (U/L)          | 20.0 (16.00–24.00) | 20.2 (17.00–25.00) *† | 21.0 (17.20–26.00) * | <0.0001 |
| γ-GT (mg/L)        | 18.0 (13.00–27.00) | 21.0 (15.00–32.00) *† | 23.0 (16.00–35.00) * | <0.0001 |
| eGFR (mL/min per 1.73m²) | 98.13 (89.97–104.54) | 95.85 (87.19–102.49) *† | 92.93 (83.72–99.67) * | <0.0001 |
| Ideal physical activity (n (%)) | 6780 (15.08) | 1397 (12.49) * | 5758 (12.57) * | <0.0001 |
| HOMA-IR            | 1.51 (1.06–2.18) | 1.7 (1.20–2.58) *† | 1.97 (1.31–2.95) * | <0.0001 |
| Family history of diabetes (n (%)) | 6318 (14.05) | 1223 (10.93) *† | 5469 (11.94) * | <0.0001 |
| Education level (n (%)) | 12995 (35.51) | 3650 (32.62) *† | 13995 (30.55) * | <0.0001 |
| Use of antihypertensive medications (n (%)) | – | – | 12913 (28.19) | <0.0001 |

Data were means±SD or medians (IQRs) for skewed variables or numbers (proportions) for categorical variables. P values were for analysis of variance or χ² analyses across the groups. *P < 0.05 was compared with non-HTN group. †P < 0.05 compared with HTN at baseline group.

ALT, alanine aminotransferase; AST, aspartate aminotransferase; BMI, body mass index; DBP, diastolic blood pressure; eGFR, estimated glomerular filtration rate; FPG, fasting plasma glucose; HDL-C, high-density lipoprotein cholesterol; HOMA-IR, homeostasis model assessment of insulin resistance; LDL-C, low-density lipoprotein cholesterol; SBP, systolic blood pressure; TC, total cholesterol; TG, triglycerides; WC, waist circumference; γ-GGT, γ-glutamyltransferase.
The adjusted HRs and 95% CIs for the risk of newly diagnosed HTN and uncontrolled HTN according to education level are shown in table 3. Compared with the elementary school and below group, the middle school group (unadjusted HR, 0.65; 95% CI, 0.63 to 0.68) had a lower risk of newly diagnosed HTN, and the high school degree or above group had the lowest risk (unadjusted HR, 0.51; 95% CI, 0.49 to 0.54). Similar trends were observed for uncontrolled HTN, with the elementary school and below group having the highest risk (unadjusted HR, 0.83; 95% CI, 0.80 to 0.85), followed by the middle school group (unadjusted HR, 0.75; 95% CI, 0.73 to 0.78), and the high school degree or above group having the lowest risk (unadjusted HR, 0.70; 95% CI, 0.67 to 0.73).

### Table 2 Baseline characteristics of study population at baseline by education level

| Education                     | n (%)          | Prevalence of HTN (n (%)) | Age (years) | Male (n (%)) | BMI (kg/m²) | WC (cm) | SBP (mmHg) | DBP (mmHg) | Current smoking (n (%)) | Current drinking (n (%)) | TG (mmol/L) | TC (mmol/L) | HDL-C (mmol/L) | LDL-C (mmol/L) | FPG (mmol/L) | ALT (U/L) | AST (U/L) | γ-GGT (U/L) | eGFR (mL/min per 1.73 m²) | Ideal physical activity (n (%)) | HOMA-IR | Family history of diabetes (n (%)) | Data were means±SD or medians (IQRs) for skewed variables or numbers (proportions) for categorical variables. | P for trend was calculated for the linear regression analysis tests across the groups. P values were for analysis of variance or χ² analyses across the groups. |
|-------------------------------|----------------|---------------------------|-------------|--------------|-------------|---------|------------|-----------|------------------------|--------------------------|-------------|-------------|----------------|----------------|-------------|----------|----------|-------------|-----------------------------|--------------------------|---------|--------------------------|-----------------------------|--------------------------|
| Elementary school and below   | 30 024 (29.45) | 35 297 (34.62) | 36 638 (35.93) | --           | --          | --      | --         | --        | --                     | --                       | --           | --           | --             | --              | --            | --        | --        | --           | --                                    | --                        | --       | --                       | --                                    | --                        |
| Middle school                 | 16 322 (54.36) | 15 491 (43.89) | 13 995 (38.20) | <0.0001     | --          | --      | --         | --        | --                     | --                       | --           | --           | --             | --              | --            | --        | --        | --           | --                                    | --                        | --       | --                       | --                                    | --                        |
| High school degree or above   | 13 352 (35.84) | 16 936 (35.64) | 12 602 (34.89) | --           | --          | --      | --         | --        | --                     | --                       | --           | --           | --             | --              | --            | --        | --        | --           | --                                    | --                        | --       | --                       | --                                    | --                        |
95% CI, 0.63 to 0.68; fully adjusted HR, 0.76; 95% CI, 0.72 to 0.80) and high school degree or above group (unadjusted HR, 0.51; 95% CI, 0.49 to 0.54; fully adjusted HR, 0.67; 95% CI, 0.63 to 0.70) were associated with relatively lower risks of incident HTN. Similarly, compared with the elementary school and below group, the middle school group (unadjusted HR, 0.83; 95% CI, 0.80 to 0.85; fully adjusted HR, 0.90; 95% CI, 0.87 to 0.92) or high school degree or above group (unadjusted HR, 0.75; 95% CI, 0.73 to 0.78; fully adjusted HR, 0.85; 95% CI, 0.82 to 0.88) had lower risks of uncontrolled HTN.

The incidences of newly diagnosed and uncontrolled HTN in the elementary school and below group were higher than those in the middle or above groups. Compared with the elementary school and below group, in the middle or above school groups, the adjusted HR for the risk of newly diagnosed HTN was lower than that of uncontrolled HTN.

**Subgroup analyses of education level with newly diagnosed HTN and uncontrolled HTN**

We further explored the influence of educational attainment on blood pressure in subgroup analyses (online supplemental table 1). Consistent results of educational attainment with both newly diagnosed HTN and uncontrolled HTN were observed in various subgroups.

**DISCUSSION**

In the present cohort study of 101,959 Chinese participants aged 40 years and older, compared with participants at the elementary school education level and below, those with educational attainment at the middle school level or above exhibited a lower risk of newly diagnosed HTN and achieved better blood pressure control in the follow-up phase. The monitoring and management of blood pressure and its control should be strengthened among middle–elderly subjects, especially those with poor educational attainment. The present study, to our best knowledge, is the largest cohort study to investigate associations of educational attainment with the risk of newly diagnosed HTN and uncontrolled HTN.

Elevated blood pressure remains a major public health challenge because it is one of the most crucial risk factors for premature death worldwide. The prevalence and incidence of HTN has rapidly increased in recent decades in China. Overall, in the present study, the prevalent HTN was 44.9% among adults aged 40 years and older at baseline from 2011 to 2012. In the same age groups, a similar pattern but a much lower rate (32.2%) was found among the American population, even though obesity was much more common in the USA from 2011 to 2014. Disparity of educational attainment could explain the differences in prevalence of HTN among developing and developed countries, even though traditional risk factors for cardiovascular diseases were generally lower in low-income countries. In a recent nationwide cross-sectional study in China, Zheng et al observed a decreased prevalence of HTN and increased HTN control among those with middle and senior education relative to those with only primary education. Accordingly, there is concern about the effects of education levels on the risk of elevated blood pressure in individuals from China, especially among those who have only a below average level of common education. At the same time, information about educational attainment in helping shape the future of blood pressure control remains sparse. We analysed data from the 4C cohort to conduct nationwide comparisons and found that subjects with higher educational attainment had lower incidence rates of HTN and better blood pressure control in the future.

We showed that education has a marked influence on HTN. On the one hand, this influence is reflected in the increase in newly diagnosed HTN in subjects with elementary school education and below. On the other hand, subjects with elementary school education and below were more likely to have difficulty meeting blood pressure control targets. Subjects with elementary school education and below may be associated with more unhealthy lifestyles, such as an unbalanced diet or a lack of exercise, which are associated with the onset of HTN. According to the German National Health Interview and Examination Survey, Finger et al estimated that adults with lower educational attainment consumed calorically dense food more frequently than those with higher educational attainment. Participants with lower education levels are more physically active, which may explain why they consume a high caloric diet. Additionally, patients with HTN with elementary school education and below may show poor compliance with antihypertensive medications. Therefore, our study suggests that the education level of patients with HTN must be considered and the education level of the people as a whole should be increased to avoid HTN and related cerebrovascular diseases.

Our study does have some important limitations. First, the study participants were followed for only a mean of 3.8 years. The relatively short follow-up time could reduce the proportion of new-onset events as well as the statistical power of the study, particularly for evaluating the incidence of HTN. Second, the follow-up rate is important for assessing the validity of the data set for research purposes. However, nearly 12.2% of study participants were lost to follow-up, which is relatively high and may devastate the reliability of the study results. Third, as the study participants had only one follow-up visit, the blood pressures were evaluated only at the baseline and follow-up visits. This could limit the sensitivity of ascertaining outcomes, especially for the timing of diagnoses for incident HTN. Fourth, the patterns of educational attainment have shown extreme imbalance and differences in regional development in China. Among all of the participants in the study, only 9972 (9.78%) had a bachelor’s degree or higher. Therefore, the findings could not be generalised to all participants worldwide.

In the present Chinese nationwide population-based cohort study, participants with elementary school education levels and below exhibited excess risks of newly diagnosed HTN and worse blood pressure control compared with individuals with educational attainment at the middle school level or above. In our study, the communities included high education and high economy communities, such as Shanghai, Beijing and Guangzhou, while poor economy communities such as Lanzhou were also included. Thus, the study was representative to analyse the effect of educational levels on HTN incidence and control in Chinese populations. Our findings emphasise the value of promoting the overall management of educational attainment in preventing HTN and related cardiovascular events, particularly in countries where education is less accessible.

**Author affiliations**

1Department of Endocrinology, Sun Yat-Sen University, Guangzhou, China
2Department of Endocrinology, Sun Yat-Sen Memorial Hospital, Guangzhou, China
3Shanghai Jiao Tong University Medical School Affiliated Ruijin Hospital, Shanghai, China
4Department of Endocrinology, Chinese People’s Liberation Army General Hospital, Beijing, China
5Department of Endocrinology, Shandong Provincial Hospital, Jinan, Shandong, China

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What is already known on this subject

▶ Educational attainment can improve life expectancy and guide healthy behaviours throughout the entire lifetime. A nationwide longitudinal study of the association of the educational status with the risk of hypertension (HTN) and its control in China is lacking.

What this study adds

▶ In the present cohort study of 101,959 Chinese participants aged 40 years and older, compared with participants at the elementary school education level and below, those with educational attainment at the middle school level or above exhibited a lower risk of newly diagnosed HTN and achieved better blood pressure control in the follow-up phase. Our findings emphasise the importance of promoting the overall management of the educational attainment in the prevention of HTN and related cardiovascular events, particularly in countries where education is less accessible.

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