INTRODUCTION

Stroke is a common cardiovascular and cerebrovascular disease with a high incidence rate, high mortality rate, and high disability rate. Among all cardiovascular and cerebrovascular disease deaths, stroke ranks second and brings a substantial economic burden to the world. According to a nationwide survey in China, stroke is the leading cause of death, and the incidence rate continues to
rise. Therefore, it is essential to identify the high-risk population of a stroke at an early stage. As far as we know, hypertension and stroke are continuously and independently positively correlated. A prospective observational study showed that for every 20 mmHg increase in SBP or 10 mmHg increase in DBP, the risk of cerebrovascular disease doubles. Asia Pacific Cohort Studies Collaboration (APCSC) showed that the relationship between blood pressure rise and stroke events in the Asian population is more robust than that in Australia and New Zealand. For every 10 mmHg increase in SBP, the stroke risk in the Asian population increases by 53%, while that in Australia and New Zealand increases by 24% and 21%, respectively. At the same time, hypertension has many people in our country, about 245 million. Thus, it was still necessary to find new detection methods to identify and prevent stroke in hypertensive patients.

Ankle–brachial index (ABI), the ratio of ankle arterial pressure and brachial arterial pressure, was obtained by measuring the posterior tibial artery’s systolic pressures or anterior tibial artery and brachial artery at ankle. A noninvasive method for diagnosis and detection of lower limb arterial diseases, and it was inexpensive and easy to use. A study has shown that ABI could also be used as a screening method for recurrent stroke. Moreover, some previous studies have shown that the decrease of ABI is related to the increase of stroke risk, but other studies have not found such a relationship. Simultaneously, the nonlinear relationship between ABI and stroke has not been discussed in the above studies.

Therefore, our current study aimed to evaluate the relation of ABI with the prevalence of stroke and any possible effect modifiers among hypertensive patients without atrial fibrillation (AF), considering that a small proportion of patients with AF can cause stroke, through the use of data from the China H-type Hypertension Registry Study to address this gap, as mentioned earlier in knowledge.

2 | METHODS

2.1 | Subject population and design

The present study was a subset of the China H-type Hypertension Registry Study (Registration number: ChiCTR1800017274); the design and methods of the China H-type Hypertension Registry Study have been reported elsewhere. The study is a real-world observational study, conducted from Wuyuan, Jiangxi province of China, beginning in July 2018. Eligible participants were men and women aged 18 years and older and were diagnosed with hypertensive. All participants gave written informed consent before data collection. The Ethics Committees of the Biomedical Institute of Anhui Medical University approved study protocols.

For this cross-sectional analysis, we included 14 268 patients at their final visit. After excluding subjects without hypertension (n = 34), and those with missing information on ABI (n = 3328), as well as with participants with AF (n = 156), considering that a small proportion of patients with atrial fibrillation can cause stroke, the final analysis included 10 750 subjects (Figure S1).

2.2 | Data collection

According to a standard protocol, physical examination and medical history were conducted for all participants. Physical examination included weight, height, waist circumference (WC), and hip circumference (HC). BMI was calculated as the body weight in kilograms divided by the square of the height in meters (kg/m²). Hypertension was defined as seated resting systolic blood pressure (SBP) ≥ 140 mmHg or diastolic blood pressure (DBP) ≥ 90 mmHg at both the screening and recruitment visit, or who were on an antihypertensive medication. Diabetes mellitus was defined as a self-reported physician diagnosis of diabetes or FBG concentration of ≥ 7.0 mmol/L or the use of glucose-lowering drugs.

2.3 | Ankle–brachial index measurement

The ankle–brachial index (ABI) was automatically measured in the supine position after having rested for more than 10 minutes, using Omron Colin BP-203RPE III device (Omron Health Care), and ABI was calculated for each leg by dividing the SBP obtained at the ankle level in the respective leg by the SBP of the brachial artery. We calculated the ABI of each leg, and the lower value of the ABI was used in the final analysis.

2.4 | Laboratory tests

Blood samples, included total serum bilirubin (TBIL), plasma homocysteine (Hcy), fasting total cholesterol (TC), triglycerides (TG) and high-density lipoprotein cholesterol (HDL-C), fasting blood glucose (FBG), liver enzymes (aspartate aminotransferases [AST], alanine transaminase [ALT]), and serum creatinine, were collected from all participants. The Chronic Kidney Disease Epidemiology Collaboration (CKD-EPI) equation was performed to calculate the estimated glomerular filtration rate (eGFR). All laboratory measurements met a standardization and certification program. All of these parameters were measured using automatic clinical analyzers (Beckman Coulter) at the Biaojia Biotechnology Laboratory, Shenzhen, China.

2.5 | Outcomes

The outcome was stroke (ischemic or hemorrhagic), excluding strokes caused by transient ischemic attacks (TIA), cranioencephalic trauma, and intracranial tumors. In this study, the medical history of stroke was a self-reported stroke, which was mainly collected through a questionnaire. By asking each participant whether there was a stroke, if so, when the stroke occurred, the symptoms at that time, what kind of treatment, and whether there were relevant medical records included discharge summary and imaging data.
2.6 | Covariates

These selected covariates are closely related to stroke. Categorical variables consisted of sex (males or females), smoking status (never, former, or current), drinking status (never, former, or current), and medication use (antihypertensive drugs [yes or no], glucose-lowering drugs [yes or no], and lipid-lowering drugs [yes or no]). Continuous variables included age, SBP, DBP, pulse, BMI, and laboratory results (FBG, TC, TG, low-density lipoprotein cholesterol [LDL-C], Hcy, AST, ALT, uric acid, TBiL, and eGFR).

2.7 | Statistical analysis

Continuous variables were described using means with standard deviations (SDs) and median with interquartile range, and categorical variables were presented as count (percentage). Student's t-test and chi-square test were performed to compare groups for continuous variables and categorical variables. Participants were divided into four groups based on participants by quartiles of ABI levels. Multivariate logistic regression was used to assess the odds ratios (ORs) and 95% confidence intervals (CI) for stroke in ABI (as a continuous variable, quartiles, and a categorical variable), without and with adjustment for age, sex, SBP, DBP, pulse, smoking status, drinking status, BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs. A generalized additive model and a smooth curve fitting (penalized spline method) were performed to characterize the shape of the relationship between ABI and stroke. If nonlinearity was detected, we first use a recursive algorithm to calculate the inflection points and then construct a two-segment binary logistic model on both sides of the inflection points. Exploratory analyses were performed to ensure the robustness of the result. Possible modifications of the association between ABI (as a continuous variable) and the risk of stroke were assessed for variables including sex (males vs. females), age (≥65 vs. <65 years), BMI (≥24 vs. ≥24 kg/m²), smoking status (never vs. former vs. current), and drinking status (never vs. former vs. current). A two-tailed p less than .05 was considered to be statistically significant.

All statistical analyses were performed using the R software (R Foundation; www.empowerstats.com; X&Y Solutions, Inc, Boston, MA, USA) and the statistical package (R) (http://www.R-project.org, The R Foundation).

3 | RESULTS

3.1 | Study participants and baseline characteristics

As shown in Table 1, the average age of our study population is 63.81 ± 9.23 years, ranging from 27 to 96 years, 690 (6.42%) had a stroke, 532 (4.95%) had coronary heart disease (CHD), and 1938 (18.03%) had diabetes mellitus. Baseline characteristics of our study participants, stratified by quartiles of ABI, were comparable between the four groups. Participants with lowest levels of ABI (Q1) had lower value in age, uric acid, had lower rates of drinker and smoker, had higher rates of CHD and stroke, and higher value in SBP, DBP, pulse, TC, TG, HDL, LDL, and eGFR (all p < .01). There were no statistically significant between the four groups in terms of BMI, Hcy, FBG, diabetes mellitus, AST, ALT, TBiL, antihypertensive drugs, glucose-lowering drugs, or lipid-lowering drugs (all p > .05).

3.2 | Associations between ABI and stroke

We constructed three models to evaluate the relationship between ABI and stroke. These three models were crude model (adjusted for none), model 1 (adjusted for age, sex, SBP, DBP, pulse, smoking status, drinking status), and model 2 (adjusted for all variables in model 1 plus adjusted for BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs), respectively. Continuous variables (ABI) were changed into quartiles and a categorical variable for sensitivity analysis. As illustrated in Table 2, ABI was negatively correlated with the prevalence of stroke (per SD increment; adjusted OR, 0.88; 95% CI, 0.82-0.94). Compared with participants in Q1, a lower prevalence of stroke was found in participants in Q2 (OR, 0.71; 95% CI: 0.56-0.90), Q3 (OR, 0.87; 95% CI: 0.70-1.08), and Q4 (OR, 0.81; 95% CI: 0.65-1.01), respectively. However, the decline of the latter two groups was not significant. To draw the real relationship between ABI and stroke, a generalized additive model and penalized spline method was performed, which showed that there is no nonlinear relationship between ABI and stroke; after adjusting age, sex, SBP, DBP, pulse, smoking status, drinking status, BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs, as shown in Figure 1.

Conversely, a significantly higher prevalence of stroke (OR, 1.26; 95% CI: 1.05-1.50) was found in participants in quartile 1 (<1.05) compared with participants in quartiles 2-4 (≥1.05).

3.3 | Subgroup analyses

To assess the relationship between ABI and stroke in various groups, we performed exploratory subgroup analyses (Figure 2). A significantly stronger inverse association between ABI (quartiles 2-4, ≥1.05 vs. quartile 1, <1.05) and the risk of stroke was observed in the elderly over 65 years old (≥65 vs. <65 years, p for interaction = .012). However, the rest of the subgroup variables included...
sex (males vs. females), BMI (< 24 vs. ≥24 kg/m²), smoking status
(never vs. former vs. current), and drinking status (never vs. for-
mer vs. current). After adjustment for age, sex, SBP, DBP, pulse,
smoking status, drinking status, BMI, FBG, TC, TG, LDL-C, Hcy,
AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-
lowering drugs, and lipid-lowering drugs, except for the stratifying
variables, there were no significant interactions in above variables
(all p for interactions >.05).

4 | DISCUSSION

In a cross-sectional study of a large sample population, the fully ad-
justed smooth curve fitting demonstrated an L-shaped correlation
between ABI and stroke among Chinese adults with hypertension
without atrial fibrillation, and the inflection point of ABI was cal-
lculated to be 1.05. Moreover, a strong negative correlation between
ABI and stroke can be observed in the elderly over 65 years old.

| Characteristics | Total | ABI     | p Value |
|----------------|-------|---------|---------|
|                |       | Q1 (<1.05) | Q2 (1.05-1.10) | Q3 (1.10-1.15) | Q4 (≥1.15) |
| N              | 10 750 | 2643 | 2452 | 2786 | 2869 | <.001 |
| Males, N       | 5072 (47.18%) | 1076 (40.71%) | 1024 (41.76%) | 1351 (48.49%) | 1621 (56.50%) | <.001 |
| Age, year      | 63.81 ± 9.23 | 64.46 ± 10.28 | 63.05 ± 9.24 | 63.14 ± 8.80 | 64.53 ± 8.48 | <.001 |
| BMI, kg/m²     | 23.60 ± 3.81 | 23.57 ± 3.89 | 23.62 ± 3.47 | 23.71 ± 4.32 | 23.49 ± 3.48 | .189 |
| Smoking status, N (%) | 2844 (26.46%) | 691 (26.15%) | 581 (23.69%) | 757 (27.17%) | 815 (28.41%) | <.001 |
| Drinking status, N (%) | 2458 (22.87%) | 496 (18.77%) | 513 (20.93%) | 681 (24.44%) | 768 (26.77%) | <.001 |

| Laboratory data | | | |
| Hcy, μmol/L     | 18.00 ± 11.05 | 18.49 ± 11.89 | 17.77 ± 11.18 | 17.76 ± 10.93 | 17.97 ± 10.20 | .057 |
| FBG, mmol/L     | 6.17 ± 1.60 | 6.19 ± 1.65 | 6.14 ± 1.54 | 6.15 ± 1.61 | 6.17 ± 1.60 | .663 |
| TC, mmol/L      | 5.15 ± 1.11 | 5.23 ± 1.14 | 5.16 ± 1.10 | 5.15 ± 1.09 | 5.06 ± 1.12 | <.001 |
| TG, mmol/L      | 1.78 ± 1.24 | 1.81 ± 1.18 | 1.84 ± 1.38 | 1.79 ± 1.26 | 1.69 ± 1.16 | <.001 |
| HDL-C, mmol/L   | 1.59 ± 0.44 | 1.61 ± 0.44 | 1.61 ± 0.43 | 1.60 ± 0.45 | 1.55 ± 0.42 | <.001 |
| LDL-C, mmol/L   | 2.99 ± 0.81 | 3.10 ± 0.84 | 3.02 ± 0.79 | 2.98 ± 0.80 | 2.89 ± 0.80 | <.001 |
| AST, U/L        | 26.79 ± 16.78 | 26.81 ± 24.37 | 26.61 ± 14.52 | 26.88 ± 14.09 | 26.85 ± 11.66 | .941 |
| ALT, U/L        | 20.47 ± 17.02 | 20.37 ± 23.31 | 20.35 ± 14.47 | 20.73 ± 15.74 | 20.39 ± 12.89 | .819 |
| TBiL, median (IQR), μmol/L | 13.30 (10.30-17.40) | 13.30 (10.30-17.40) | 13.10 (10.20-17.40) | 13.40 (10.30-17.30) | 13.40 (10.30-17.70) | .370 |
| Uric acid, mmol/L | 414.93 ± 120.89 | 411.84 ± 121.05 | 404.88 ± 120.47 | 413.41 ± 120.82 | 427.84 ± 120.15 | <.001 |
| eGFR, ml/min/1.73 m² | 88.77 ± 20.38 | 87.56 ± 22.78 | 90.65 ± 19.44 | 90.03 ± 18.88 | 87.04 ± 20.03 | <.001 |

| Comorbidities, N (%) | Stroke | CHD | Diabetes mellitus a | PAD | | |
|---------------------|--------|-----|---------------------|-----|--------|
|                     | 690 (6.42%) | 197 (7.45%) | 130 (5.30%) | 181 (6.50%) | 182 (6.34%) | .020 |
|                     | 532 (4.95%) | 148 (5.60%) | 95 (3.87%) | 133 (4.77%) | 156 (5.44%) | .018 |
|                     | 1938 (18.03%) | 494 (18.69%) | 406 (16.56%) | 507 (18.20%) | 531 (18.51%) | .181 |
|                     | 343 (3.19%) | 343 (12.98%) | 0 (0.00%) | 0 (0.00%) | 0 (0.00%) | <.001 |

| Medication use, N (%) | Antihypertensive drugs | Glucose-lowering drugs | Lipid-lowering drugs |
|-----------------------|------------------------|------------------------|--------------------|
|                       | 7044 (65.53%) | 562 (5.23%) | 370 (3.44%) |
|                       | 1758 (66.54%) | 137 (5.18%) | 100 (3.78%) |
|                       | 1619 (66.03%) | 119 (4.85%) | 79 (3.22%) |
|                       | 1818 (65.25%) | 154 (5.53%) | 100 (3.59%) |
|                       | 1849 (64.45%) | 152 (5.30%) | 91 (3.17%) |

| Abbreviations: ABI, ankle-brachial index; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BMI, body mass index; CHD, coronary heart disease; DBP, diastolic blood pressure; eGFR, estimated glomerular filtration rate; FBG, fasting blood glucose; Hcy, homocysteine; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; SBP, systolic blood pressure; TBiL, total bilirubin; TC, total cholesterol; TG, Triglycerides. |

aDiabetes mellitus was defined as self-reported physician diagnosis of diabetes or FBG concentration ≥ 7.0 mmol/L or use of glucose-lowering drugs.
As far as we know, most previous studies had explored the relationship between ABI and stroke. Meves et al.\(^{10}\) conducted an open, prospective, noninterventional cohort study, including 6880 patients (age ≥ 65 years), and the result showed that low ABI (ABI < 0.9) independently raised risk of stroke (HR, 1.6; 95% CI, 1.1-2.2), during the 5-year follow-up. A Honolulu Heart Program of 2767 males aged 71-93 years showed that follow-up for 3-6 years, ABI is inversely related to stroke incidence.\(^{11}\) A Framingham Study included 251 men and 423 women (mean age: 80 years) and followed up for four years, and evaluating the relationship between low ABI (ABI < 0.9) and stroke, coronary heart disease and death, reported that a statistically significant increase in the risk of stroke or transient ischemic attack in persons with a low ABI (hazards ratio, 2.0; 95% CI, 1.1-3.7).\(^{12}\) A recent meta-analysis involving ten studies also showed that a lower ABI level is a risk predictor for increased stroke.\(^{21}\) However, the ARIC study\(^{13}\) showed that the lowest group (ABI < 0.80) had a hazard ratio of 1.93 (95% CI 0.78 to 4.78) in the risk of stroke. Newman et al.\(^{14}\) noted that low ABI (<0.9) was not associated with stroke, whether it is the population with CVD at baseline (RR 1.39 [95% CI: 0.85, 2.25]) or the population without CVD at baseline (RR 1.12 [95% CI: 0.74, 1.70]). The above study’s conclusions were not consistent. Moreover, none of the previous studies had assessed the nonlinear relationship between ABI and stroke.

In the current study, we have a new and more in-depth understanding of the relationship between ABI and stroke. There is an L-shaped correlation between ABI and stroke, not a simple linear correlation or irrelevance. The inflection point of ABI was calculated to be 1.05. The result showed that on the left side of the inflection point, the prevalence of the stroke gradually increased with the decreased ABI level; however, the stroke’s prevalence no longer decreased with the increased ABI level in the right of an inflection point. At the same time, the subgroup analysis of this study suggests that there is a strong negative correlation between the ABI and stroke in the elderly aged 65 or older. Some researcher has confirmed that arterial stiffness is increasing with the aging process.\(^{22}\) Not surprisingly, older adults are more likely at higher risk of stroke.\(^{21}\) The reasons for these inconsistent findings might be research design. Specifically, previous studies only discussed the relationship between PAD and stroke, and only used ABI as an indicator to diagnose PAD, but did not evaluate the relationship between ABI and stroke. However, this study uses ABI as a continuous variable to evaluate its relationship with stroke.

However, the actual situation is that the prevalence rate of PAD is meager (<5%) in the general population.\(^{23}\) The prevalence of PAD was only 3.19% in the hypertensive population in our study. In the current study, we use ABI as a continuous variable to analyze its relationship with stroke. We found an L-shaped association between ABI and stroke, indicating that the low ABI cutoff point marking an increased stroke prevalence was about 1.05 in Chinese hypertensive population. As a possible reflection of the damage caused by the comprehensive

| ABI          | N     | Events (%) | Stroke OR (95% CI), p Value |
|--------------|-------|------------|----------------------------|
|              |       |            | Crude model   | Model 1      | Model 2      |
| Per SD increment | 10,750 | 690 (6.42%) | 0.87 (0.81, 0.93) <.001 | 0.87 (0.81, 0.93) <.001 | 0.88 (0.82, 0.94) <.001 |
| Quartiles     |       |            | 1 | 1 | 1 |
| Q1 (<1.05)    | 2,643 | 197 (7.45%) | 0.70 (0.55, 0.87) .002 | 0.70 (0.55, 0.88) .003 | 0.71 (0.56, 0.90) .004 |
| Q2 (1.05-1.10) | 2,452 | 130 (5.30%) | 0.86 (0.70, 1.06) .167 | 0.86 (0.70, 1.07) .175 | 0.87 (0.70, 1.08) .208 |
| Q3 (1.10-1.15) | 2,786 | 181 (6.50%) | 0.84 (0.68, 1.04) .104 | 0.78 (0.63, 0.97) .026 | 0.81 (0.65, 1.01) .058 |
| Q4 (≥1.15)    | 2,869 | 182 (6.34%) | 0.80 (0.68, 0.95) .013 | 0.79 (0.66, 0.94) .007 | 0.80 (0.66, 0.95) .014 |
| Categories    |       |            | 1 | 1 | 1 |
| Q1 (<1.05)    | 2,643 | 197 (7.45%) | 1.24 (1.05, 1.48) .013 | 1.27 (1.07, 1.52) .007 | 1.26 (1.05, 1.50) .014 |
| Q2 - Q4 (≥1.05) | 8,107 | 130 (5.30%) | 1 | 1 | 1 |

Note: Crude model was adjusted for none. Model 1 was adjusted for age, sex, SBP, DBP, pulse, smoking status, and drinking status. Model 2 was adjusted for all variables in Model 1 plus adjusted for BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBil, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs.

Abbreviations: ABI, ankle-brachial index; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BMI, body mass index; DBP, diastolic blood pressure; eGFR, estimated glomerular filtration rate; FBG, fasting blood glucose; Hcy, homocysteine; LDL-C, low-density lipoprotein cholesterol; SBP, systolic blood pressure; TBil, total bilirubin; TC, total cholesterol; TG, Triglycerides.
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action of risk factors. ABI can be used as a useful clinical tool for risk stratification of hypertensive patients, not just as a diagnostic criterion for PAD. Furthermore, Previous studies have shown that a lower ABI level is closely related to systemic atherosclerosis load.\textsuperscript{24,25} This provides a shred of strong evidence for ABI to predict stroke.

Both lower (ABI $\leq 0.90$) and higher (ABI $> 1.40$) ABI levels are independently related to all-cause mortality and cardiovascular disease (CVD) in clinical practice.\textsuperscript{26-28} However, in our study, there are only six patients with ABI $> 1.4$, and the sample size is too small to achieve statistical efficiency.

The potential limitations of our study should also be noted. First, this is an observational study, and we cannot wholly exclude uncontrolled confounding or reverse causality. Second, as a cross-sectional study, we cannot obtain the causal relationship between ABI and stroke. Lastly, this study was conducted in Chinese adults with hypertension; whether the results can be extended to other populations remains discussed.

5 | CONCLUSIONS

In summary, our results suggest that an L-shaped association between ABI and the prevalence of stroke was found in general hypertensive patients, with a cutoff point at about 1.05. A significantly stronger inverse association between ABI ($\geq 1.05$ vs. $< 1.05$) and the risk of stroke was observed in the elderly over 65 years old. The current study demonstrated that ABI measurement is rapid and straightforward, and can be carried out in the general practice of stroke screening, especially in hypertensive subjects, to reduce the risk of stroke.

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CONFLICT OF INTEREST
The authors declare that they have no conflict of interest.

AUTHOR CONTRIBUTIONS
YMS participated in the literature search, data analysis, and data interpretation, and wrote the manuscript. LHH extracted and collected data. LHH, MHL, CCD, WZ, TW, and LJZ conceived the study and participated in its design and coordination. HHB and XSC participated in the study design and provided critical revision. All authors read and approved the final manuscript.

INFORMED CONSENT
Informed consent was obtained from all individual participants included in the study.
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FIGURE 2 Stratified analysis for the stroke and ankle-brachial index (ABI). *Each subgroup analysis adjusted for age, sex, SBP, DBP, pulse, smoking status, drinking status, BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs, except for the stratifying variable.

| Subgroup     | N  | ABI ≥ 1.05 Event,n (%) | ABI < 1.05 Event,n (%) | OR(95%CI) | P for interaction |
|--------------|----|------------------------|------------------------|-----------|-------------------|
| Sex          |    |                        |                        |           |                   |
| males        | 5072 | 295 (7.38%)            | 97 (9.01%)             | 1.22 (0.94, 1.57) | 0.518             |
| females      | 5678 | 198 (4.82%)            | 100 (6.38%)            | 1.32 (1.02, 1.71) |                   |
| Age, years   |    |                        |                        |           |                   |
| <65          | 6083 | 246 (6.08%)            | 69 (5.60%)             | 1.01 (0.75, 1.36) | 0.012             |
| ≥65          | 4665 | 247 (6.09%)            | 128 (9.08%)            | 1.55 (1.22, 1.96) |                   |
| BMI, Kg/m2   |    |                        |                        |           |                   |
| <24          | 5281 | 282 (6.16%)            | 114 (7.56%)            | 1.22 (0.96, 1.55) | 0.653             |
| ≥24          | 5469 | 211 (5.98%)            | 83 (7.31%)             | 1.28 (0.97, 1.69) |                   |
| Smoking status|    |                        |                        |           |                   |
| Never        | 6182 | 236 (5.16%)            | 106 (6.60%)            | 1.33 (1.04, 1.70) | 0.833             |
| Former       | 1723 | 131 (9.50%)            | 39 (11.34%)            | 1.21 (0.80, 1.83) |                   |
| Current      | 2844 | 126 (5.85%)            | 52 (7.53%)             | 1.20 (0.83, 1.72) |                   |
| Drinking status|    |                        |                        |           |                   |
| Never        | 6278 | 304 (6.09%)            | 127 (7.32%)            | 1.28 (1.03, 1.61) | 0.977             |
| Former       | 1562 | 119 (10.35%)           | 50 (12.14%)            | 1.20 (0.82, 1.76) |                   |
| Current      | 2458 | 69 (3.52%)             | 20 (4.03%)             | 1.20 (0.71, 2.05) |                   |

0 1 2

FIGURE 2 Stratified analysis for the stroke and ankle-brachial index (ABI). *Each subgroup analysis adjusted for age, sex, SBP, DBP, pulse, smoking status, drinking status, BMI, FBG, TC, TG, LDL-C, Hcy, AST, ALT, uric acid, TBiL, eGFR, antihypertensive drugs, glucose-lowering drugs, and lipid-lowering drugs, except for the stratifying variable.
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**SUPPORTING INFORMATION**

Additional supporting information may be found online in the Supporting Information section.

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