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## 1 Incidence and Burden of Cardiovascular Disease Attributable to

### 2 Extreme Heat in China

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## Abstract

- 5 Aim The increasing frequency of heat events driven by climate change poses a serious
- 6 challenge to cardiovascular health. This study aimed to investigate the relationship between
- 7 high temperature and acute cardiovascular disease (CVD) incidence in China, and to
- 8 quantify the heat-related burden of CVD.
- 9 Methods 856,357 incident acute CVD cases were used from CVD surveillance in China
- in 2023. A distributed lag non-linear model was applied to evaluate the dose-response
- 11 relationship between temperature and CVD incidence, as well as the best linear unbiased
- 12 prediction of the minimum incidence temperature (MIT). The heat-related burden was
- 13 quantified using attributable fraction and attributable number for two temperature ranges:
- 14 all heat and extreme heat.
- 15 Results Higher temperatures were associated with an increased risk of acute CVD
- incidence. Extreme heat exposure resulted in a cumulative relative risk (RR) of 1.17 (95%)
- 17 confidence interval[CI] 1.05-1.30), with the highest cumulative risk observed on the fourth
- day following exposure. All heat exposure accounted for 3.19% of CVD cases, while
- 19 extreme heat contributed to 0.08%. The burden was particularly pronounced among
- individuals aged  $\geq$  65 years (RR = 1.20, 95% CI 1.05-1.37), rural populations (RR = 1.18,
- 21 95% CI 1.01-1.37), and those living in temperate monsoon and temperate continental
- climates (RR = 1.25, 95% CI 1.05-1.49). MIT varied geographically, ranging from 16.0 °C
- 23 in northern regions to 26.2 °C in southern regions, with the highest MITs concentrated in
- 24 tropical areas.
- 25 Conclusions These findings emphasize the urgent need for region-specific public health
- 26 strategies that integrate climate change adaptation and CVD prevention to mitigate the
- 27 growing health risks associated with rising temperatures.

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# Lay summary

- 3 This study examined the relationship between high temperatures and acute cardiovascular
- 4 disease (CVD) incidence in China and quantified the heat-related burden.
- 5 Higher temperatures increased the risk of acute CVD, with extreme heat exposure
- leading to a 17% higher risk, peaking on the fourth day after exposure.
- 7 The heat-related burden was more pronounced among older adults (≥65 years), rural
- 8 populations, and residents of the northern region, highlighting the need for targeted
- 9 public health strategies.
- 10 Keywords: Attributable fraction, Cardiovascular disease, Environmental exposure,
- 11 Extreme heat, Incidence and burden

Introduction

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- 14 Significant public health achievements in cardiovascular disease (CVD) prevention have
- been achieved in recent decades, largely attributed to interventions targeting traditional risk
- 16 factors such as high blood pressure, elevated cholesterol, and smoking 1-3. Despite this
- progress, the burden of CVD remains substantial, particularly with the increasing trends of
- urbanization and aging populations. In 2022, CVD was responsible for an estimated 19.8
- 19 million deaths worldwide, resulting in 396 million years of life lost (YLLs) and an

additional 44.9 million years lived with disability (YLDs)<sup>4</sup>. Entering the era of precision medicine, further reductions of the CVD burden require expanding prevention strategies beyond conventional risk factors to incorporate nontraditional ones, such as environmental exposures, which can provide novel insights into CVD prevention. Due to increased greenhouse gas emissions from human activities, global warming has become increasingly severe. This lead to a significant rise in both the frequency and the intensity of extreme

heat<sup>5</sup>, imposing a substantial burden on cardiovascular health.

- High temperature can lead to physiological changes, including increased heart rate, heightened blood viscosity, and dehydration due to excessive sweating, which collectively place additional strain on the heart and increase the risk of CVD<sup>6</sup>. Numerous studies have documented heat-related CVD mortality, consistently showing trends that align with established physiological mechanisms<sup>7-10</sup>. With the shift in CVD prevention strategies toward early health interventions, it becomes even more critical to explore the relationship between high temperature and CVD incidence, and the burden on the healthcare system should also be measured in terms of incidence. However, relatively few studies have investigated the relationship between high temperature and the risk of incident CVD<sup>11-13</sup>, and even fewer have quantified heat-related incidence burden in terms of attributable fraction (AF) or attributable number (AN)<sup>14</sup>, particularly across multiple counties in China. Furthermore, the effect of ambient temperature on CVD hospitalizations varies by population characteristics<sup>12</sup>. This highlights the importance to better understand heat-related risks in the context of CVD and underscores the need for targeted, population-specific prevention strategies.
- To address these gaps, we use the most up-to-date and comprehensive registration data to investigate the relationship between heat exposure and acute CVD incidence in China. Moreover, we calculate the AF and AN, which serve as burden measures that extend

- 1 beyond RR by estimating the actual impact of heat exposure on population health. These
- 2 measures are critical for informing targeted interventions and policies to ultimately
- 3 strengthen resilience against the health impacts of climate change.

#### Methods

#### Data sources

This study was based on the CVD surveillance in China, a nationwide surveillance launched in 2021. This system covered approximately 238 million people, accounting for 16.9% of the population of China, and aimed to assess the incidence of acute CVD among Chinese residents aged 18 years and older. All medical institutions within the surveillance network are required to report CVD cases using a standardized case report form. In-hospital CVD cases—including those from inpatient, emergency, and outpatient/clinic settings—are recorded by the responsible clinicians or trained staff at each institution. For fatal out-of-hospital CVD events and cross-regional medical incidents, certified physicians from primary healthcare institutions conduct on-site investigations during routine public health activities to complete case reports. Additionally, annual mortality registration data are cross-referenced with the surveillance database, with subsequent verification and supplementary data collection conducted also by primary care physicians. More detailed information can be found in the Supplementary Material.

Major cardiovascular outcomes were monitored, classified using the International Classification of Diseases, 10th Revision (ICD-10) coding system, ensuring precise and standardized categorization. Stroke encompassed a spectrum of conditions, ranging from subarachnoid hemorrhage to ischemic stroke, broadly classified under codes I60, I61, I63,

- 1 and I64, while explicitly excluding code I62. Coronary heart diseases (CHD) with accurate
- 2 and reliable diagnoses were selected, including acute myocardial infarction, angina
- 3 pectoris and sudden cardiac death. Acute myocardial infarction was classified as codes I21-
- 4 I22, and angina pectoris was represented by code I20, specifically highlighting cases
- 5 treated with percutaneous transluminal coronary angioplasty (PTCA), stent implantation,
- 6 and/or coronary artery bypass grafting (CABG). Sudden cardiac death was categorized
- 7 under code I46.1. Cases with chronic or recovering CVD were explicitly excluded.
- 8 All districts were defined as urban areas, and all counties, including county-level cities and
- 9 banners, were defined as rural areas. The classification was based on the participant's
- 10 habitual residence, defined as having lived in the location for six months or more.
- 11 To ensure data reliability, the quality of each monitoring site was rigorously evaluated
- 12 through assessments of validity, completeness, uniqueness, and comparability. Data for
- cases diagnosed between January 1, 2023, and December 31, 2023, were extracted from
- the database. After data quality control, a total of 856,357 CVD onset cases were included
- in this study.

## Environmental exposure

- We acquired daily meteorological observations in 2023, including maximum, minimum,
- 18 and mean temperatures, along with relative humidity, from over 2,400 meteorological
- 19 stations via the China Meteorological Data Sharing Service System (Figure S1). Bilinear
- 20 interpolation was applied to estimate specific daily environmental exposures for each
- 21 participant. We also evaluated the results using other interpolation methods, including
- 22 bicubic and inverse distance weighting. The Pearson correlation coefficients between these
- 23 results and those obtained using bilinear interpolation were all close to 1, and statistically
- significant at the 99% confidence level. The extreme heat was defined as the 95th percentile

- 1 of daily maximum temperatures. In addition, daily air pollution data for particulate matter
- 2 less than 2.5 μm in diameter (PM<sub>2.5</sub>) and maximum 8-h average ozone in 2023 were sourced
- 3 from the Environmental Monitoring Centers of each county. The surface elevation data
- 4 was obtained from ETOPO 1 Arc-minute global relief model<sup>15</sup>.

#### Statistical analysis

- 6 This study employed a two-stage approach to examine the relationship between high
- 7 temperature and the incidence of acute CVD. In the first stage, we applied a distributed lag
- 8 nonlinear model (DLNM)<sup>16</sup> to assess the county-specific effects of daily maximum
- 9 temperatures on acute CVD incidence. A cross-basis function has been refined to integrate
- 10 exposure-response and lag-response associations. To address overdispersion, we employed
- 11 quasi-Poisson regression models, adjusting for confounders such as PM<sub>2.5</sub>, relative
- humidity, seasonality, and day-of-week effects. A natural cubic spline with four degrees of
- freedom (df) was used for both temperature exposure and lag effects.
- 14 In the second stage, risk estimates from individual counties were pooled to obtain an overall
- estimate. A two-level random-effects model was constructed to capture both between- and
- 16 within-province variability by nesting counties within their respective provinces. We
- 17 computed best linear unbiased predictions (BLUP)<sup>17</sup> at the county level using the
- multilevel model. This approach enables geographically clustered counties within the same
- 19 province to share information effectively with each other.
- 20 The BLUP values were used to determine the MIT for each county and at the national level.
- 21 The MIT corresponds to the temperature associated with the lowest risk of CVD incidence,
- 22 providing critical insights for identifying vulnerable populations and guiding public health
- 23 interventions aimed at mitigating heat-related health risks.

- 1 To examine the delayed effects of high temperature, we analyzed lag periods ranging from
- 2 lag 0-2 to lag 0-7 days, and used the lag period with the largest cumulative effect in the
- 3 model. Furthermore, we performed stratified analyses to explore the differential effects of
- 4 high temperature on various demographic groups. Specifically, we compared the impacts
- 5 of extreme heat across genders, age groups, urban versus rural populations, and climate
- 6 zones.
- 7 A series of sensitivity analyses were also performed to assess the robustness of the findings.
- 8 First, we tested alternative definitions of extreme heat, using the 90th, 92.5th and 99th
- 9 percentiles of daily maximum temperatures. Second, given that previous studies have
- shown that ozone<sup>18</sup> and altitude<sup>19</sup> may modify the effect of temperature on CVD, we
- additionally introduced ozone and altitude as linear terms in the main model. Furthermore,
- we varied the df for meteorological variables from 3 to 5 to check if there was any change
- in the results.
- To quantify the heat-related burden of incident acute CVD, we followed the methodology
- that Gasparrini<sup>20</sup> developed to calculate AF and AN. AF was computed as

$$AF_x = 1 - \exp(-\beta x)$$

17 and AN was calculated as

$$AN_x = n \cdot AF_x$$

- where n is the total number of CVD onset cases and  $\beta x$  represents the exposure-response
- 20 coefficient for a given temperature  $\mathcal{X}$ . Two temperature ranges were considered: 1) all
- 21 heat, referring to the temperature effects above the MIT, and 2) extreme heat, referring to
- 22 temperatures above the 95th percentile of the daily maximum temperature distribution.
- 23 Sub-group analysis was performed for AF, and empirical CIs (95%) were calculated

- 1 through 1,000 Monte Carlo simulations, assuming a multivariate normal distribution of the
- $2 \quad BLUP^{21}$ .
- 3 All statistical analyses in this study were carried out using the "dlnm" and "mixmeta"
- 4 packages in R Version 4.4.0. Continuous variables were described as means  $\pm$  standard
- 5 deviation (SD), and categorical variables as frequencies and percentages. All tests were
- 6 two-sided, and a *P*-value of less than 0.05 was considered statistically significant.

#### 7 Results

## 8 Demographic Characteristics

- 9 In this study, we analyzed 856,357 incident acute CVD cases. The majority of cases were
- male, comprising 58.82% (n = 503,780). In terms of age distribution, 68.65% (n = 587,973)
- were elderly (aged ≥65 years). The overwhelming majority cases were of Han ethnicity,
- accounting for 97.60% (n = 835,907). Almost half (48.97%, n = 410,682) were from rural
- areas. The cardiovascular outcomes monitored included acute stroke (79.36%, n =
- 14 679,436) and acute CHD (20.64%, n = 176,921)(Table 1).
- 15 Given China's extensive geographical diversity, cases were distributed across various
- climate zones with distinct characteristics. Nearly half (41.92%, n = 358,892) occurred in
- 17 temperate monsoon climate, characterized by moderate temperatures and seasonal rainfall.
- Another 54.12% (n = 463,267) occurred in subtropical monsoon climate, marked by hot
- 19 and humid summers. A smaller proportion of cases were located in tropical monsoon
- 20 climate (0.98%, n = 8,399) and in temperate continental climate (3.00%, n = 25,688).
- 21 In China, across all study counties in 2023, the mean daily maximum temperature was
- 22 22.01°C (ranged from -30.03°C to 41.09°C), with mean relative humidity of 70.64%, PM<sub>25</sub>
- 23 concentration of 31.30μg/m³, and ozone concentration of 97.86μg/m³ (*Table 2*).

## 1 Lag pattern of the incidence response at extreme heat

- 2 Figure 1 illustrated the lag effects of extreme heat on CVD incidence, with the
- 3 corresponding effect estimates provided in *Table S1*. The cumulative RR of CVD incidence
- 4 exhibited an initial upward trend, peaking around lag 0-4 days, and then gradually declined.
- 5 At lag 0-6 days, the RR was 1.10 (95% CI 0.95-1.27), indicating that the elevated risk
- 6 associated with extreme heat exposure persists for up to one week. These findings
- 7 suggested that the impact of elevated temperatures on CVD incidence was acute, with the
- 8 highest risk occurring within the initial few days following exposure.

## 9 Temperature-incidence relationship cumulated over 4 days

- 10 As shown in Figure 2, the pooled relationship between temperature and acute CVD
- 11 incidence demonstrated a non-linear association, cumulated over a 4-day lag period. As the
- 12 temperature exceeded approximately 30°C, a notable rise in risk was observed. The pooled
- 13 RR of CVD incidence associated with extreme heat (95th percentile vs. MIT, around
- 14 38.9°C vs.19.3 °C) was 1.17 (95% CI 1.05-1.30). Subgroup analysis further showed that
- the RR of CVD incidence due to extreme heat was higher in individuals aged 65 years and
- above (RR = 1.20, 95% CI 1.05-1.37, p<0.01), female (RR = 1.19, 95% CI 1.01-1.40,
- p=0.78), the rural population (RR = 1.18, 95% CI 1.01-1.37, p=0.03), and those living in
- 18 Temperate Monsoon/Continental climates (RR = 1.25, 95% CI 1.05-1.49, p=0.04),
- 19 compared to their respective counterparts (Figure 2 and Figure 4A).

## Geographical distribution of the MIT ( $\mathcal{C}$ )

- 21 The geographical distribution of the MITs varied in China (Figure 3), ranging from 16.0 to
- 22 26.2 °C. The MIT distribution showed a clear increasing trend from north to south, with all
- values exceeding 24°C south of the Tropic of Cancer. This spatial pattern highlighted the

- 1 variation in temperature-related health risks across different climatic zones in China, with
- 2 highest MITs concentrated in tropical regions with warmest climates.

### Burden of acute CVD incidence attributable to all heat and extreme heat

- 4 In 2023, the AF of acute CVD incidence due to all heat exposures was 3.19% (95% CI
- 5 1.34%-4.98%), which corresponded to 27,298 CVD onset cases, resulting in a large heat
- 6 burden. Regarding extreme heat, we found that 0.08% (95% CI 0.02%-0.15%) of the total
- 7 sample occurred during the 5% hottest days of the year, representing 685 CVD onset cases.
- 8 Figure 4 illustrated that rural residents and the elderly were more affected by high
- 9 temperature and experienced a higher health burden of CVD both during all heat and
- 10 extreme heat (Figure 4 and Table S2).

## 11 Sensitivity Analyses

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- 12 Sensitivity analyses were performed to validate our main findings. The association between
- 13 extreme heat and acute CVD incidence remained consistent when using alternative
- 14 thresholds to define extreme heat, specifically by replacing the 95th percentile of daily
- maximum temperature with the 90th, 92.5th and 99th percentiles. Additionally, altering the
- 16 df for meteorological variables from 3 to 5, or extending the main model to include ozone
- and altitude as covariates to account for potential confounding, did not change the findings
- substantially, further indicating the robustness of the results(*Table S3*).

#### Discussion

- 20 This study was a national-level investigation in China to investigate the relationship
- 21 between heat exposure and acute CVD incidence and estimated the heat burden across
- 22 diverse geographical regions. The study also examined the MIT for CVD incidence,

1 accounting for both cumulative and delayed health impacts. Our primary findings 2 demonstrate that extreme heat significantly elevates the risk of acute CVD incidence, with 3 the risk peaking within the first four days following exposure. We also found that the heatrelated risk and burden varied by age, and urbanization level. These results advanced the 4 5 existing body of knowledge by emphasizing the broader health impacts of extreme heat exposure on incidence. Importantly, our findings highlighted the urgent need for tailored 6 7 public health interventions, especially in light of ongoing climate change. 8 Our findings regarding the association between high temperatures and CVD aligned closely with previous research from different climate zones. Achebak and colleagues 12 conducted 9 10 a quantitative analysis on the association between heat and CVD hospitalizations in Spain, 11 showing that elevated temperatures slightly increased the risk of hospitalization for diseases of arteries, arterioles, and capillaries (RR: 1.104 [1.038 to 1.175]) and ischemic 12 heart diseases (RR: 1.083 [1.052 to 1.114]). Although the demographic characteristics of 13 patients varied, Achebak's study mainly focused on Whites with a lower proportion of 14 elderly (aged ≥65 years), at 18.9%. Similarly, Tao and colleagues<sup>22</sup> observed that excess 15 hot days were associated with an increased risk of CVD in China (RR: 1.10 [1.05 to 1.15]). 16 17 However, Tao's study was limited to data solely from the largest hospitals in 15 cities, excluding rural areas and other provinces. 18 Previous studies on heat-related health burden have mainly focused on mortality and all-19 20 cause incidence, while few studies so far have specifically examined the heat-related health 21 burden of CVD incidence<sup>14</sup>. Concentrating on all-cause outcomes could dilute the heat-22 incidence relationship, making it challenging to detect the specific effects of heat<sup>23</sup>. Our 23 study extended this understanding by demonstrating that total and extreme heat contributed

to 3.19% and 0.08% of CVD incidence, respectively. Bai and colleagues<sup>24</sup> reported that in

Ontario, Canada, 1.2% of CHD hospitalizations and 1.8% of stroke hospitalizations were

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due to heat exposure, with 0.16% and 0.19% attributable to extreme heat. However, their analysis focused solely on hospitalizations (both new and preexisting cases), excluding outpatient settings such as emergency visits. Another study in Shenzhen, a southern Chinese city, found that 2.5% of first-ever stroke incidence was attributable to heat<sup>25</sup>, underscoring the significant burden of heat on CVD in urban areas. These emphasizes the importance of calculating the AF and AN, which, by translating individual risk (represented by RR) into population-level impact, could be indispensable for public health service evaluation and planning.

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Another key finding of this study was the geographic variation in MIT, showing a northsouth gradient that likely reflects human physiological<sup>26-28</sup>, behavioral and cultural adaptations to the local climate<sup>29</sup>. Individuals in tropical regions, accustomed to consistently warmer climates, generally exhibit higher MITs, while those in northern areas were more susceptible to the heat's effects. This corresponds with our subgroup analysis, which showed that populations in northern climate zones had higher risk compared to those in more southern climate zones. Although no previous studies have specifically examined the MIT for CVD, similar trends were observed in studies of the minimum mortality temperature. A study found that countries with hotter climates or close proximity to the equator tend to have higher minimum mortality temperatures, suggesting a degree of population adaptation to local climatic conditions<sup>30</sup>. Yin and colleagues<sup>31</sup> analyzed 420 locations and found that minimum mortality temperatures closely aligned with the most frequently observed local temperatures during the same period. This correlation highlights how long-term climate exposure shapes physiological and behavioral adaptations to environmental conditions. Recognizing these geographic differences highlights the importance of incorporating regional climate characteristics into public health strategies. Our findings confirmed that extreme heat exerts acute and short-term lag effects on cardiovascular health. Consistent with existing literature from China, we observed that the

- 1 heat effect on CVD was immediate and can persist for up to approximately one week
- 2 following exposure<sup>32,33</sup>. This lag effect indicated that individuals might remain at
- 3 heightened risk for CVD even after extreme heat had dissipated, underscoring the necessity
- 4 for timely public health interventions during heatwave events and continued monitoring in
- 5 the days afterward.
- 6 The findings of our study were in line with previous meta-analyses<sup>34,35</sup>, which demonstrate
- 7 that the association between heat and CVD varied by population characteristics and
- 8 geographical location. Subgroup analyses further revealed disparities in the effects of
- 9 extreme heat across different demographic and regional groups. Older individuals appeared
- 10 to be at greater risk, likely due to physiological differences in thermoregulation. Elderly
- 11 individuals were more vulnerable because of age-related impairments in temperature
- 12 regulation and the increased prevalence of comorbidities, which amplified their
- susceptibility to heat-related health impacts<sup>29,36</sup>. Rural populations also appeared more
- vulnerable to extreme heat, likely due to limited access to healthcare services and greater
- exposure to outdoor work. Specifically, many rural residents in China are farmers or
- laborers and live in underdeveloped housing<sup>29,37</sup>, which can exacerbate heat exposure. In
- 17 2023, the average number of air conditioning units in rural areas was 105.7 units per 100
- households, significantly lower than the 171.7 units per 100 households in urban areas<sup>38</sup>.
- 19 Our findings are supported by physiological plausibility. Exposure to extreme heat induces
- various stress responses in the cardiovascular system. To dissipate heat, the body activates
- 21 mechanisms such as increased skin blood flow and sweating, which elevate cardiac output
- 22 to maintain blood pressure and support thermoregulation<sup>39,40</sup>. Dehydration caused by
- 23 extreme heat further exacerbates cardiovascular stress by reducing blood volume and
- 24 decreasing the heart's filling pressure, which can lead to ischemia or infarction<sup>41</sup>. Heat
- 25 exposure can trigger an inflammatory response, with mediators like cytokines disrupting

- 1 cellular homeostasis and impairing organ function, thereby exacerbating damage to
- 2 vulnerable tissues and organs<sup>42</sup>. In addition, extreme heat may also impact cardiovascular
- 3 health through indirect behavioral and psychological pathways. The E(e)SEEDi lifestyle
- 4 framework integrates environmental exposures with four core domains of health—Sleep,
- 5 Emotion, Exercise, and Diet<sup>43</sup>. According to this framework, environmental stressors such
- 6 as extreme heat can compromise the quality of these essential lifestyle factors, thereby
- 7 increasing cardiovascular vulnerability.
- 8 Although this study focused on the adverse effects of extreme ambient heat exposure, it is
- 9 important to acknowledge that not all forms of heat exposure are harmful. For instance,
- 10 controlled and time-limited exposures such as Finnish sauna bathing have been associated
- 11 with cardiovascular benefits, including improved vascular function, reduced blood
- pressure, and lower risk of cardiovascular events and mortality<sup>44</sup>.
- 13 This study has several notable advantages. First, it stands as one of the largest studies to
- date to examine the effects of extreme heat on CVD incidence in China, covering multiple
- 15 counties across a range of climate zones. Such extensive cases, combined with broad
- 16 geographic coverage, allow for a more comprehensive and robust assessment of the
- 17 relationship between high temperature and CVD risk. Secondly, this study explored MIT
- 18 variations across different climate zones, providing novel insights into CVD prevention
- 19 and serving as a valuable reference for region-specific public health interventions.
- 20 Additionally, few studies have conducted subgroup analyses that differentiate urban and
- 21 rural populations<sup>6,12,45</sup>, and this study addressed this gap. By identifying specific vulnerable
- groups, such as rural residents and elderly individuals, this study contributed to the design
- of targeted adaptation strategies for more effective health outcomes.
- However, this study has several limitations. First, as an ecological study, this study was
- subject to ecological fallacies, and caution was needed when generalizing the findings to

other contexts or regions. Second, although we controlled for environmental confounding factors such as humidity and air pollution, individual-level factors such as lifestyle and behavior still influence the results. Additionally, since most individuals spent the majority of their time indoors<sup>46,47</sup>, relying on outdoor temperature observations might not fully reflect actual heat exposure at the individual level. Beyond coronary heart disease and stroke, extreme heat may also increase the risk of other major adverse cardiovascular events (MACE), including arrhythmias and heart failure. However, as these conditions are not included in the current scope of the CVD surveillance system in China, they were not assessed in this analysis. Furthermore, the study was based on data from one single year, limiting its ability to assess long-term trends over time. Future research should address these limitations by incorporating more precise individual-level exposure data, exploring compound exposure to other environmental risk factors, using composite biometeorological indicators such as net effective temperature<sup>48</sup>, and examining long-term trends across diverse global contexts to provide a more comprehensive understanding of heat-related cardiovascular risks.

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In conclusion, our study demonstrated that extreme heat significantly elevated the risk of acute CVD incidence in China, with notable variations across geographic regions and demographic groups. These findings underscore the urgent need of implementing targeted measures to mitigate heat exposure, particularly for vulnerable populations. Such as, establishing early warning systems and enhancing public awareness can enable communities to better anticipate and respond to extreme heat events. And, developing tailored interventions to reduce heat exposure for high-risk populations is essential. Together, these strategies provide a comprehensive approach to reduce the adverse health effects of extreme heat on the Chinese population, mitigate CVD risk, and enhance community resilience in the context of ongoing climate change.

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#### **Conflicts of interest**

- 9 The authors declared that they have no known competing financial interests or personal
- relationships that could have appeared to influence the work reported in this paper.

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### **Authors' Contributions**

- 2W had full access to all of the data in the study and takes responsibility for the integrity
- of the data and the accuracy of the data analysis. ZW: conceptualization, data curation,
- methodology, review & editing, supervision, funding acquisition; XP: methodology, formal
- analysis, writing-original draft preparation; HT: methodology and writing-original draft
- preparation; CZ, YZ, MS, QJ, SL, XW, YT, XC, YT, XY, NT, CC, YZ, FL and JY:
- 18 investigation.

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## Data Availability Statement

- 21 The health data underlying this article will be shared on reasonable request to the
- 22 corresponding author. The environmental data used in this study are publicly available from

- 1 their respective sources. The daily meteorological observations from over 2,400
- 2 meteorological stations were obtained from the China Meteorological Data Sharing Service
- 3 System (CMDC, available at http://data.cma.cn). Air pollution data were sourced from the
- 4 Environmental Monitoring Centers of each county in China (available at
- 5 https://quotsoft.net/air/). Surface elevation data were derived from the ETOPO 1 Arc-
- 6 Minute Global Relief Model, provided by the National Centers for Environmental
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1	Figure	legends	
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- 2 Figure 1. Cumulative Lag Effects of Extreme Heat on CVD Incidence Over One Week. RR, relative
- 3 risk. The effect of extreme heat was summarized as the relative risk (RR) of incidence at the 95th
- 4 temperature percentiles vs. the minimum incidence temperature (MIT).
- 5 Figure 2. Temperature-Incidence Relationship and RR of CVD Over a 4-Day Lag Period. (A) Pooled
- 6 Non-Linear Relationship Between Temperature and CVD Incidence. (B) RR of CVD Incidence by
- 7 Gender. (C) RR of CVD Incidence by Age Group. (D) RR of CVD Incidence by Urbanicity. (E) RR of
- 8 CVD Incidence by Climate Zone. RR, relative risk. Dashed vertical lines in each panel indicate the
- 9 threshold for extreme heat, defined as the 95th percentile of daily temperature distribution. Asterisk (\*)
- indicates P < 0.05, and double asterisk (\*\*) indicates P < 0.01 for differences in RR between extreme
- 11 heat and CVD incidence.
- 12 Figure 3. Geographical Distribution of Minimum Incidence Temperature Across China.
- 13 Figure 4. Subgroup Analysis by Age, Sex, Urbanicity and Climate Zone (A) RR and 95%CI of CVD
- 14 Incidence Associated with Extreme Heat. (B) AF and 95%CI of Extreme Heat Exposures. (C) AF and
- 15 95%CI of All Heat Exposures. RR, relative risk. AF, Attributable fraction.

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# 1 Tables

# 2 Table 1. Demographic Characteristics and CVD Outcomes of the Study Population

				D
Characteristics	Female	Male	Total	P-Value
	n=352,577 (%)	n=503,780 (%)	n=856,357(%)	
Age				
18-64	81,159 (23.02)	187,225 (37.16)	268,384 (31.35)	
≥65	271,418 (76.98)	316,555 (62.84)	587,973 (68.65)	< 0.001
Ethnic				
Han	344,968 (97.84)	490,939 (97.45)	835,907 (97.60)	
National minority	7,609 (2.16)	12,841 (2.55)	20,450 (2.40)	< 0.001
Urbanity				
Urban	177,252 (50.27)	268,423 (53.28)	445,675 (52.03)	
Rural	175,325 (49.73)	235,357 (46.72)	410,682 (48.97)	< 0.001
Career				
White collar	12,099 (3.43)	24,180 (4.80)	36,279 (4.23)	
Administrator	3,702 (1.05)	13,075 (2.60)	16,777 (1.96)	
Blue collar	3,864 (1.10)	5,526 (1.10)	9,390 (1.10)	
Other	221,880 (62.93)	306,746 (60.89)	528,626 (61.76)	
Missing	111,032 (31.49)	154,253 (30.62)	265,285 (30.96)	< 0.001
Marital status	,			
Married/remarried /living together	214,743 (60.91)	324,193 (64.35)	538,936 (62.98)	
Separated or divorced	1,481 (0.42)	3,790 (0.75)	5,271 (0.62)	
Single	3,735 (1.06)	12,556 (2.49)	16,291 (1.90)	
Widowed	23,844 (6.76)	12,192 (2.42)	36,036 (4.21)	
Missing	108,774 (30.85)	151,049 (29.98)	259,823 (30.32)	<0.001
Diseases				
CHD	61,447 (17.43)	115,474 (22.92)	176,921 (20.64)	
Stroke	291,130 (82.57)	388,306 (77.08)	679,436 (79.36)	< 0.001

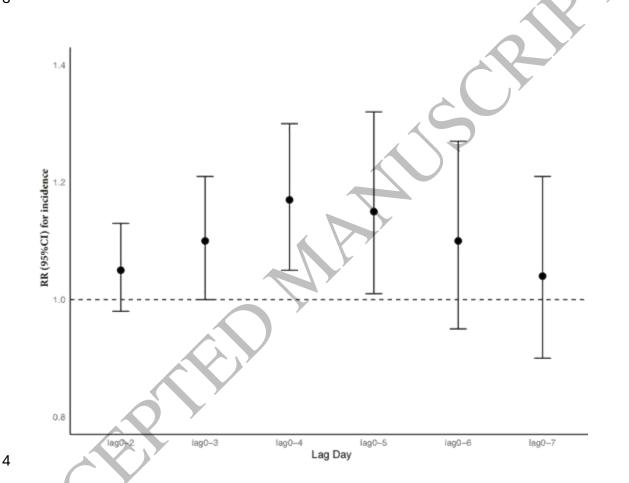
Climate zones			
Temperate Monsoon Climate	150,835 (42.84)	208,057 (41.30)	358,892 (41.92)
Subtropical Monsoon Climate	187,545 (53.33)	275,722 (54.62)	463,267 (54.12)
Tropical Monsoon Climate	3,418 (0.97)	4,981 (0.99)	8,399 (0.98)
Temperate Continental Climate	10,086 (2.86)	15,602 (3.10)	25,688 (3.00) <0.001
			, ,

<sup>1 \*</sup>P-values represent comparisons between groups for each characteristic.

# 2 Table 2. Distributions of Daily Ambient Environment Conditions in China in 2023

Environmental factors	Mean	Minimum	Q1	Q2	Q3	Maximum
Maximum temperature(°C)	21.00	-30.03	14.20	22.91	29.19	41.09
Relative humidity(%)	70.64	5.80	56.29	75.72	87.99	100.00
$PM_{2.5}(\mu g/m^3)$	31.30	1.00	14.50	24.50	39.80	334.90
Ozone(µg/m³)	97.86	5.90	71.80	91.70	120.10	274.00
altitude (m)	232.43	1.70	18.30	49.70	244.10	2524.80





**Figure 1 Cumulative Lag Effects of Extreme Heat on CVD Incidence Over One Week.** RR, relative risk. The effect of extreme heat was summarized as the relative risk (RR) of incidence at the 95th temperature percentiles vs. the minimum incidence temperature (MIT).

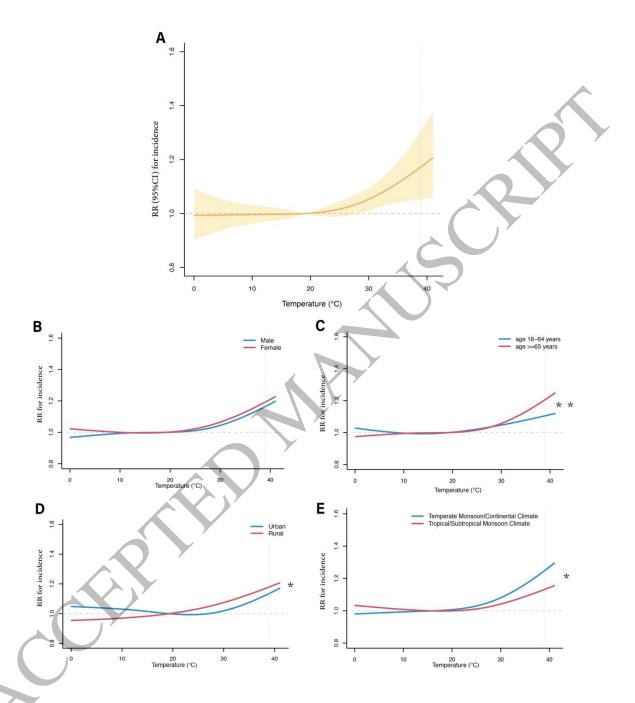
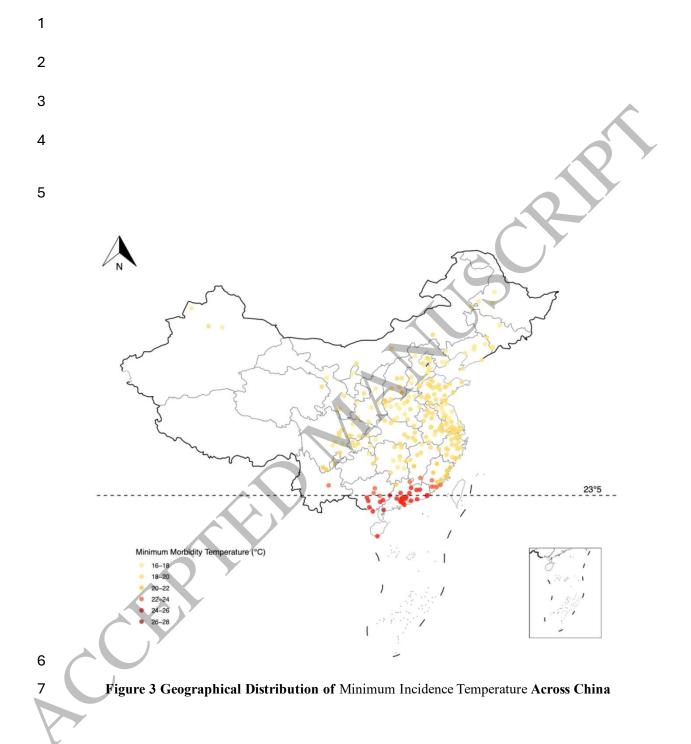


Figure 2 Temperature-Incidence Relationship and RR of CVD Over a 4-Day Lag Period. (A) Pooled Non-Linear Relationship Between Temperature and CVD Incidence. (B) RR of CVD Incidence by Gender. (C) RR of CVD Incidence by Age Group. (D) RR of CVD Incidence by Urbanicity. (E) RR of CVD Incidence by Climate Zone. Dashed vertical lines in each panel indicate the threshold for extreme heat, defined as the 95th percentile of daily temperature distribution. Asterisk (\*) indicates P < 0.05, and double asterisk (\*\*) indicates P < 0.01 for differences in RR between extreme heat and CVD incidence.



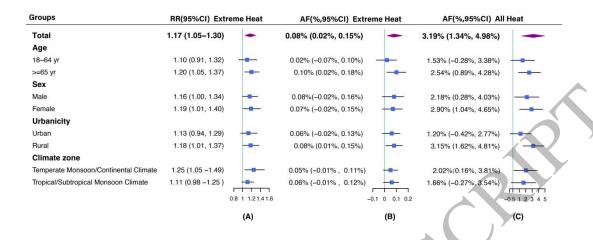
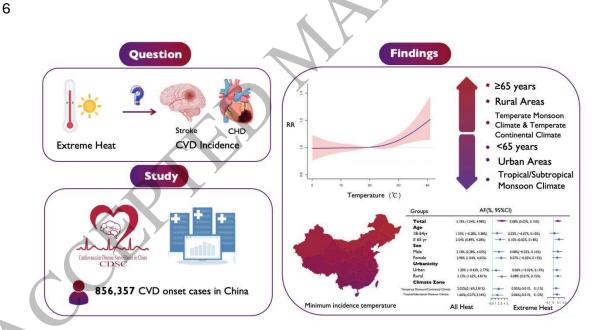


Figure 4 Subgroup Analysis by Age, Sex, Urbanicity and Climate Zone (A) RR and 95%CI of CVD Incidence Associated with Extreme Heat. (B) AF and 95%CI of Extreme Heat Exposures. (C) AF and 95%CI of All Heat Exposures. RR, relative risk. AF, Attributable fraction.



**Graphical abstract**