

Trends in stroke prevalence caused by an elevated BMI in the European population

Keywords

BMI, Europe, Epidemiology, Stroke

Abstract

Introduction

This study investigated the complex relationship between an elevated body mass index (BMI) and stroke risk in Europe amidst rising obesity rates and an aging population.

Material and methods

A joinpoint model was used to fit the data on stroke mortality rates attributed to an elevated BMI. The age period cohort model was used to analyze the effects of age, period, and birth cohort on trends in stroke prevalence related to an elevated BMI from 1990–2019 in the European context. A Bayesian model was used to predict stroke mortality from increased BMI in Europe in 2020-2044.

Results

Stroke mortality rates attributed to an elevated BMI displayed a wave-like declining trend in Europe, decreasing from 21.73/100,000 in 1990 to 14.01/100,000 in 2019. The most substantial decline in stroke mortality rate occurred between 2003 and 2013 (-4.10% , $P < 0.001$). Male mortality rates decreased from 21.48/100,000 in 1990 to 15.45/100,000 in 2019, while female rates declined more significantly from 21.40/100,000 in 1990 to 12.48/100,000 in 2019. Age-standardised stroke mortality due to increased BMI is expected to decline in Europe over the next 25 years.

Conclusions

From 1990 to 2019, stroke mortality rates associated with elevated BMI in Europe steadily increased. However, projections indicate that this trend may shift, with a slight decline expected in the near future. Given these patterns, there is an urgent need in Europe to enhance weight management strategies for stroke patients and increase public health awareness, particularly focusing on men and the elderly, to help reduce stroke-related mortality.

Trends in Stroke Prevalence Caused by an Elevated BMI in the European Population

Abstract

Background: This study investigated the complex relationship between an elevated body mass index (BMI) and stroke risk in Europe amidst rising obesity rates and an aging population.

Methods: A joinpoint regression model was used to fit the data on stroke mortality rates attributed to an elevated BMI. The age period cohort model was used to analyze the effects of age, period, and birth cohort on trends in stroke prevalence related to an elevated BMI from 1990–2019 in the European context. A Bayesian Age-Period-Cohort model was used to project age-standardized stroke mortality attributed to elevated BMI in Europe from 2020 to 2044.

Results: Stroke mortality rates attributed to an elevated BMI displayed a wave-like declining trend in Europe, decreasing from 21.73/100,000 in 1990 to 14.01/100,000 in 2019. The most substantial decline in stroke mortality rate occurred between 2003 and 2013 (age period cohort analysis: -4.10%, 95% *CI*: -4.44 to -3.75; $P < 0.001$). Male mortality rates decreased from 21.48/100,000 in 1990 to 15.45/100,000 in 2019, while female rates declined more significantly from 21.40/100,000 in 1990 to 12.48/100,000 in 2019. Age-standardized stroke mortality due to elevated BMI is projected to decline in Europe over the next 25 years. **Age-standardized stroke mortality due to increased BMI is expected to continue declining in Europe over the next 25 years, with projected reductions of 12.99% in males and 13.01% in females by 2044.**

Conclusion: From 1990 to 2019, stroke mortality rates associated with elevated BMI in Europe steadily increased. However, projections suggest a slight decline in the near future. Given these trends, there is an urgent need to enhance weight management for stroke patients and increase public health awareness, particularly among men and the elderly, to reduce stroke-related mortality.

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29 **1. Background**

30 Stroke, an acute focal neurologic deficit syndrome caused by vascular injury in the
31 central nervous system, is a leading cause of mortality and disability worldwide.
32 Accounting for approximately 6 million deaths annually, stroke ranks as the 2nd
33 leading cause of deaths globally [1, 2]. The burden of stroke in terms of patients,
34 survivors, related deaths, and global disability-adjusted life years is increasing,
35 especially in Europe [3, 4]. Specifically, the European population of 509 million had
36 120,000 new strokes and 530,000 stroke-related deaths in 2017, with projections
37 indicating a substantial increase by 2047 [5].

38 Body Mass Index (BMI) is a well-established independent predictor of stroke risk,
39 even after accounting for age, cardiovascular comorbidities (e.g., hypertension,
40 diabetes), and lifestyle factors (e.g., smoking, physical activity) [6]. A pooled analysis
41 of 97 prospective cohorts found that each 5 kg/m² increase in BMI raised stroke
42 hazard ratios (HR) to 1.18 (95% *CI* 1.14–1.22) after adjusting for various confounders
43 [7]. **Elevated BMI** is likely to contribute to stroke risk through mechanisms such as
44 chronic inflammatory responses and metabolic dysregulation, which may promote the
45 onset of stroke [8].

46 Interestingly, some studies report an 'obesity paradox,' suggesting that individuals
47 with higher BMI may exhibit lower mortality in certain conditions, including stroke.
48 This paradox adds complexity to the relationship between BMI and stroke, indicating
49 that BMI's role in stroke outcomes may be more nuanced than traditionally
50 understood and warrants further investigation. Nonetheless, large-scale studies
51 consistently show a strong association between elevated BMI and increased stroke
52 risk. For example, a meta-analysis of 5,798,826 participants reported that higher BMI
53 (overweight or obese) significantly increases stroke risk, with pooled relative risk (RR)
54 values of 1.25 (95% *CI* 1.16-1.34) and 1.47 (95% *CI* 1.02-2.11) [9].

55 **Age-period-cohort effects have become an essential framework for understanding the**
56 **complex relationship between BMI, stroke, and temporal trends. While age effects on**
57 **stroke mortality are well-documented, the impact of period (time) and cohort (birth**

58 group) effects remains less understood. Recent research suggests that period effects—
59 such as shifts in public health strategies, medical interventions, and lifestyle
60 changes—may explain some of the temporal variations in stroke rates. In addition,
61 cohort effects may help to understand how generational differences in environmental
62 exposures, lifestyle factors, and medical advancements influence stroke risk over
63 time.

64 As elevated BMI rates rise across Europe, understanding the full scope of its role in
65 stroke risk becomes increasingly critical [10]. Therefore, this study utilized
66 Europe-wide stroke and BMI data from the Global Burden of Disease (GBD) database
67 (1990–2019) to examine this association comprehensively [11, 12]. At present,
68 relatively few studies have examined trends in stroke prevalence due to increased
69 BMI in Europe, and traditional descriptive analyses of age-specific morbidity or
70 mortality data at different time periods have not been able to eliminate or control for
71 interactions between age, period, and cohort factors. The Jointpoint regression (JPR)
72 model is widely used to analyze the time trend of morbidity and mortality and the
73 burden of disease, which can better reflect the change of epidemic trend and its
74 impact. Age-period-cohort model improves the traditional descriptive analysis method
75 to estimate the risk and trend of disease morbidity or mortality while adjusting for age,
76 period and cohort [13].

77 This study aimed to provide a comprehensive analysis of the relationship between
78 elevated BMI and stroke within the European context. By examining trends in stroke
79 incidence and disability-adjusted life years rates (DALYs) associated with high BMI,
80 the JPR model and Age-Period-Cohort analysis were employed to assess the impact of
81 obesity on stroke prevalence. Our findings are intended to inform public health
82 policies and clinical practices, supporting the development of targeted preventive
83 measures and interventions. Such strategies have the potential to reduce the burden of
84 stroke and obesity-related health issues, ultimately improving health outcomes at the
85 population level.

86

87 **2. Data Sources and Methods**

88 2.1 Study Design

89 This population-based observational study was designed to determine the relationship
90 between an elevated BMI and stroke incidence within the European population.
91 Subsequently, data on stroke mortality rates attributed to elevated BMI in the
92 European population aged 20-94 years were collected from the GBD 2019 study. **The
93 selection of this age group is based on the relatively low stroke mortality rates in
94 populations under 20 years of age, as well as the limited data reliability for those over
95 94 years old. Therefore, the age group 20-94 years was chosen to ensure more
96 accurate representation and consistency in the analysis.** The age-period-cohort model
97 was then applied to analyze the risk of stroke mortality caused by an elevated BMI.

98 2.2 Global Burden of Disease

99 **The data for this study were obtained from the GBD 2019 study [14], which provides
100 comprehensive estimates of stroke mortality attributed to an elevated BMI across
101 different regions, including Europe. Specifically, we extracted data on stroke deaths
102 related to high BMI (defined as BMI ≥ 25 kg/m²) for the European population aged
103 20-94 years.**

104 **The GBD 2019 study reports age-standardized stroke mortality rates, as well as
105 estimates of DALYs for different age groups and sex. The stroke mortality data was
106 standardized based on the GBD 2019 world standard population, ensuring consistency
107 in comparison across different populations and time periods.**

108 2.3 Joinpoint Regression Model (JRM)

109 The JRM was selected to identify significant trend shifts over time, making it
110 particularly suitable for analyzing temporal changes in stroke mortality. Unlike
111 traditional models, which may not capture precise points of trend changes, the JRM
112 can detect and quantify specific inflection points in mortality rates attributed to
113 elevated BMI. The JRM was used to fit the data on stroke mortality rates, with
114 statistical significance determined through the Monte Carlo permutation test. This
115 approach allows for the calculation of annual percentage change (APC) and average
116 annual percentage change (AAPC), along with 95% confidence intervals (CIs).
117 Positive APC or AAPC values indicate an increasing mortality trend, while negative

118 values suggest a decline.

119 2.4 Age-Period-Cohort Model

120 To address potential collinearity between period and cohort effects in the
121 Age-Period-Cohort model, we used the intrinsic estimator method, which is designed
122 to manage collinearity among these variables. **This method allows for the independent**
123 **estimation of age, period, and cohort effects, ensuring more accurate and transparent**
124 **analysis of stroke trends over time.** Compared to traditional models, the APC model
125 provides several advantages. It allows for the separation of age, period, and cohort
126 effects, offering a more nuanced understanding of each factor's influence on stroke
127 mortality.

128 The Age-Period-Cohort model is based on the Poisson distribution and analyzes the
129 risk of stroke mortality attributed to an elevated BMI in the European population
130 based on the following: age; period; and cohort. With every 5-year increment grouped
131 as 1 age group, individuals 20–94 years of age resulted in 15 groups. To meet the
132 structural requirement of the model that the age group interval equals the period group
133 interval, the years 1990–2019 were divided into 6 groups, with each interval 5 years
134 long. The birth cohort was calculated as the period minus age, resulting in 20 cohorts.
135 **This model helps us to understand how the effects of BMI on stroke mortality vary**
136 **not only by age but also by temporal factors such as public health changes and**
137 **cohort-specific risk factors.** To understand the linear relationship between age, period,
138 and cohort in the model, the intrinsic estimator method was used to estimate the effect
139 coefficients.

140

141 2.5 Statistical Analysis

142 The APC and AAPC were calculated using Joinpoint Regression Software (version
143 4.9.0.0), which provides advanced methodologies for detecting trend shifts in stroke
144 mortality over time. Age, period, and birth cohorts were each segmented into 5-year
145 intervals to ensure consistency in the Age-Period-Cohort analysis. Effect coefficients
146 and relative risks were calculated with the intrinsic estimator package in Stata 17.0,
147 chosen for its robust capabilities in handling complex regression analyses. Relative

148 risk (RR) was determined as $[RR = \exp(\text{effect coefficient})]$, with a higher RR
149 indicating a greater risk of stroke incidence. Additionally, R software (version 4.2.1)
150 was used to project stroke mortality rates in Europe from 2020 to 2049. Statistical
151 significance was set at $P < 0.05$.

152

153 **3. Results**

154 **3.1 Trends in Stroke Mortality Attributed to an Elevated BMI in the European** 155 **Population**

156 The standardized mortality rate of stroke attributed to an elevated BMI in the overall
157 European population showed a wave-like declining trend (AAPC=-1.47%, 95% CI:
158 -1.86 to -1.08, $P < 0.001$), decreasing from 21.73/100,000 in 1990 to 14.01/100,000 in
159 2019 (Figure1). The most significant decline was observed between 2003 and 2013
160 (APC=-4.10%, 95% CI: -4.44 to -3.75, $P < 0.001$). The standardized mortality rate in
161 males was generally higher than females. The male standardized mortality rate
162 decreased from 21.48/100,000 in 1990 to 15.45/100,000 in 2019 (AAPC=-1.07%,
163 95% CI: -1.67 to -0.66, $P < 0.001$). The decline in the female standardized mortality
164 rate was more substantial (AAPC=-1.84%, 95% CI: -2.23 to -1.44, $P < 0.001$),
165 dropping from 21.40/100,000 in 1990 to 12.48/100,000 in 2019 (Table 1).

166 Figure 1. Age-Standardized Mortality Rate (ASMR) Trends by Gender from 1990 to 2019 Using
167 Joinpoint Regression Models

168 Table 1. Changing trends in standardized stroke mortality rates attributed to an elevated BMI in
169 Europe from 1990–2019

170

171 **3.2 Trends in Stroke Mortality Attributed to an Elevated BMI by Age, Period,** 172 **and Cohort**

173 3.2.1 Age Period Mortality Rate

174 The stroke mortality in the European population 20–94 years of age showed a similar
175 trend in all age groups. In the 20–54-year-old age group, the stroke mortality rate
176 gradually increased for males and females, and after 55–79 years of age, the mortality
177 rate tended to increase. For males and females, the mortality rate was highest in the

178 75–79 age group, decreasing in the 80-84 age group and continuing to increase in the
179 85-94-year-old age group. The RR for stroke mortality in the 75-79 age group was
180 2.8-fold higher compared to the 20-54 age group (95% CI: 2.5 to 3.2, $P<0.001$),
181 highlighting the significant increase in stroke risk with age (Figure 2). Compared to
182 other periods, the mortality was higher for males and females in the period 1990-1994
183 (Figure 2).

184 Figure 2. Age-period changing trends in stroke mortality attributed to an elevated BMI in Europe
185 (A: male; B: female).

186

187 3.2.2 Age-Cohort Mortality Rate

188 Age-specific stroke mortality in the European population tended to decrease as the
189 birth cohort increased. Males and females in the elevated BMI cohort had an
190 increasing, then decreasing trend in stroke mortality after the 55–59-year-old age
191 group. Mortality varied by birth cohort at the same age and this difference was more
192 pronounced in the earlier birth cohorts. In the 90-94-year-old age group, stroke
193 mortality due to an elevated BMI was 1.26-fold higher in males and 1.24-fold higher
194 in females born in 1900 - 1904 compared to those born in 1925 - 1929 (95% CI for
195 males: 1.18 to 1.34, $P<0.001$; 95% CI for females: 1.15 to 1.33, $P<0.001$) (Figure 3).

196 Figure 3. Age-cohort changing trends in stroke mortality attributed to an elevated BMI in Europe
197 (A: male; B: female).

198

199 **3.3 Age-Period-Cohort Analysis of Stroke Mortality Attributed to an Elevated** 200 **BMI in the European Population**

201 3.3.1 Age effect

202 The age effect showed that stroke mortality due to an elevated BMI generally
203 increased with age in European males and females aged 20-79 years, with a
204 decreasing then increasing trend after 80 - 94 years of age. For example, the RR of
205 stroke death in the 75-79-year-old age group was 63.04-fold higher compared to the
206 20-24-year-old age group (95% CI: 60.22 to 66.30, $P<0.001$). This high RR reflects
207 the strong age-related effect of BMI on stroke mortality. The RR for females in the

208 75-79 age group was even higher at 87.38-fold compared to the 20-24 age group
209 (95% CI: 83.29 to 91.56, $P<0.001$).

210 Figure 4. Age effect of stroke mortality attributed to an elevated BMI in Europe.

211 Table 2. Age-period-cohort model analysis of stroke mortality rate attributed to an elevated BMI
212 in Europe

213

214 3.3.2 Period Effect

215 The stroke mortality risk attributed to an elevated BMI for males and females
216 exhibited consistent changes over the years. Except for a decline in the risk between
217 2010 and 2014, the mortality risk for other periods increased progressively over time.
218 Between 2005 and 2009, the relative risk for stroke mortality was 1.38-fold higher for
219 males and 1.26-fold higher for females compared to the 1990 - 1994 period (95% CI
220 for males: 1.32 to 1.45, $P<0.001$; 95% CI for females: 1.18 to 1.34, $P<0.001$) (Figure
221 5, Table 2).

222 Figure 5. Period effect of stroke mortality attributed to an elevated BMI in Europe.

223

224 3.3.3 Cohort effect

225 The cohort effect of an elevated BMI on stroke mortality risk varied significantly
226 across different birth cohorts. Specifically, the RR for stroke death in males born
227 between 1910 and 1914 was 8.29-fold higher than in males born between 1995 and
228 1999. Similarly, the RR for stroke death in females born between 1910 and 1914 was
229 9.10-fold higher than in those born between 1995 and 1999 (Figure 6, Table 2). These
230 differences were statistically significant, with 95% confidence intervals for males
231 (95% CI: 8.01 to 8.57) and females (95% CI: 8.72 to 9.49) confirming the large
232 disparities in stroke mortality across cohorts.

233 Figure 6. Cohort effect of stroke mortality attributed to an elevated BMI in Europe.

234

235 3.4 Prediction

236 According to the Bayesian age-period-cohort model, the age-standardized stroke
237 mortality due to an elevated BMI is expected to gradually decline in Europe from

238 2020–2044. By 2044, the mortality rate for males in Europe is expected to continue to
239 decline, decreasing by 12.99% compared to 2019. Similarly, stroke mortality in
240 females is expected to continue a downward trend over the next 20 years, decreasing
241 by 13.01% compared to 2019 (Figure 7).

242 Figure 7. Prediction of mortality rate of stroke in Europe from 2020–2044(A: male; B: female).

243

244 **4. Discussion**

245 From 1990–2019, the stroke mortality rate attributed to BMI in the European
246 population showed a declining trend. The Age-Period-Cohort model revealed that the
247 mortality rate trends in stroke caused by BMI in European males and females are
248 influenced by age, period, and cohort effects. The joinpoint regression results showed
249 that the stroke mortality rate attributed to an elevated BMI in Europe is generally
250 declining, which was consistent with the overall mortality results in Europe [15]. The
251 most significant decline in stroke-related mortality occurred between 2003 and 2013.
252 Some studies have suggested that widespread changes in healthy behaviors and
253 treatments for these risk factors are the reasons for the sharp decline in vascular
254 mortality rates in high-income countries [15]. The higher mortality rate in males
255 compared to females may be attributed not only to differences in hormone levels but
256 also to lifestyle factors and behavioral patterns. Males are more likely to engage in
257 habits such as smoking and alcohol consumption, which can increase the risk of
258 overweight and obesity, thereby contributing to higher stroke mortality.

259 The age effect showed that the standardized mortality risk for stroke caused by an
260 elevated BMI in the European population 20–79 years of age generally increased with
261 age. The peak mortality for males and females involved the 75–79-year-old age group,
262 which is consistent with the global trend of population growth and aging. As the
263 global population continues to grow and age, the absolute number of stroke deaths has
264 been increasing. The risk of stroke increases with age, and in males and females, the
265 risk of stroke doubles after 55 years of age [16]. Moreover, as age advances and aging
266 accelerates, population immunity decreases and risk factors related to stroke
267 accumulate in the body. Cellular and vascular aging with collagen deposition

268 accelerate the formation of mature atherosclerotic plaques and interact with an
269 elevated BMI, thereby increasing the risk of death [17].

270 Notably, the risk of stroke mortality caused by an elevated BMI in the 80–84-year-old
271 age group gradually decreased with age, showing contradictory results compared to
272 previous studies. The obesity paradox has received much attention in recent years.

273 Several studies have shown that an elevated BMI may have a protective effect on
274 survival after a stroke, a concept known as the obesity paradox [18]. For example, Gu
275 et al. [19] found that obese patients were less likely to experience fatigue during the
276 acute phase of ischemic stroke, suggesting that elevated BMI may provide a
277 protective energy reserve. Similarly, Chaudhary et al. [20] observed that overweight
278 and obese ischemic stroke patients had a significantly lower risk of 1-year mortality
279 compared to those with normal weight. A cross-sectional study in Korea also found
280 that obesity was negatively correlated with adverse outcomes in all stroke patients,
281 and the obesity paradox may vary by stroke subtype [21]. In European populations,
282 elevated BMI may offer protective benefits through greater nutritional reserves and
283 increased energy stores during stroke recovery. Additionally, lifestyle factors such as
284 higher physical activity levels could mitigate the negative effects of obesity on stroke
285 outcomes. Exploring these biological and lifestyle mechanisms in European
286 populations could provide deeper insights into the obesity paradox and its role in
287 stroke survival and recovery.

288 The period effect showed that the risk of stroke mortality caused by an elevated BMI
289 in Europe has shown a trend of first rising, then falling, then rising again over the
290 years, but the overall trend was not significant. The cohort effect showed that the
291 mortality risk for males and females declined with the progression of the birth cohort.
292 The World Health Organization 2015 World Health Statistics report showed that the
293 overall obesity rate for adults was 21.5% for males and 24.5% for females in Europe
294 [22]. An abundance of clinical and epidemiologic evidence indicated that obesity is
295 associated with a wide range of cardiovascular diseases (CVDs). Obesity can directly
296 or indirectly increase the incidence and mortality of CVDs. The direct effects are
297 through cardiovascular system structural and functional adaptations induced by

298 obesity to accommodate the excess weight and the impact of adipose factors on
299 inflammation and vascular homeostasis, leading to a pro-inflammatory and
300 pro-thrombotic environment. Indirect effects are mediated by accompanying CVDs
301 risk factors, such as insulin resistance, type 2 diabetes mellitus, visceral fat,
302 hypertension, and dyslipidemia. Therefore, there should be an emphasis on
303 controlling the overweight and obesity rates, advocating for a healthy weight [22].
304 People from later birth cohorts, with the advancement of economy and technology,
305 pay more attention to pursuing a healthy lifestyle, enhancing health literacy, and
306 maintaining a healthy BMI. Socioeconomic and biological factors can have a
307 significant impact on stroke risk, severity, and outcomes. For example, approximately
308 68% of the variability in stroke incidence rates in different European countries can be
309 explained by differences in the gross domestic product, reflecting the living standards
310 and well-being of the population [23].

311 These findings hold important public health implications. The observed decline in
312 stroke mortality linked to BMI, together with the obesity paradox, suggests the need
313 for a nuanced approach to stroke prevention. Policymakers should promote healthy
314 lifestyle modifications for individuals with elevated BMI, while acknowledging that
315 BMI alone may not fully predict stroke outcomes. Integrating these insights into
316 preventive strategies could inform more effective health policies, especially for
317 high-risk populations.

318 Although this study primarily examines the relationship between BMI and stroke
319 mortality, it is essential to consider the role of socioeconomic factors. Income level,
320 education, and healthcare access likely influence obesity and stroke outcomes across
321 European regions. Individuals with fewer resources may have limited access to
322 preventive care and face barriers to adopting healthier lifestyles, which can impact
323 both BMI and stroke risk. Additionally, other contributors to the decline in stroke
324 mortality, such as advancements in stroke management (e.g., improved treatment
325 protocols and rehabilitation strategies) and expanded healthcare access, likely interact
326 with BMI-related trends, collectively shaping stroke outcomes. Future research
327 incorporating socioeconomic dimensions and other health determinants would deepen

328 our understanding of these complex interactions, providing valuable insights for
329 targeted public health interventions.

330

331 **5. Conclusion**

332 From 1990–2019, the stroke mortality rate caused by an elevated BMI in Europe has
333 continuously increased. Stroke mortality was affected by age, period and cohort.
334 Europe urgently needs to strengthen weight control for stroke patients and raise public
335 health awareness through health education, with a particular focus on males and the
336 elderly to reduce the death burden caused by stroke. There should be an emphasis on
337 strengthening the three levels of stroke prevention and early screening to achieve
338 early detection, control, and treatment to avoid unnecessary waste of public health
339 resources and decreased public health.

340

341 **Limitations**

342 This study's findings, based on the GBD database, may be influenced by inherent
343 biases and data accuracy limitations. We restricted the age range to 20 - 94 years due
344 to low stroke incidence in those under 20 and limited data for those over 94, which
345 affects the generalizability of our results, particularly to younger and older
346 populations. Additionally, we did not analyze visceral vs. peripheral obesity, which
347 may impact stroke risk differently. While we acknowledge the obesity paradox, its
348 underlying mechanisms were not explored in this study, representing a gap in
349 understanding.

350 Furthermore, our stroke mortality projections are based on pre-pandemic data, and the
351 COVID-19 pandemic may have influenced stroke risk through changes in healthcare
352 access and lifestyle behaviors. Post-pandemic data would be valuable for refining
353 these projections. Lastly, the GBD database lacks detailed subgroup data, limiting our
354 ability to conduct more granular analyses of specific populations, such as ethnic
355 minorities or socioeconomically disadvantaged groups.

356

357 **Ethical statement**

358 Ethical approval and consent were not required as this study was based on publicly
359 available data.

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361 We appreciate the work of the GBD study 2019 collaborators. We also want to thank
362 the staff and platforms that provide open data.

363 **Conflict of interest**

364 None declared.

365 **Data availability statement**

366 The data that support the findings of this study are openly available in Global Burden
367 of Disease study 2019 collaborators.

368 **Authors' contributions**

369 MJT and YW organised the whole sentinel project, and made the first draft of the
370 manuscript. MJT, YW, HRC and HP made the design and planning of the project.
371 MJT, YW were responsible for the organisation, sample handling and data
372 management. MJT did the data analysis, HRC did the statistical analysis and editing,
373 and the corresponding author HP did the review. All Authors revised the manuscript
374 critically and approved the final version.

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378

379 **Abbreviations**

380 **AAPC:** Average Annual Percentage Change

381 **APC:** Annual Percentage Change

382 **BMI:** Body Mass Index

383 **CI:** Confidence Interval

384 **CVD:** Cardiovascular Disease

385 **DALY:** Disability-Adjusted Life Year

386 **GBD:** Global Burden of Disease

387 **HR:** Hazard Ratio

388 **JRM:** Joinpoint Regression Model

389 **RR:** Relative Risk

390

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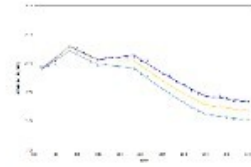
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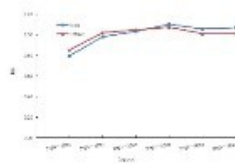
Stroke mortality associated with elevated BMI in Europe



Joinpoint Model

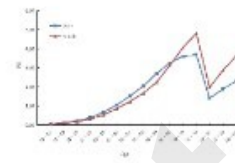


Age effect

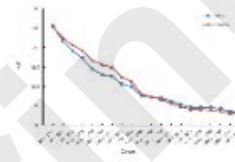


Age-Period-Cohort (APC) Effects

Period effect

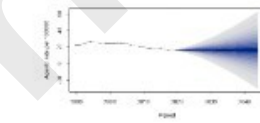


Cohort effect

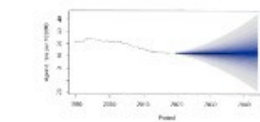


Projection

Male



Female



Preprint

Table 1. Changing trends in standardized stroke mortality rates attributed to an elevated BMI in Europe from 1990–2019

Group	Period	APC (95% CI)	P value	AAPC (95% CI)	P value
Both	1990-1994	4.00(2.72, 5.29)	< 0.001	-1.47(-1.86, -1.08)	< 0.001
	1994-1998	-2.48(-4.37, -0.56)	0.015		
	1998-2003	0.05(-1.18, 1.18)	0.939		
	2003-2013	-4.10(-4.44, -3.75)	< 0.001		
	2013-2019	-1.15(-1.80, -0.50)	0.002		
Male	1990-1994	4.41(3.07, 5.76)	< 0.001	-1.07(-1.47, -0.66)	< 0.001
	1994-1998	-2.38(-4.35, -0.56)	0.023		
	1998-2003	0.68(-0.61, 1.98)	0.284		
	2003-2013	-3.52(-3.87, -3.16)	< 0.001		
	2013-2019	-1.05(-1.73, -0.36)	0.005		
Female	1990-1994	3.63(2.72, 4.95)	< 0.001	-1.84(-2.23, -1.44)	< 0.001
	1994-1998	-2.54(-4.46, -0.58)	0.015		
	1998-2003	-0.48(-1.73, 0.78)	0.430		
	2003-2013	-4.65(-4.99, -4.30)	< 0.001		
	2013-2019	-1.29(-1.96, -0.63)	0.001		

Table 2. Age-period-cohort model analysis of stroke mortality rate attributed to an elevated BMI in Europe

Class	Male				Female			
	Coefficient	Se	P	95% CI	Coefficient	Se	P	95% CI
Age								
20~24	-2.83	0.44	< 0.001	(-3.69,-1.98)	-2.90	0.53	< 0.001	(-3.94,-1.86)
25~29	-2.11	0.28	< 0.001	(-2.67,-1.55)	-2.26	0.36	< 0.001	(-2.96,-1.55)
30~34	-1.51	0.22	< 0.001	(-1.94,-1.09)	-1.66	0.27	< 0.001	(-2.19,-1.13)

35~39	-0.95	0.17	< 0.001	(-1.28,-0.61)	-1.13	0.22	< 0.001	(-1.55,-0.71)
40~44	-0.39	0.14	0.006	(-0.66,-0.11)	-0.61	0.18	< 0.001	(-0.96,-0.27)
45~49	0.06	0.12	0.609	(-0.17,0.29)	-0.13	0.14	0.367	(-0.41,0.15)
50~54	0.42	0.10	< 0.001	(0.23,0.62)	0.22	0.12	0.072	(-0.02,0.45)
55~59	0.72	0.08	< 0.001	(0.56,0.88)	0.52	0.10	< 0.001	(0.33,0.72)
60~64	0.99	0.07	< 0.001	(0.86,1.13)	0.81	0.08	< 0.001	(0.65,0.97)
65~69	1.18	0.06	< 0.001	(1.07,1.30)	1.14	0.07	< 0.001	(1.02,1.27)
70~74	1.28	0.05	< 0.001	(1.18,1.38)	1.40	0.06	< 0.001	(1.29,1.51)
75~79	1.31	0.05	< 0.001	(1.21,1.41)	1.57	0.06	< 0.001	(1.46,1.68)
80~84	0.35	0.07	< 0.001	(0.22,0.47)	0.68	0.07	< 0.001	(0.54,0.82)
85~89	0.64	0.07	< 0.001	(0.50,0.78)	1.05	0.08	< 0.001	(0.89,1.22)
90~94	0.83	0.08	< 0.001	(0.67,1.00)	1.29	0.10	< 0.001	(1.09,1.48)
Period								
1990—1994	-0.23	0.05	< 0.001	(-0.33,-0.13)	-0.16	0.06	0.008	(-0.28,-0.04)
1995—1999	-0.02	0.04	0.609	(-0.09,0.05)	0.02	0.04	0.654	(-0.06,0.10)
2000—2004	0.03	0.03	0.302	(-0.03,0.09)	0.05	0.03	0.118	(-0.01,0.11)
2005—2009	0.10	0.03	0.002	(0.03,0.16)	0.07	0.03	0.024	(0.11,0.13)
2010—2014	0.05	0.04	0.175	(-0.02,0.13)	0.01	0.04	0.808	(-0.08,0.10)
2015—2019	0.07	0.05	0.178	(-0.03,0.17)	0.01	0.06	0.825	(-0.10,0.13)
Cohort								
1900~1904	1.12	0.13	< 0.001	(0.87,1.38)	1.11	0.15	< 0.001	(0.82,1.40)
1905~1909	0.96	0.11	< 0.001	(0.75,1.17)	0.99	0.12	< 0.001	(0.75,1.23)
1910~1914	0.83	0.09	< 0.001	(0.65,1.01)	0.90	0.11	< 0.001	(0.70,1.11)
1915~1919	0.73	0.08	< 0.001	(0.57,0.89)	0.84	0.09	< 0.001	(0.66,1.02)
1920~1924	0.54	0.07	< 0.001	(0.40,0.69)	0.69	0.08	< 0.001	(0.53,0.86)
1925~1929	0.45	0.07	< 0.001	(0.31,0.59)	0.63	0.08	< 0.001	(0.47,0.79)
1930~1934	0.41	0.08	< 0.001	(0.26,0.56)	0.59	0.09	< 0.001	(0.41,0.76)

1935~1939	0.25	0.08	0.003	(0.08,0.41)	0.39	0.10	< 0.001	(0.20,0.59)
1940~1944	0.18	0.09	0.056	(0.00,0.36)	0.30	0.11	0.007	(0.08,0.52)
1945~1949	-0.11	0.11	0.319	(-0.32,0.10)	-0.03	0.13	0.791	(-0.29,0.22)
1950~1954	-0.15	0.12	0.207	(-0.39,0.08)	-0.14	0.15	0.363	(-0.43,0.16)
1955~1959	-0.17	0.14	0.208	(-0.44,0.10)	-0.22	0.17	0.192	(-0.56,0.11)
1960~1964	-0.29	0.15	0.062	(-0.59,0.11)	-0.39	0.19	0.047	(-0.77,-0.01)
1965~1969	-0.45	0.17	0.009	(-0.79,-0.11)	-0.56	0.22	0.012	(-0.99,-0.12)
1970~1974	-0.60	0.20	0.003	(-1.00,-0.21)	-0.69	0.25	0.007	(-1.19,-0.19)
1975~1979	-0.60	0.23	0.010	(-1.06,-0.14)	-0.71	0.30	0.018	(-1.30,-0.12)
1980~1984	-0.60	0.29	0.036	(-1.16,-0.04)	-0.76	0.37	0.043	(-1.49,-0.02)
1985~1989	-0.69	0.39	0.077	(-1.45,0.08)	-0.85	0.51	0.093	(-1.84,0.14)
1990~1994	-0.82	0.59	0.166	(-1.98,0.34)	-0.99	0.77	0.197	(-2.51,0.52)
1995~1999	-0.99	1.24	0.425	(-3.42,1.44)	-1.10	1.53	0.473	(-4.10,1.90)
AIC		6.14				5.967		
BIC		-223.10				-223.726		
Deviance		0.21				0.197		

Multiple Joinpoint Models

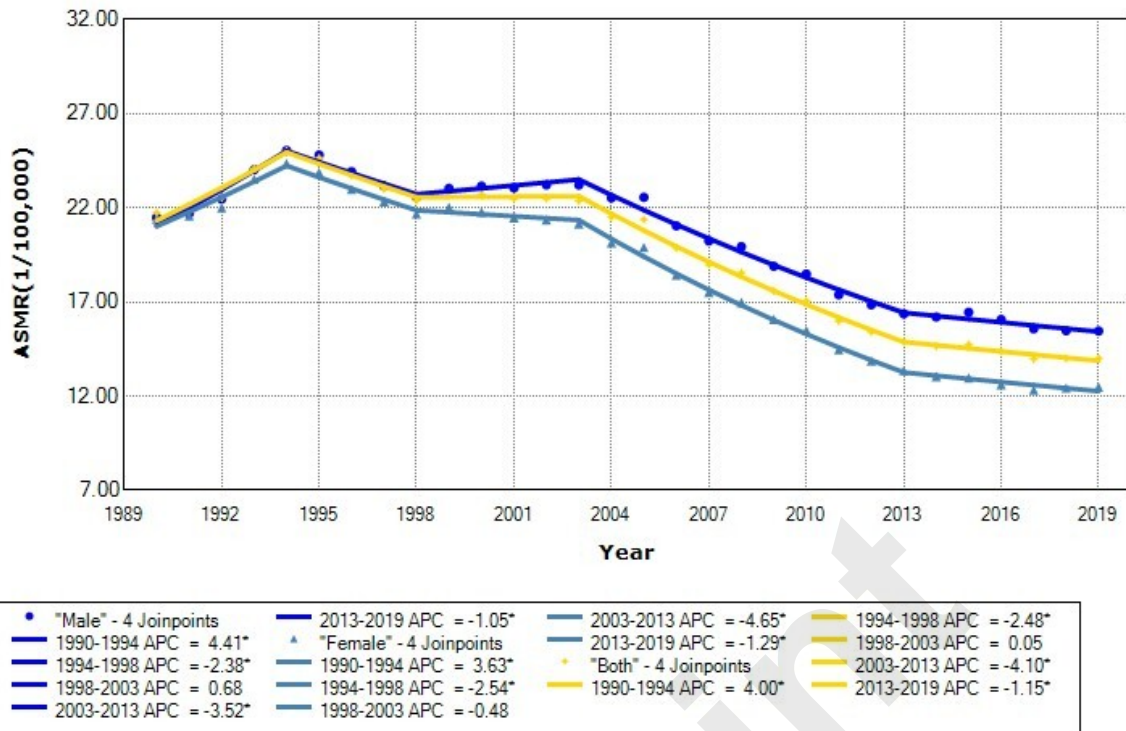


Figure 1. Age-Standardized Mortality Rate (ASMR) Trends by Gender from 1990 to 2019 Using Joinpoint Regression Models

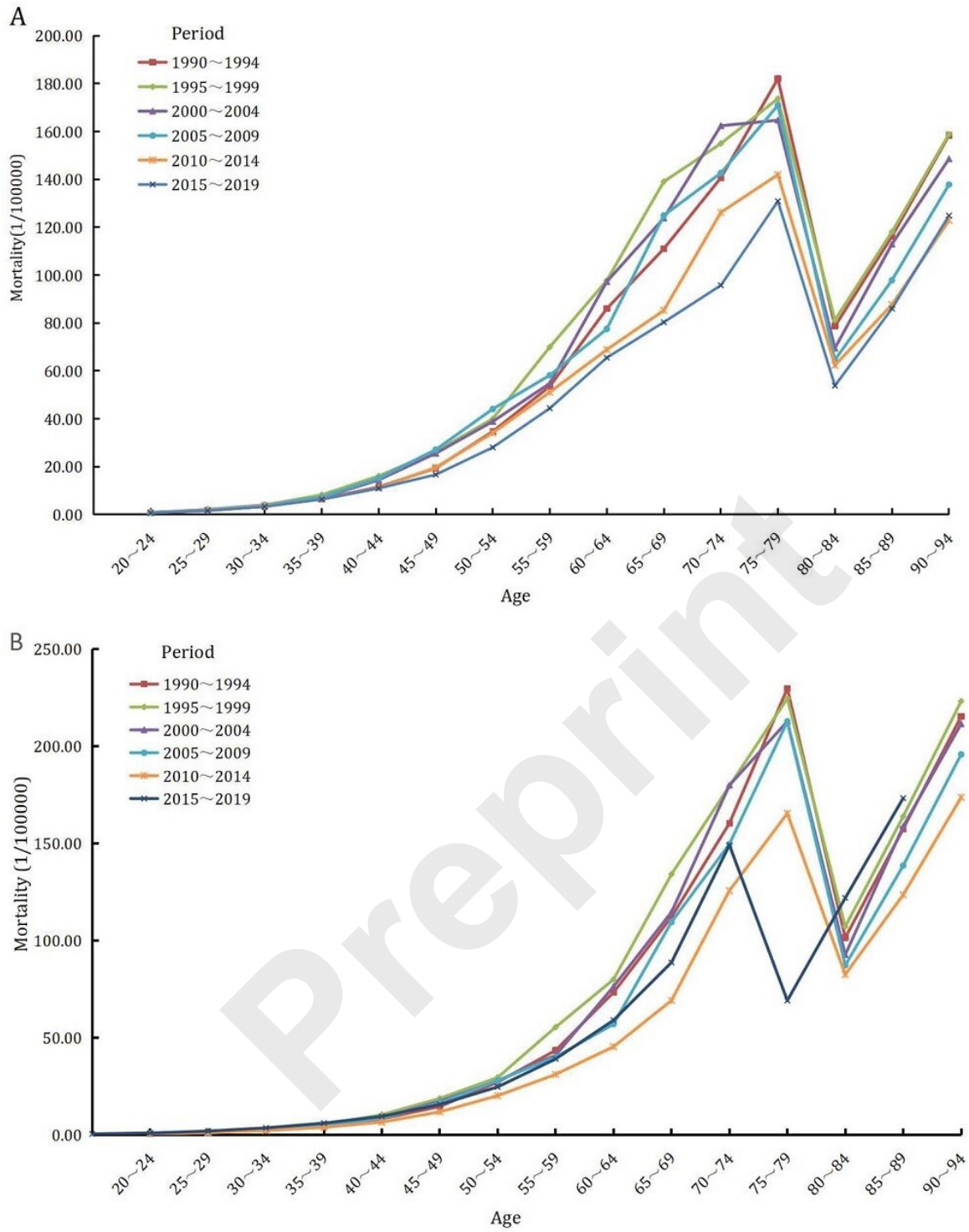


Figure 2. Age-period changing trends in stroke mortality attributed to an elevated BMI in Europe (A: male; B: female)

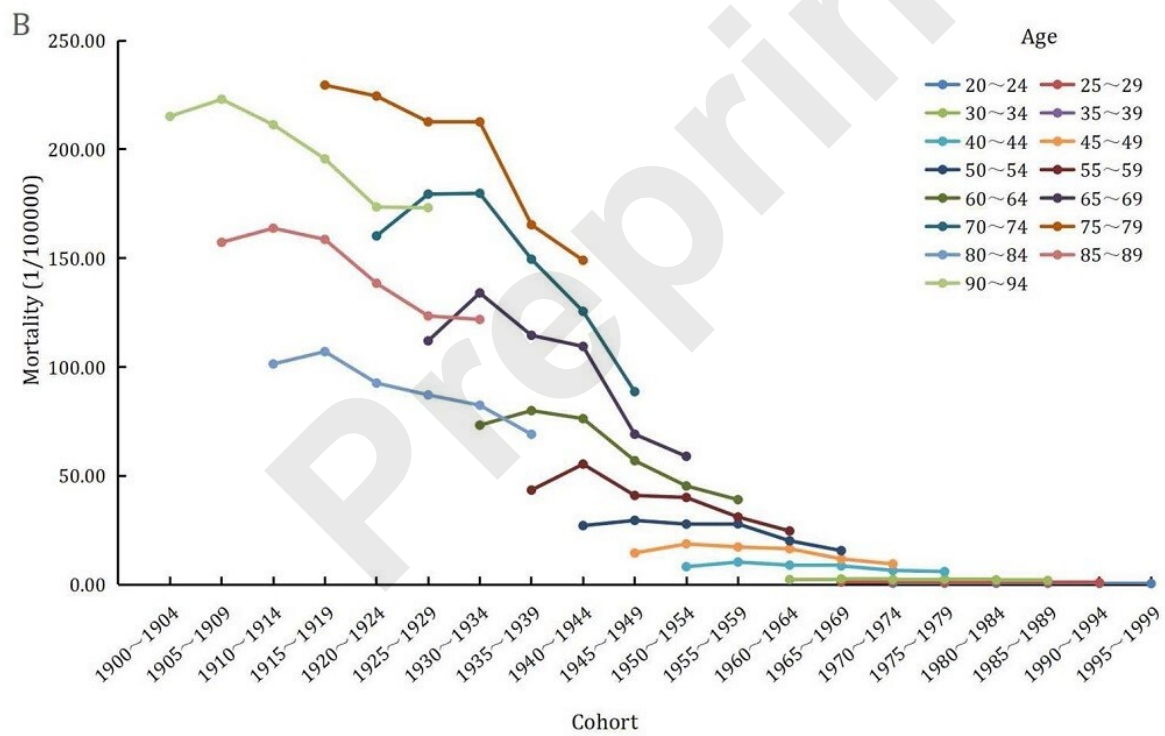
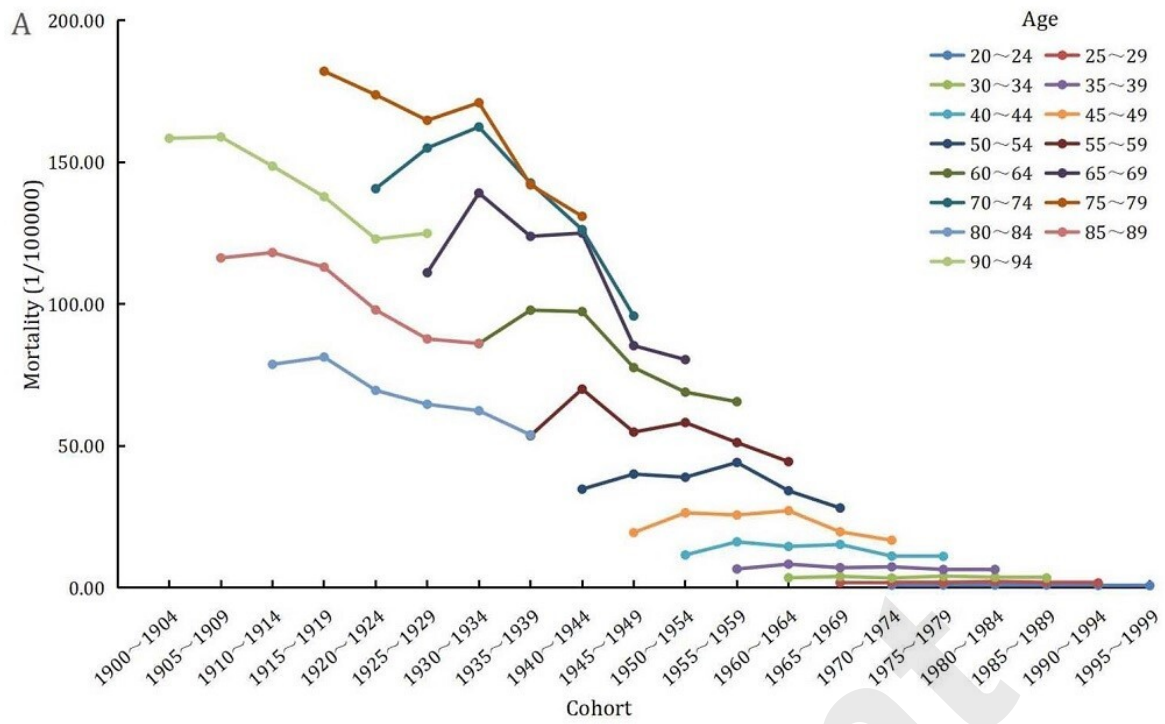


Figure 3. Age-cohort changing trends in stroke mortality attributed to an elevated BMI in Europe (A: male; B: female)

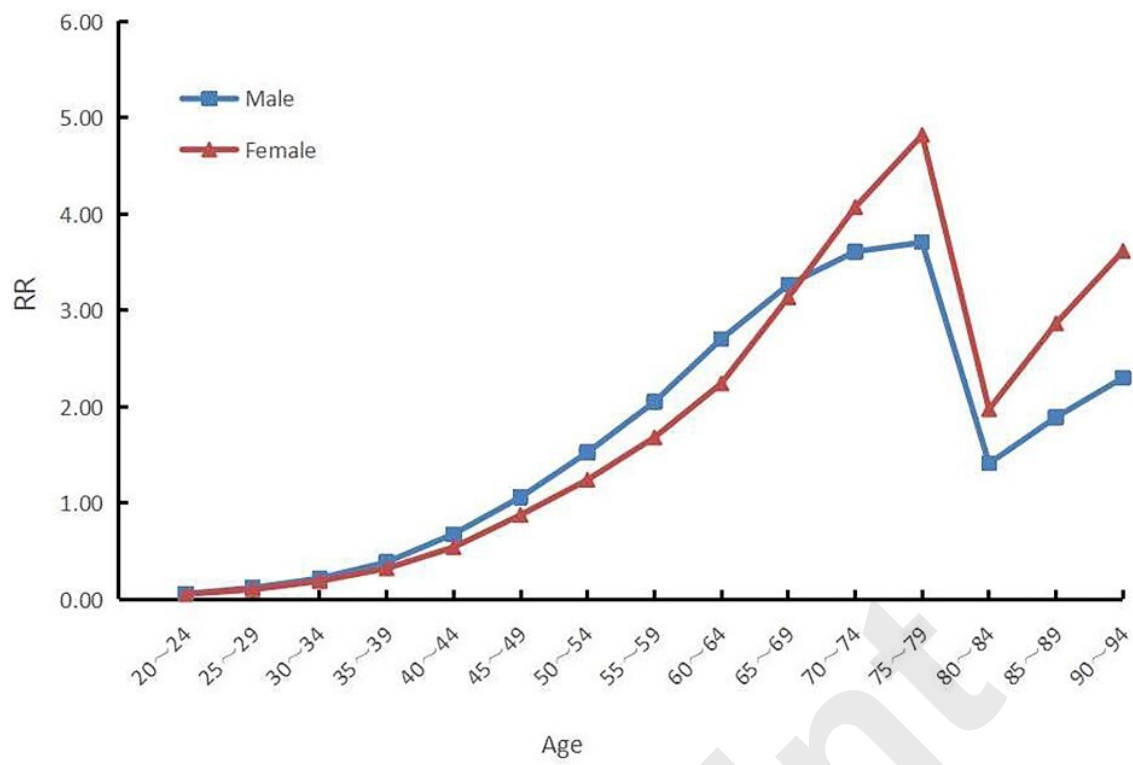


Figure 4. Age effect of stroke mortality attributed to an elevated BMI in Europe

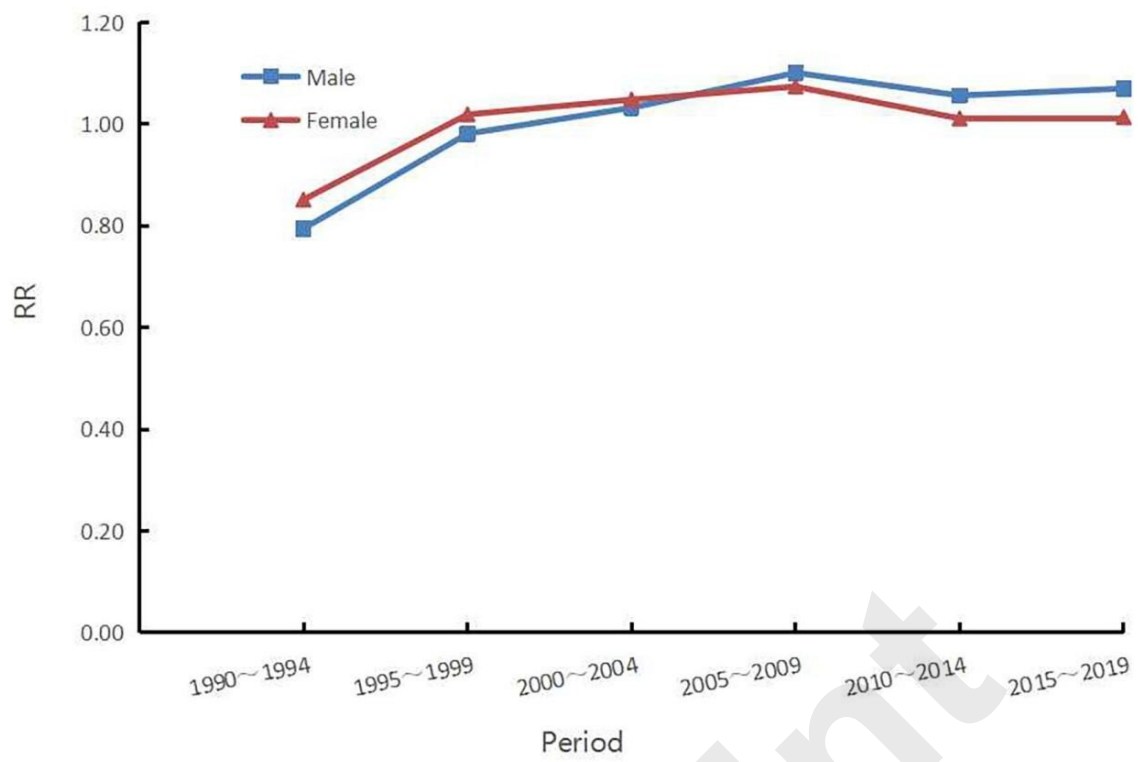


Figure 5. Period effect of stroke mortality attributed to an elevated BMI in Europe

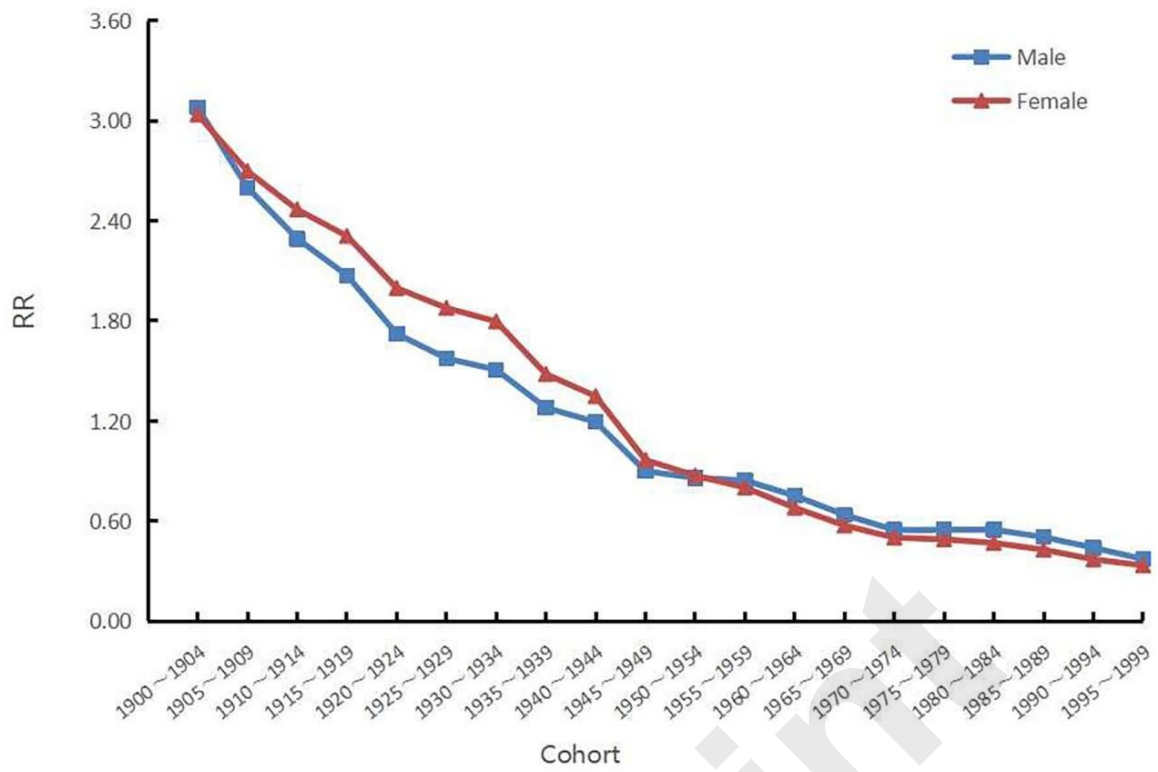


Figure 6. Cohort effect of stroke mortality attributed to an elevated BMI in Europe

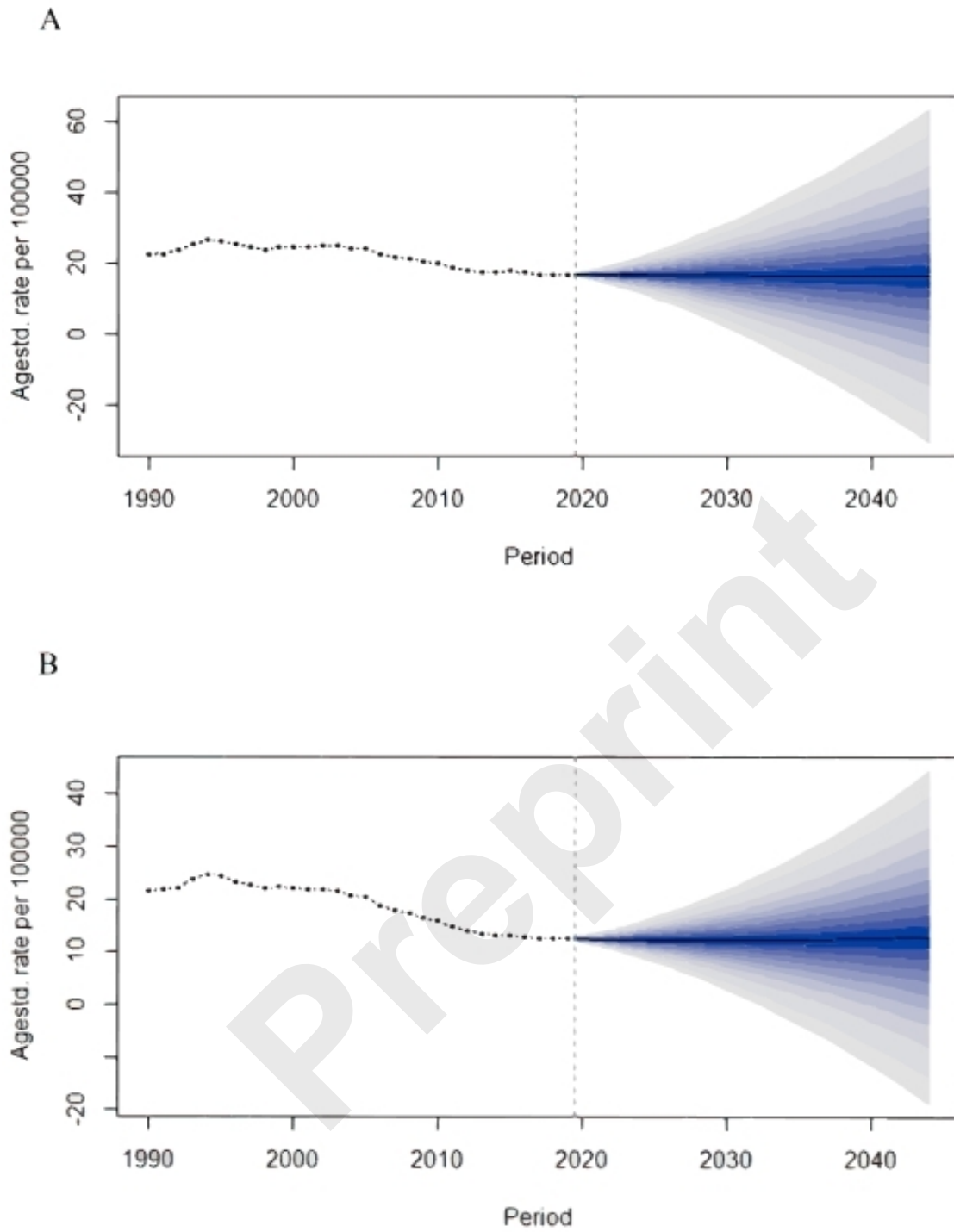


Figure 7. Prediction of mortality rate of stroke in Europe from 2020–2044(A: male; B: female)