

Trend of declining stroke mortality in China: reasons and analysis

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ABSTRACT

Introduction There is a downward trend of stroke-related mortality in the USA. By reviewing all published articles on stroke mortality in China, we analysed its trend and possible factors that have influenced the trend.

Methods Both English and Chinese literatures were searched on the mortality of stroke or cerebrovascular diseases in China. Potential papers related to this topic were identified from PubMed, Medline, Embase, Cochrane Library, Wanfang Database, SINOMED and China National Knowledge Infrastructure databases.

Results Comparing the results from the most recent population-based epidemiological survey and databank from the national Center for Disease Control and Prevention, the age-adjusted stroke mortality rate has shown a downward trend among both urban and rural population in the past 30 years in China. Comparing with 30 years ago, the rate of stroke mortality has decreased by more than 31% in urban/suburban population and 11% in rural population. In men, the age-adjusted stroke mortality rate decreased by 18.9% and in women by 24.9% between 1994 and 2013. Factors that may have contributed to the trend of decreased stroke mortality rate include (1) improved healthcare coverage and healthcare environment: (2) improved treatment options and medical technology; (3) support by government to educate the public on stroke and stroke prevention; and (4) improved public knowledge on stroke.

Conclusions The age-adjusted stroke mortality rate in China has shown a downward trend among both urban and rural population in the past 30 years. The major influencing factors that helped in reducing stroke mortality in China included improved healthcare coverage, healthcare environment, the updated treatment options and modern medical technology.

INTRODUCTION

Stroke is the second leading cause of death above the age of 60 years and the fifth leading cause of death in people aged 15–59 years old in the world.¹ In China, about 2.4 million people have a new stroke each year,² and 75% live with varying degrees of disability from stroke.³ However, in several Western countries such as the USA, the age-adjusted mortality rate of stroke has decreased by 37% from 1999 to 2009.⁴ In China, even with increased stroke incidence, stroke mortality rate has also been decreased in the past decades. In 2005, stroke was found to be the first leading cause of death in the third national survey on all causes of death.⁵ Then, what was the trend in stroke mortality in China over the past 30 years? Was it trending up or down? By reviewing all published articles on stroke mortality in China, we analysed its trend and possible underlying factors that have influenced the trend.

METHODS

Both English and Chinese literatures were searched on the mortality of stroke or cerebrovascular diseases in China. Potential papers related to this topic were identified from PubMed, Medline, Embase, Cochrane Library, Wanfang Database, SINOMED and China National Knowledge Infrastructure databases. The reference lists of all relevant studies and reviews were also checked in order to identify any pertinent information. Our search strategy consisted of MeSH terms and the following keywords: stroke mortality, cerebral vascular disease mortality, death from stroke or death from cerebral vascular disease in Chinese or China. Published data were tabulated based on their different variables. Reported rates on the mortality of stroke were tabulated and analysed.

RESULTS

Data from the national population-based epidemiological study

The results from the national epidemiological survey of stroke in China in 2013 were published in *Circulation* at the beginning of 2017.² The survey was implemented and conducted by the Beijing Neurosurgical Institute and the National Office for Cerebrovascular Disease Prevention and Control, National Health and Family Planning Commission of China, and the Chinese Center for Disease Control and Prevention (CDC), and supported by the Ministry of Science and Technology. It was conducted at 157 national disease monitoring sites in 31





provinces (autonomous regions and directly governed municipalities) using multistage stratified clusters sampling method. The sampling was calculated according to a 1% prevalence rate of stroke, which required a total of 600000 populations (across all ages) that should be surveyed. This study project was completed by two teams in two steps. The first step was the screening, completed by the investigators at the CDC of provincial and district/ county level. The second step was an onsite visit of those who were diagnosed with either a stroke, possible stroke or died from stroke from 1 September 2012 to 31 August 2013 by the local neurologists. This method helped to ensure the accuracy and reliability of the study results.

Death due to stroke was determined according to the following criteria: patient's history, imaging test results and hospital records when alive, or the death certificate. In addition, entry would be confirmed if one of the following criteria was met: (1) bedridden after stroke and died from infection or organ failure; (2) dysphagia after stroke and died from aspiration pneumonia; (3) death from recurrent stroke; and (4) death from complications such as pneumonia, gastrointestinal haemorrhage, cardiopulmonary failure and so on. Those who have had partial recovery from stroke but died from other known causes, such as suicide, motor vehicle accident or cancer, were excluded from this survey. For those cases that had no clear medical history, or a diagnosis of death from old age or unclear causes, two neurologists would adjudicate their causes of death.

To determine the death attributable to stroke, the investigator obtained information by an onsite visit and combined it with the data on the causes of stroke. Onsite review was required by neurologists so that all documented deaths from 1 January 2012 to the day of the survey could be accurately recorded. The statistical analysis was performed on the mortality data obtained between 1 September 2012 and 31 August 2013. The mortality rate was calculated as the total number of death/average number of population during that period. Table 1 shows the total (cross all ages) mortality of stroke. Of all population investigated (595711 people), the overall crude mortality rate was 127.2/100000 (95% CI 118.2 to 136.3), and the weighted mortality rate was 85.9/100000 (95% CI 71.8 to 100.1). Among them, the crude mortality rate for men was 139.1/100000 (95% CI 125.8 to 152.5), and the weighted mortality rate was $90.0/100\ 000\ (95\%)$ CI 72.8 to 107.3). In women, the crude mortality rate of stroke was 115.2/100000 (95% CI 103.0 to 127.4), and the weighted mortality rate was 81.6/100000 (95% CI 64.5 to 98.8). The age-adjusted stroke mortality rate (age-adjusted by the WHO world population) in rural areas was 116.8/100 000, significantly higher than that in the urban areas, which was 74.9/100000.

Table 2 lists the weighted mortality rate of stroke in adults (\geq 20 years) in 31 provinces in China. The top three provinces with the highest stroke mortality rate were Liaoning province (282.0/100 000), city of Tianjin (260.5/100 000) and Hubei province (211.9/100 000).

The two regions with the lowest stroke mortality rate were Shanghai $(7.3/100\,000)$ and Hainan province $(8.5/100\,0000)$.

Early data from the population-based survey

In 1980s, several large-scale, cross-sectional, population-based surveys on stroke were conducted.^{6 7} These surveys were also conducted by onsite visit by neurologists. The methodology and the way to define stroke mortality were similar to the study employed in 2013. However, in 1980s, CT scan was just introduced to China and the diagnosis of stroke was mostly based on clinical findings. Therefore, the accuracy in stroke classification was lacking. In the 2013 survey, over 90% of patients with stroke had imaging examinations, which helped with the diagnostic accuracy, better stroke classification and determination of the cause of death from stroke.

In 1983, the survey of six Chinese cities $(63\,195\,\text{people})$ showed that the age-adjusted mortality rate of stroke (age-adjusted by WHO world population)⁸ was about 108.5/100000. In 1985, the survey of rural areas in 22 provinces (246812 people) showed that the age-adjusted mortality rate of stroke was 131.8/100000. In 2013, the survey in urban residents showed that the age-adjusted mortality rate of stroke was 74.9/100000. To compare the two different rates of stroke mortality from these two different periods, it appeared that stroke mortality rate in China was lowered by 31.0% (six cities 30 years ago).⁵ The 2013 survey also showed that the age-adjusted stroke mortality rate in all ages of residents in rural areas was 116.8/100000, a 11.4% decrease compared with 30 years ago (figure 1).

In 1985, the other large-scale, population-based study on stroke mortality was conducted by the Armed Forces Epidemiology Research Group. It was a cross-sectional onsite survey of 5 814 851 people, which showed a stroke mortality rate of $77.2/100\,000^{9\,10}$ Since the sample size was too large, quality control of the survey was difficult. This might have contributed to the findings of lower rate of stroke prevalence, incidence and mortality than those in the other surveys during the same period in 1980s.

Since 1970s, the Ministry of Health of China has organised three national retrospective surveys of the causes of death. The second survey was completed during 1990-1992 and the third one between 2004 and 2005.⁴ The survey recorded the findings based on findings from a combined search by clue and onsite visit findings. The second survey (1990-1992) found the stroke death rate (crude) was about 101.93/100000, age-adjusted (by China census population 1982) was 84.54/100000. At that time, cerebrovascular disease was the third leading cause of death in China. However, the third national survey (2004–2005) revealed that the stroke death rate (crude) was 136.64/100000, age-adjusted to 81.19/100000. It was this survey that has found cerebrovascular disease or stroke had become the leading cause of death in China (table 3).

Table 1 Mor	tality rates (wi	th 95% CI)	Table 1 Mortality rates (with 95% Cl) of stroke per 100000 person-years among Chinese in 2013	rson-years a	mong Chines	se in 2013			
	Men			Women			Total		
Age Popula group (years) at risk	Population at risk	Stroke (n	Stroke (n) Rate* (95% CI)	Population at risk	Stroke (n)	Stroke (n) Rate* (95% Cl)	Population at risk		Stroke (n) Rate* (95% Cl)
<20	63442	0	0.0 (0.0 to 0.0)	56113	0	0.0 (0.0 to 0.0)	119555	0	0.0 (0.0 to 0.0)
20–29	45961	0	0.0 (0.0 to 0.0)	47 553	0	0.0 (0.0 to 0.0)	93514	0	0.0 (0.0 to 0.0)
30–39	45508	ę	6.6 (0.0 to 14.1)	44 462	-	2.2 (0.0 to 6.7)	89 970	4	4.4 (0.1 to 8.8)
40-49	56068	20	35.7 (20.0 to 51.3)	55 022	14	25.4 (12.1 to 38.8)	111 090	34	30.6 (20.3 to 40.9)
50-59	39171	38	97.0 (66.2 to 127.8)	40 017	28	70.0 (44.1 to 95.9)	79 188	66	83.3 (63.2 to 103.4)
60-69	29 056	102	351 (283.0 to 419.1)	30352	51	168.0 (122.0 to 214.1)	59 408	153	257.5 (216.8 to 298.3)
62-02	15737	147	934.1 (783.8 to 1084.4)	16561	104	628.0 (507.7 to 748.3)	32 298	251	777.1 (681.4 to 872.9)
80+	4782	107	2237.6(1818.4 to 2656.8)	5906	143	2421.3(2029.3 to 2813.3)	10688	250	2339.1 (2052.5 to 2625.6)
Total	299725	417	139.1 (125.8 to 152.5)	295 986	341	115.2 (103.0 to 127.4)	595 711	758	127.2 (118.2 to 136.3)
Weighted*	I	I	90.0 (72.8 to 107.3)	I	I	81.6 (64.5 to 98.8)			85.9 (71.8 to 100.1)
*Weighted rates to China census population 2010	to China censu	is populatior	n 2010.						

Table 2 Mortality rates (with 95% CI) of stroke per 100 000 person-years among adults (≥20 years) in 31 provinces of China							
Provinces	Population at risk	Death (n)	Mortality rates	Rate* (95% CI)			
Shanghai	7354	1	13.6	7.3 (0 to 21.7)			
Hainan	5829	2	34.3	8.5 (0 to 26.8)			
Guangdong	16297	13	79.8	48.7 (22.7 to 74.7)			
Shanxi	18563	19	102.4	56.9 (12.2 to 101.6)			
Sichuan	20945	22	105.0	90.8 (47.1 to 134.5)			
Jiangsu	20054	23	114.7	97.7 (8.2 to 187.3)			
Beijing	6652	8	120.3	65.7 (16.1 to 115.4)			
Xinjiang	9612	12	124.8	150.0 (39.1 to 260.9)			
Guangxi	19881	25	125.7	106.9 (31.6 to 182.2)			
Jiangxi	14926	20	134.0	87.9 (49.9 to 125.9)			
Shaanxi	15930	22	138.1	143.6 (73.9 to 213.3)			
Chongqing	6433	9	139.9	86.9 (48.4 to 125.4)			
Yunnan	17040	25	146.7	89.9 (13.1 to 166.7)			
Fujian	14912	22	147.5	104.3 (40.7 to 167.9)			
Guizhou	15429	23	149.1	171.3 (40.5 to 302.1)			
Hebei	23232	35	150.7	128.9 (26.2 to 231.6)			
Anhui	20260	32	157.9	120.3 (88.4 to 152.3)			
Qinghai	8723	14	160.5	124.4 (100.9 to 147.8)			
Jilin	16526	27	163.4	50.1 (0 to 128.7)			
Henan	24448	41	167.7	178.7 (117.3 to 240.1)			
Shandong	24929	43	172.5	99.1 (45.6 to 152.6)			
Hunan	20803	36	173.1	123.1 (103.7 to 142.5)			
Zhejiang	19240	34	176.7	136.9 (91.9 to 181.8)			
Heilongjiang	22122	45	203.4	90.6 (44.5 to 136.7)			
Inner Mongolia	15612	32	205.0	145.3 (91 to 199.6)			
Ningxia	5962	13	218.0	147.8 (80.3 to 215.3)			
Tibet	5666	13	229.4	157.4 (72.4 to 242.4)			
Gansu	13393	31	231.5	135.0 (12 to 257.9)			
Hubei	19260	46	238.8	211.9 (129 to 294.9)			
Tianjin	6905	17	246.2	260.5 (125.7 to 395.2)			
Liaoning	19218	53	275.8	282.0 (99.6 to 464.5)			

*Weighted rates to China census population 2010.

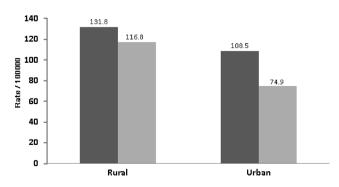


Figure 1 Age-standardised mortality of stroke in rural and urban areas of China compared between the survey in 2013 and 1983–1985.

Data from the national disease surveillance system on stroke mortality

Filing a report on death began in 1980s in China. There were two systems: the national disease surveillance points (DSP) and registry on cause of death by the Ministry of Health at that time.

The DSP system began in early 1980s. Professor He Guan-qing, a well-known epidemiologist, established a few pilot surveillance sites in the Dongcheng District of Beijing and Tongxian county. Surveillance sites were increased to 71 in 29 provinces (autonomous zone and municipality) in 1989. The project later became the official national DSP. In 1990, with the support from the World Bank, the Chinese Academy of Preventive Medicine employed multistage stratification random sampling

Table 3 Death rates of six main diseases in the second (1990–1992) and the third survey of the causes of death (2004–2005) in China

	1990–1992				2004–2005				
Rank	Disease	Death rate	Age- standardised* rate	Percentage in all deaths	Disease	Death rate	Age- standardised* rate	Percentage in all deaths	
1	Diseases of the respiratory system	137.52	115.47	22.79	Cerebrovascular disease	136.64	81.19	22.45	
2	Malignant neoplasms	108.26	94.22	17.94	Malignant neoplasms	135.88	91.24	22.32	
3	Cerebrovascular disease	101.93	84.54	16.89	Diseases of the respiratory system	96.28	56.27	15.81	
4	Injury and poisoning	66.16	61.44	10.97	Heart disease	90.23	52.2	14.82	
5	Heart disease	52.70	43.84	8.73	Injury and poisoning	61.51	49.48	10.10	
6	Diseases of the digestive system	30.37	26.5	5.03	Diseases of the digestive system	16.78	11.14	2.76	

*Age-standardised by China census population 1982.

process and built representative sampling sites in 31 provinces, municipalities and autonomous zones, and formed systematically 145 DSPs. This new DSP system covers 10 million populations and routinely collects data on any birth and mortality (about 1% of Chinese population). In 2003, the DSP system was expanded to 160 sites and covered 70 million populations.¹¹

The Ministry of Health became responsible for all mortality registries in 1987. After 2000, this registry covered 90 counties in 15 provinces and eastern rural areas of China, about 100 million people, 8% of the Chinese population. Among them, 50% lived in the eastern area, 40% in central China and only 10% in the western region. The statistical centre at the Ministry of Health would collect and analyse the mortality rate of all kinds of diseases annually.

Recently, based on the data on global disease burden in 2013 (data from Chinese DSP system of CDC), Wang et al¹² from the School of Public Health of Wuhan University published in Stroke about the downward trend of stroke mortality among different age-adjusted groups in China from 1994 to 2013. Overall, in 20 years, stroke mortality decreased by 18.9% in men, but increased a little after 2010. In women, stroke mortality showed a consistent downward trend, decreased by 24.9% (figure 2). Professor GH Yang of the Chinese CDC stated that the age-adjusted stroke mortality rate was about 167.0/100000 in 1990 and decreased to 126.8/100000 by 2010. In this report, the mortality from ischaemic stroke was markedly less than that from intracerebral haemorrhage. Among them, the age-adjusted mortality rate of ischaemic stroke was 56.3/100000 in 1990 and reduced to 46.7/100000 in 2010. Mortality from intracerebral haemorrhage was 110.7/100000 in 1990 and dropped to 80.2/100000 in 2010.¹³ However, the other Chinese CDC document reported that the age-adjusted stroke

mortality rate was 199.7/100000 in 1990 and reduced to 157.3/100000 in 2013. Among them, death from ischaemic stroke was 51.6/100000 in 1990 but increased to 62.9/100000 in 2013. Death related to intracerebral haemorrhage was 148.1/100000 in 1990 and dropped to 94.4/100000 in 2013.¹⁴ Although both documents came from Chinese CDC, there was a significant discrepancy between the two. The Statistical and Information Center of the Ministry of Health reported stroke mortality from the survey results of 36 cities and 80 counties in China from 1990 to 1998. During this period, the age-adjusted (by China census population 1982) stroke mortality among the urban residents was 84.11/100000 and 74.05/100000 in rural areas.¹⁵ During a 9-year period, stroke mortality in urban areas was showing a downward

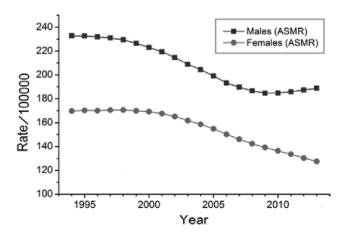


Figure 2 Trends of ASMR of stroke by sex in China, 1994–2013. ASMR, age-standardised mortality rates; CMR, crude mortality rate. (This figure is partly quoted from published paper Wang *et al.*¹²)



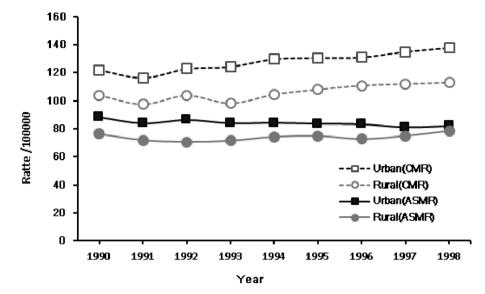


Figure 3 Trends of the crude mortality rates and the age-standardised mortality rates (ASMR) per 100 000 population for stroke in urban and rural in China, 1990–1998.

trend, but in rural areas it was showing a slightly upward trend (figure 3).

Data from the Monitoring Trend and Determinants in Cardiovascular Disease (MONICA) study

Designed by the expert group of WHO, the MONICA study was a project to monitor the trend of cardiovascular disease and stroke incidence and mortality in different countries. The MONICA study in China monitored people between the ages of 35 and 64 in 17 provinces from 1987 to 1993 and found that the mortality rate varied significantly. Anhui province had the lowest male stroke mortality rate at 27.0/100 000. The highest rate was in the province of Heilongjiang, at 107.3/100000. Fujian province had the lowest stroke mortality rate in women at 14.0/100000. Overall, stroke mortality rate did not change significantly between 1987 and 1993.¹⁶

DISCUSSION

Literature has reported that in countries of high income, age-adjusted stroke mortality rate has decreased by 37% in the past 20 years.¹⁷ In America, age-adjusted stroke mortality rate has always shown a declining trend in the past 50 years. Stroke has dropped down to be the fourth common death of diseases.¹⁸ The 2013 scientific statement by the AHA/ASA mentioned that the factors that have caused the lower stroke mortality rate included lower incidence of stroke and fewer people dying from stroke. Looking back to 1970s, the strategy of controlling blood pressure better probably had the biggest impact on stroke mortality. Tightly controlling other risk factors such as diabetes and hyperlipidaemia, lowering rate of smoking, together with blood pressure control have all helped in lowering stroke mortality in the USA.

In China, the research on this newly discovered lowering trend in stroke mortality is lacking. It is even more difficult to explain why stroke has become the leading cause of death in China since 2005 while the report on stroke mortality has shown a downward trend over the past 30 years. Nevertheless, we have postulated the following possible explanations:

The improvement of overall healthcare conditions in China

Improvement in healthcare environment and technology are two important factors that lowered stroke mortality in China. In the past 30 years, Chinese economy has had rapid growth. Healthcare access has continued to improve among urban and rural residents. The new healthcare insurance plan for the rural area helped rural residents get treated much more readily. Parry¹⁹ reported that the health insurance coverage in China has increased from 45% in 2006 to around 90% in 2009. With the improved healthcare insurance coverage, more patients with stroke could get timely treatment in the hospital, which would reduce poststroke complications and death.

Hypertension is considered the silent killer since it is the most important risk factor for all vascular diseases including stroke. A report by Liao *et al*²⁰ showed that in China, treatment rates among those diagnosed with hypertension have increased to around 90% by 2009. And such improved awareness was more prominent in elderly.

Advancement of the clinical diagnosis and treatment technology

Significant improvements on techniques for diagnosis, treatment and management, accompanied by more and better healthcare professionals, played a critical role in the decline of case fatality of stroke, and thus lowered the overall stroke mortality.^{12 21} Data from the Chinese National Stroke Registry showed that during the beginning stage of offering intravenous rt-PA treatment to patients with acute ischaemic stroke, about one in five patients had the treatment if they presented within 3 hours.²² Early treatment has translated into improved

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neurological outcome and hence lowered mortality of stroke. In recent years, the number of patients with acute ischaemic stroke treated with intravenous rt-PA has increased, especially in many cities in China. In addition, the establishment of stroke unit has improved stroke care and might have contributed to lowered stroke mortality. A cross-sectional survey in 2004 revealed that 7.4% of the 124 hospitals have established stroke units in Beijing. The report also stated that at the time of survey, around 300 stroke units have been established across the country.^{23 24}

Many county hospitals in rural areas have started to offer intravenous thrombolysis to treat acute ischaemic stroke. In the past 20 years, a new treatment technique, minimally invasive transcranial needle aspiration of haematoma for patients with intracerebral haemorrhage, has been widely implemented in China.²⁵ By the end of 2013, nearly 10000 neurosurgeons, neurologists and doctors working at local emergency rooms (ERs) were trained on this technique, which has benefited more than one million patients with intracerebral haemorrhage.²⁶

The decline of stroke incidence in urban residents and epidemiological transition of stroke subtype in China

The current (2013) study has found that stroke incidence was higher than the findings in previous surveys in China, even substantially increased in rural areas over the last three decades. However, in urban residents, the age-ad-justed incidence of stroke (168.2/100000) declined significantly than that in the 1983 survey (205.4/100000), a decrease by 18.1%. This must have had a significant impact on the decline of stroke mortality in China.

A prospective population-based registry showed that the incidence of ICH has decreased significantly at a rate of 12.0% per year in Beijing, 4.4% in Shanghai and 7.7% in Changsha during the 1990s, although the incidence of ischaemic stroke has increased in Beijing (5.0% per year) and Shanghai (7.7%).²⁷ In recent times, the prevalence of obesity, hypertension, diabetes and hypercholesterolaemia in Chinese populations has increased because of increased calorie intake, alcohol consumption, cigarette smoking and reduced physical activity. There is a trend of stroke epidemiological transition, with increased IS and decreased ICH in China. This trend might have also played a role in the decline of stroke mortality over the past 30 years in both urban and rural areas.

The impact on improvements of stroke knowledge

Improved stroke education and awareness of stroke in the past 30 years have helped in reducing stroke mortality. The Chinese government has invested heavily in the prevention and treatment of non-communicable diseases, which has rapidly improved the healthcare conditions and upgraded technology for all levels of healthcare institutions. Modern stroke prevention and treatment guidelines have facilitated the standardisation of stroke care for physicians and improved their skills.^{28 29} Furthermore, public knowledge on stroke prevention and seeking timely treatment when stroke strikes have improved significantly in China. Currently, there is ongoing national campaign to shorten the prehospital delay of recognising stroke, transporting patient to the nearest ER, and treatment with intravenous thrombolysis and intra-arterial thrombectomy. It is likely that with improved public knowledge on stroke, fast transport of stroke patient and expedited treatment, stroke mortality rate may decrease further in China.

CONCLUSION

From the results obtained in the most current population-based epidemiological survey and data from the national CDC, the age-adjusted stroke mortality rate has shown a downward trend among both urban and rural population in the past 30 years. Comparing with 30 years ago, the rate of stroke mortality has decreased by more than 31% in urban/suburban population and 11% in rural population. In men, the age-adjusted stroke mortality rate decreased by 18.9% and in women by 24.9% between 1994 and 2013.

Factors that may have contributed to the trend of decreased stroke mortality rate in the past 20–30 years in China include (1) improved healthcare coverage and healthcare environment; (2) improved treatment options and medical technology; (3) support of government to educate the public on stroke prevention; and (4) better public knowledge on stroke. More research is needed to monitor the trend of stroke mortality in China.

Correction notice This paper has been amended since it was published Online First. Owing to a scripting error, some of the publisher names in the references were replaced with 'BMJ Publishing Group'. This only affected the full text version, not the PDF. We have since corrected these errors and the correct publishers have been inserted into the references.

Competing interests None declared.

Provenance and peer review Commissioned; internally peer reviewed.

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