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## Trends in stroke incidence in high-income countries in the 21<sup>st</sup> century: population-based study and systematic review

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### Abstract

**Background and Purpose**—Population-based studies provide most reliable data on stroke incidence. A previous systematic review of population-based studies suggested that stroke incidence in high-income countries decreased by 42% between the 1970s and early 2000s. However, it is uncertain whether this trend of steady decline has been maintained in more recent periods.

**Methods**—Data from the Oxfordshire Community Stroke Project (OCSP; 1981-1986) and the Oxford Vascular Study (OXVASC; 2002-2017) along with other published population-based stroke incidence studies that reported temporal trends of stroke incidence since 1990 in high-income countries were included. Age-standardised relative incidence rate ratios (IRR) were calculated for each study and then pooled with inverse variance weighted random effects meta-analysis. Projection estimates were calculated for the number of incident stroke patients in the UK from year 2015 to 2045.

**Results**—In Oxfordshire, stroke incidence fell by 32% from OCSP to OXVASC, with a similar trend before or after year 2000. With the projected aging population, if the age-specific stroke incidence continued to decrease at its current rate (6% every 5 years), there would still be a 13% increase of the number of first-ever strokes in the UK up to year 2045. Incorporating the Oxfordshire data with other 12 population-based studies, stroke incidence declined steadily between the 1990s and 2010s within each study, resulting in a 28% decline over an average period of 16.5 years (pooled IRR=0.72, 95% CI 0.66-0.79,  $p<0.0001$ ). The trend was the same for men (0.69; 0.61-0.77,  $p<0.0001$ ) and for women (0.66; 0.59-0.74,  $p<0.0001$ ), and remained consistent after year 2010 in OXVASC. Proportion of disabling or fatal stroke also decreased over time (early vs. later period 53.6% vs. 46.1%,  $p=0.02$ ).

**Conclusions**—Stroke incidence is continuing to decline with steady rate in Oxfordshire and in other high-income settings. However, the absolute number of strokes occurring is not falling.

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#### Disclosures

Drs. Li, Scott and Rothwell reports no disclosures.

## Keywords

stroke; incidence; population studies; trends; systematic review

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## Introduction

Stroke is the second leading cause of death worldwide, and the main cause of long-term neurological disability in adults.<sup>1</sup> In high-income countries, increased use of preventive treatments and major reductions in premorbid risk factors between the 1970s and early 2000s have coincided with significant falls in stroke incidence during this period.<sup>2</sup> However, it is uncertain whether this trend of steady decline has been maintained in more recent years, particularly in view of the continuing ageing of the population and the rise of diabetes and obesity in recent decades.<sup>3–5</sup> Reliable estimates and projections of the future stroke burden are important for long-term healthcare infrastructure planning, policy making, and are important for the organisation of stroke services and prevention activities.

Some studies based on administrative data reported that the reduction of stroke incidence was less marked in the early 2000s.<sup>6, 7</sup> However, accuracy of coding of stroke in routinely collected data can also vary over time.<sup>8, 9</sup> Prospective population-based incidence studies provide the most reliable data on stroke incidence.<sup>2</sup> Therefore in a population-based study in Oxfordshire, UK, we determined the change of stroke incidence in the last four decades and estimated the potential burden of incidence stroke in the UK from year 2015 to 2045. A few other population-based studies also maintained “ideal” case-ascertainment procedures over a long period of time to provide comparable estimates of stroke incidence change over time. Since the publication of the most recent systematic review of temporal trend of stroke incidence in high-income countries,<sup>10</sup> several population-based studies have reported new data on stroke incidence, particularly after year 2010.<sup>11–18</sup> We therefore also combined our results in an updated systematic review to determine the temporal trends of stroke incidence between 1990s and 2010s in high-income settings.

## Methods

*Requests for access to data from the Oxford Vascular Study will be considered by the corresponding author.*

## Study Population

The Oxford Vascular Study (OXVASC) is an ongoing prospective population-based study of the incidence and outcome of all acute vascular events in a mixed rural/urban population. The study population comprises all 92,728 individuals, irrespective of age, registered with approximately 100 general practitioners (GPs) in nine general practices in Oxfordshire, UK. Ascertainment of acute vascular events started on 1 April 2002 and is ongoing. This analysis reports all first-ever incident strokes during the first 15-years (2002-2017).

The detailed study methodology and case definitions have been published before (supplementary methods), and multiple overlapping methods of “hot” and “cold” pursuit were used to achieve near complete and consistent ascertainment of all individuals with

stroke.<sup>19, 20</sup> To ensure consistency, one senior stroke neurologist (PMR) has prospectively reviewed all cases over the 15 years. Stroke was defined as an event with appropriate symptoms lasting longer than 24 hours.

Stroke incidence in OXVASC was also compared with that in the Oxfordshire Community Stroke Project (OCSP), which was a high-quality, population-based study of all first-ever stroke in 1981 to 1986 in the same general practice population. The methodology of OCSP has also been published before (supplementary methods),<sup>21</sup> and the case diagnosis, assessment, and follow-up were similar to those in OXVASC. In particular, to ensure consistency of clinical diagnosis between OCSP and OXVASC, summaries of all potential cases in the first 2 years of OXVASC were reviewed by the principal investigator of OCSP to ensure that the application of definitions of strokes was comparable between the two studies.<sup>19</sup>

In both OCSP and OXVASC, patients were ascertained and assessed face-to-face by study physicians as soon as possible after the initial presentation in hospital, an emergency clinic or at home. If a patient died before assessment, we obtained an eyewitness account of the clinical event and reviewed any relevant clinical records. In OCSP, CT brain or autopsy was the first-line imaging modality and 12% of the cases did not have brain imaging or autopsy. In OXVASC, patients routinely had brain imaging (CT or MR), vascular imaging (Carotid Doppler or CT-angiography/MR-angiography or digital subtraction angiography), 12-lead electrocardiography (ECG) and standard blood tests. Echocardiography, 24-hour ECG (HOLTER) and 5-day ambulatory ECG monitoring were done when clinically indicated. Less than 5% of the cases in OXVASC did not have any brain imaging or autopsy.

All patients were followed up face to face at 1, 6, 12, 60 and 120 months by a study nurse or physician to determine their functional status (modified Rankin Scale, mRS).<sup>22</sup> Disabling or fatal stroke is defined as new disability (mRS>2) or death or progression of disability (one score increase in mRS) in those with premorbid disability (pre-morbid mRS>2) using the 1 month mRS. For patients who had moved out of the study area, follow up was performed by telephone or email. All patients were flagged for the Office for National Statistics mortality data and all deaths during follow-up were recorded with causes.

### Systematic review

We followed the PRISMA guideline and identified all stroke incidence and prevalence studies referenced in previous published reviews and hand-searched PUBMED and EMBASE for any follow-up or secondary studies using the author and study names from the primary studies. We also performed a further search of PUBMED (1950-December 2017) and EMBASE (1974-December 2017) using the following search terms without restrictions: “exp STROKE/” OR “first ever stroke\*.ti,ab.” AND “incidence/” OR “registries/”. The abstracts of all papers identified from initial searches were reviewed by two authors (LL and CS), and both authors reviewed the full text of all eligible studies. In cases of disagreement about the eligibility of studies or data extraction, consensus was reached through joint reassessment.

Studies were included in the main analysis if they fulfilled the following criteria: 1) published incidence studies satisfying the quality criteria of “ideal” population-based study;<sup>23, 24</sup> 2) conducted in high-income countries;<sup>10</sup> 3) reported stroke incidence, incidence rate ratio or raw numbers sufficient to calculate incidence, or change in incidence at two or more time periods; 4) reported stroke incidence between 1990 ( $\pm 5$  years) and 2017, with the latest time period after 2010; 6) English language. Studies that fulfilled criteria 1) to 3) and reported the latest time point between year 2000 and 2010 were also included in subgroup analyses. Population-based studies confined to only one pathological stroke subtype were excluded. Where there was more than one publication on a cohort of patients, data on incidence rate were taken from the most recent publication or the publication with the most complete raw numbers. Updated data from OXVASC were also included.

Information was extracted from each report on the population studied, study period (duration), observed person-years, and total number of incident stroke cases. Where possible, we also documented the reported numbers of incident stroke cases by sex, stroke subtype (ischaemic stroke, intracerebral haemorrhage and subarachnoid haemorrhage) and stroke severity (disabling/fatal vs. non-disabling). Where available crude and standardised incidence rates (IR) with confidence intervals, as well as raw numbers (numerator, n and denominator N) were extracted for each time period reported. Where IR, n or N were not reported, when possible these were calculated based on reported incidence rate ratio (IRR).

### Statistical analyses

For Oxfordshire and other included studies, age-specific crude incidence rates (/100,000 population/year) were calculated with confidence intervals estimated assuming a Poisson distribution. For studies which reported stroke incidence in different age groups, sexes, ethnicities or stroke subgroups, where raw numbers were not available, we used inverse variance weighted fixed effects meta-analysis to generate summary incidence rates.

All crude incidence rates were standardised to the European population using the direct method.<sup>25</sup> Standardised relative incidence rate ratios (IRR) were then calculated for the chosen two periods within each study using Poisson regression models adjusting for the age structure of the two populations. IRR from individual studies were then pooled with inverse variance weighted random effects meta-analysis to generate a pooled IRR with 95% confidence intervals. We estimated the percentage of variability across studies attributable to heterogeneity beyond chance using the  $I^2$  statistic.

In the main analysis, pooled estimates for temporal trends of stroke incidence between 1990s and 2010s were calculated. We also performed subgroup analyses stratified by stroke subtypes (ischaemic stroke, intracerebral haemorrhage and subarachnoid haemorrhage), sex, stroke severity (disabling/fatal vs. non-disabling) and study periods (1990-2000 vs. after year 2000). Sensitivity analyses were also carried out in relation to study duration.

Finally, in order to estimate the number of incident stroke patients on a national basis in the UK from 2015 up to 2045, we used the age-specific incidence during 2014-2017 in OXVASC for standardisation to the projected UK population.<sup>26</sup> Two different scenarios for changes in age-specific incidence were used: stable rates over time or maintaining the

current 6% decrease every 5 years. Moreover, we calculated how much the incidence would have to decrease every 5 years to maintain a stable number of incident stroke patients in the UK in 2045 compared with 2015.

All analyses were performed using SPSS version 22.

### Standard Protocol Approvals, Registrations, and Patient Consents

Written informed consent or assent from relatives was obtained in all participants. OXVASC was approved by the local research ethics committee (OREC A: 05/Q1604/70).

### Results

Based on 2811 incident stroke cases in OCSF (n=557) and OXVASC (n=2254), stroke incidence fell by 32% from 1981-1986 to 2014-2017 (IRR=0.68, 95% CI 0.60-0.78,  $p<0.0001$ ), with no evidence of heterogeneity before and after the year 2000 ( $p_{het}=0.54$ ) (OXVASC 2002-2005 vs. OCSF 1981-1986: 0.80, 0.70-0.91,  $p=0.0006$ ), and after (2014-2017 vs. 2002-2005: 0.85, 0.74-0.98,  $p=0.03$ ).

Using the above estimates, we applied different scenarios for the projection of the number of first-ever stroke patients in the UK between 2015 and 2045. The UK population is projected to increase by 3.6 million (5.5%) over the next 10 years and reaching 72.9 million in mid-2041 with doubling number of individuals aged 85 years.<sup>26</sup> If the age-specific incidence remained stable over the next 30 years, the number of incident stroke would increase by 66% from year 2015 to 2045. If the age-specific stroke incidence continued to decline with its current magnitude (OXVASC estimates: 6% every 5 years), there would be a 13% increase of the number of first-ever strokes in the UK up to year 2045. To maintain a stable number of incident stroke patients in the UK in 2050, the incidence would have to decrease by 8% every 5 years.

16,638 citations were identified and 13 population-based studies from 9 high-income countries were included in the systematic review (web appendix I).<sup>11-16, 19, 27-29, 30-31, online-only</sup> Of the 13 studies, nine (including unpublished data from OXVASC)<sup>11-19, 29-31</sup> reported temporal trends of stroke incidence including at least one period after year 2010 and were thus included in the main analysis (table 1). An additional four studies<sup>online-only</sup> reporting either temporal trends between 1990 and 2000 or after year 2000 were also included in the sensitivity analysis (web appendix II). No published study reported change of stroke incidence beyond year 2010.

Combining results from Oxfordshire and the systematic review, 31,351 new stroke cases occurred during approximately 18,453,235 person-years of observation. Despite noticeable variation in absolute age-standardised stroke incidence rates between studies, stroke incidence declined significantly between the 1990s and 2010s in all studies (web appendix I&II), resulting in less between-study difference in absolute rates after year 2010 (highest vs. lowest incidence before year 2000: 183 vs. 81 per 100,000 population; after year 2010: 143 vs. 98 per 100,000 population).

Although the rates of reduction in stroke incidence varied slightly between studies, the trend of steady decline was consistent across studies (figure 1). Consequently, the pooled estimate from the nine studies between 1990s and 2010s suggested a 28% decline of stroke incidence during an average study duration of 16.5 years (IRR=0.72, 95%CI 0.66-0.79,  $p<0.0001$ ; figure 2), with no differences between men and women (8 studies,  $p_{\text{difference}} = 0.56$ ; table 2). The results were also largely consistent by stroke subtypes (table 2). Sensitivity analyses excluding the four studies with the longest study duration (Oxfordshire, Sweden<sup>17, 18, 32, 33</sup> and Auckland<sup>11, 12</sup>) also showed consistent results (mean study duration 12 years; IRR=0.78, 0.70-0.86,  $p<0.0001$ ).

Two studies of similar duration (Oxfordshire and Porto<sup>13</sup>) reported the temporal trends of stroke incidence stratified by stroke severity (figure 3). In contrast to a 30% reduction of disabling or fatal stroke during a mean study duration of 11.5 years, there was no statistically significant reduction in non-disabling stroke (pooled IRR=0.98, 95%CI 0.85-1.12,  $p=0.73$ ; figure 3) due apparently to reductions in the proportion of disabling or fatal stroke (2 studies; early vs. later period 53.6% vs. 46.1%, OR=0.77, 0.62-0.96,  $p=0.02$ ; figure 3).

In the subgroup analysis stratified by study periods, although more heterogeneity was observed post year 2000 (web appendix III), the pooled estimates for the reduction of stroke incidence in each period were consistent (1990 to 2000: 9 studies, mean study duration= 9 years, IRR=0.84, 95%CI 0.74-0.94,  $p=0.003$  versus post year 2000:11 studies, mean study duration= 10 years, IRR=0.82, 0.73-0.93,  $p=0.001$ ; web appendix III), and was maintained after year 2010 in OXVASC (figure 1). There was also no difference in the trends between men and women within each period ( $p_{\text{difference}} = 0.22$  for 1990-2000 and  $p_{\text{difference}} = 0.73$  post year 2000; table 2). Again, results were also largely comparable for ischaemic vs. haemorrhagic strokes, although the reduction of subarachnoid haemorrhage was more marked post year 2000 (table 2) whereas the reduction of intracerebral haemorrhage became less marked in more recent periods (table 2).

## Discussion

In this updated systematic review of all population-based stroke incidence studies in high-income countries, we showed that the previously suggested 1.1% yearly percentage reduction in stroke incidence from the 1970s to early 2000s<sup>2</sup> is maintained and stroke incidence continued to decline at an annual rate of 1.0% to 1.5% in the last three decades. However, with the aging population, even if the age-specific stroke incidence continued to decrease at its current rate, there would still be a 13% increase of the number of first-ever strokes in the UK in year 2045.

Our findings based on population-based stroke incidence studies in high-income countries are supported by data from hospital-based registries, which also show a steady decline of hospitalised stroke in recent years in southeast Asia,<sup>34</sup> Western Europe<sup>35-37</sup> and North America<sup>38-40</sup>. The results are also consistent with the data modelling results from the Global Burden of Disease analysis.<sup>5</sup> Moreover, it is encouraging that the previously suggested 1.1% yearly percentage reduction in stroke incidence from the 1970s to early 2000s was

maintained,<sup>2</sup> possibly due to continued effort in implementing preventive treatment at the population-level to reduce smoking, hypertension and other vascular risk factors.<sup>19, 36, 41–42</sup>

However, even if stroke incidence continued to decline at its current rate, there would still be a 13% increase of the number of new strokes in the UK in year 2045 due to the aging population. Similar projections have also been reported for other European countries.<sup>27, 32, 33</sup> Moreover, reduced stroke case-fatality in high-income countries would lead to growing numbers of stroke survivors, contributing to the increase of the overall burden of stroke.<sup>10</sup>

There was variation in the annual reduction rates between studies, with Dijon<sup>16</sup> showing the least steep slope and Örebro<sup>18</sup>, Martinique<sup>15</sup> and Porto<sup>13</sup> showing the most prominent reduction. In other studies based on administrative data, one Swedish study and two Japanese registries also reported less marked reduction rates in the early 2000s than those from other high-income countries.<sup>6, 7, 43</sup> The overall difference in the time trends may reflect the difference by stroke subtypes between studies. Whilst some studies found consistent decline for all types of strokes,<sup>13, 15</sup> others reported diverging trend for ischaemic stroke versus haemorrhagic stroke.<sup>12, 17, 19, 44–46</sup> Moreover, although consistent decline for men and women was found in our pooled analysis and also reported in other hospital-based registries,<sup>39</sup> the Erlangen study only found a significant decline for men.<sup>14</sup> Given that the included population-based studies had broadly similar ascertainment methods and also maintained consistent methodology over time, the observed differences between studies would appear to be real and may be accounted for by differences in life expectancy, risk factor prevalence, control of risk factors and accessibility of health services.<sup>30</sup>

Notably, in contrast to a steady decline of stroke incidence for ischaemic stroke, our estimate suggested less marked change of stroke incidence for intracerebral haemorrhage in more recent years. This apparent divergent trend could perhaps be explained by increasing use of antithrombotic treatment at older ages, especially with the increasing burden of atrial fibrillation.<sup>44</sup>

Both OXVASC and the Porto stroke registry<sup>13</sup> found no significant reduction in non-disabling stroke in the past 10 years. Cerebrovascular events could have become less severe due to implementation of preventive treatment, or less disabling due to improved acute stroke care. On the other hand, increasing public awareness of stroke symptoms and growing use of neuroimaging and diagnostic awareness among healthcare providers might have also resulted in improving ascertainment of minor strokes over time.

Although we consider our findings to be valid, our study has limitations. Firstly, although we confined our analysis to high-quality population-based stroke incidence studies that maintained the “ideal” methodology over time, completeness of ascertainment might have changed over time.<sup>47</sup> However, we found a consistent and steady decline across all studies. Secondly, we only had aggregated data from published results and were therefore unable to assess thoroughly potential reasons for the observed overall temporal trends or the heterogeneity between studies, particularly if any demographic change, such as change in occupation or standard of living, had any role in explaining the observed trends across

studies. Thirdly, proportion of unclassified stroke differed considerably between studies, with further variation over time. Therefore it was difficult to provide accurate pooled estimates comparing the temporal change of incidence in ischaemic versus haemorrhagic strokes. Moreover, most studies did not provide data on the change of aetiological subtypes of ischaemic strokes over time. We were also unable to compare the incidence change by stroke aetiology. However, one recent systematic review of population- and hospital-based studies suggested an increasing trend of cardioembolic stroke and a decrease for lacunar stroke in high-income countries in the last 30 years.<sup>48</sup>

In conclusion, stroke incidence is continuing to decline in Oxfordshire and in other high-income settings in the last 30 years. However, with the aging population, even if the age-specific stroke incidence continued to decrease at its current rate, the number of new stroke cases annually in high-income countries would continue to increase in the next 30 years.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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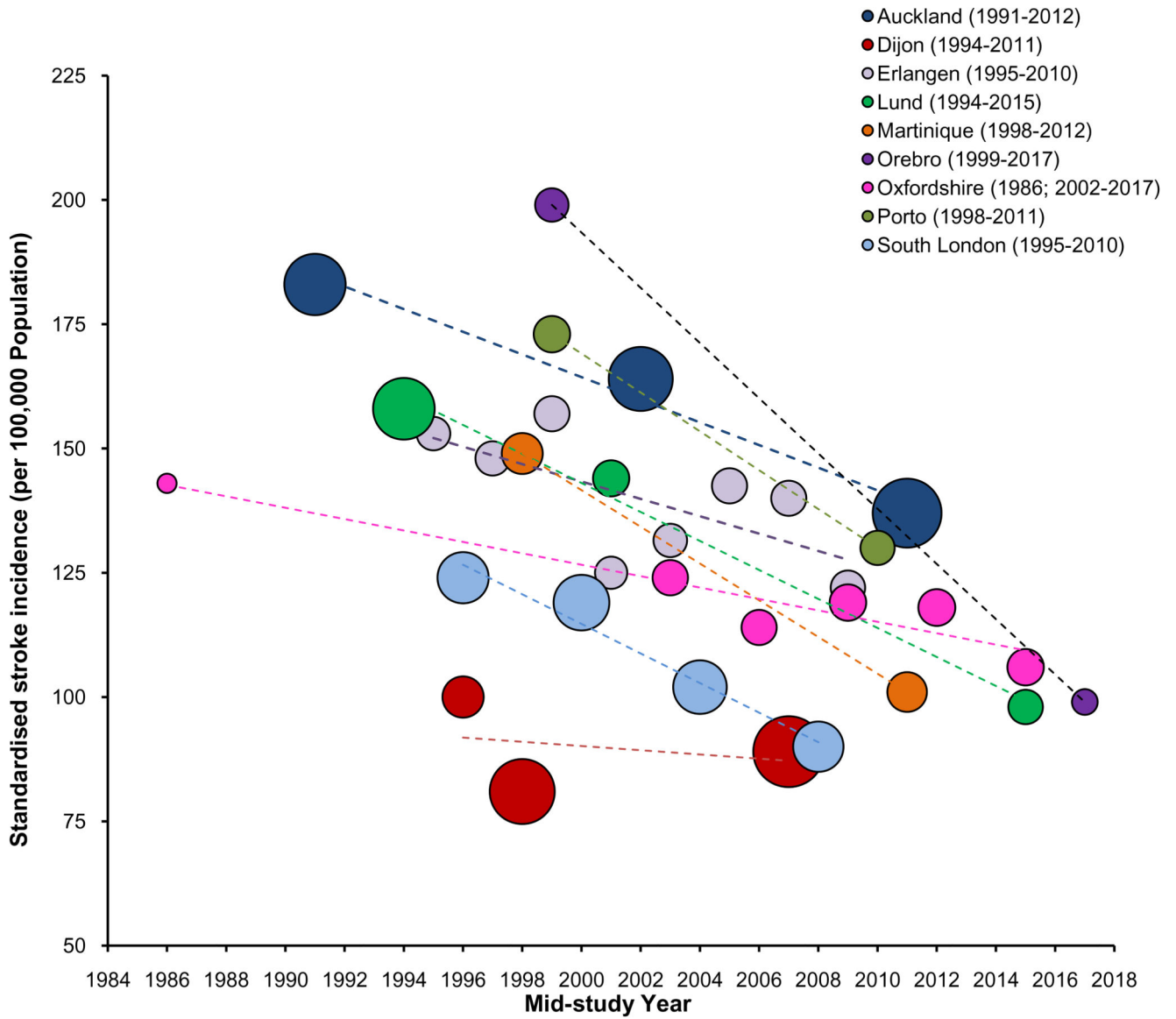
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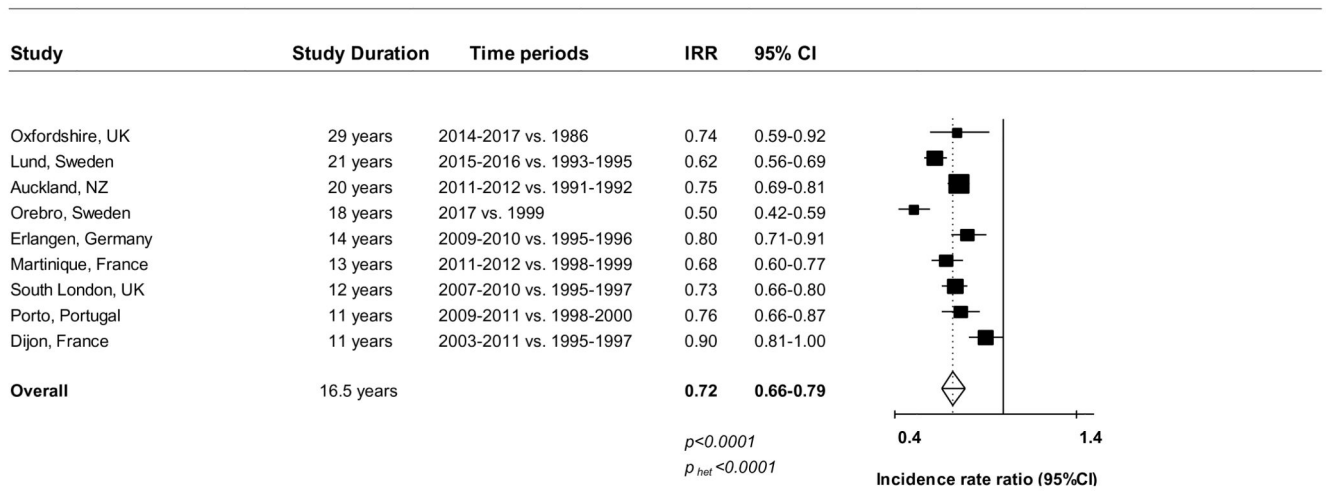
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**Figure 1.** Temporal trends of standardised stroke incidence in population-based studies reporting at least one time point after year 2010 (1990s-2010s)



IRR=Incidence Rate Ratio.

**Figure 2.**

Meta-analysis (random effects) of standardised incidence rate ratio (temporal trend) in population-based studies of first-ever stroke (1990s – 2010s)

†Raw numbers of incident stroke cases for each study for the two time periods included are presented in table 1.

IRR=Incidence Rate Ratio.

### A. Change of the proportions of disabling or fatal stroke

Study	Disabling or fatal / Total (%)		OR	95% CI
	Later periods	Early periods		
Oxfordshire, UK (2014-2017 vs. 2002-2005)	185 / 462 (40.0)	201 / 430 (46.7)	0.76	0.58-0.99
Porto, Portugal (2009-2011 vs. 1998-2000)	122 / 204 (59.8)	169 / 260 (65.0)	0.80	0.55-1.17
<b>Total</b>	307 / 666 (46.1)	370 / 690 (53.6)	0.77	0.62-0.96

$p=0.02$   
 $p_{het}=0.83$

**B. Change of IRR** †Raw numbers of incident stroke cases for each study for the two time periods included can be derived from figure 3A.

Study	Study Duration	Time periods	IRR	95% CI
<b>Disabling or fatal</b>				
Oxfordshire, UK	12 years	2014-2017 vs. 2002-2005	0.68	0.55-0.85
Porto, Portugal	11 years	2009-2011 vs. 1998-2000	0.71	0.60-0.84
<b>Overall</b>	11.5 years		<b>0.70</b>	<b>0.61-0.80</b>
$p<0.0001$ $p_{het}=0.72$				
<b>Non-disabling</b>				
Oxfordshire, UK	12 years	2014-2017 vs. 2002-2005	1.02	0.85-1.23
Porto, Portugal	11 years	2009-2011 vs. 1998-2000	0.91	0.73-1.13
<b>Overall</b>	11.5 years		<b>0.98</b>	<b>0.85-1.12</b>
$p=0.73$ $p_{het}=0.44$				

IRR=Incidence Rate Ratio.

**Figure 3. Meta-analysis (random effects) of standardised incidence rate ratio (temporal trend) in population-based studies of first-ever stroke stratified by stroke severity**  
IRR=Incidence Rate Ratio.

A. Change of the proportions of disabling or fatal stroke

B. Change of IRR †Raw numbers of incident stroke cases for each study for the two time periods included can be derived from figure 3A.

**Table 1**  
**Study characteristics and incidence (95%CI) of first-ever stroke (per 100,000 population) in population-based studies included in main analyses stratified by study periods**

Study (country)	Study duration	Number of stroke	Person-years	Age range	Crude incidence	Standardised incidence <sup>‡</sup>	Standardised population <sup>‡</sup>	Standardised to European population <sup>‡</sup>
Auckland, NZ <sup>11, 12</sup>	1991-1992	1305	945369	15y	181 (171-191)	156 (145-167)	WHO	183 (181-185)
	2002-2003	1423	897882	15y	158 (150-167)	139 (132-147)	WHO	164 (162-166)
	2011-2012	1643	1119192	15y	147 (140-154)	119 (114-125)	WHO	137 (136-138)
Dijon, France <sup>16, 30</sup>	1985-1993	1302	1316497	All	99 (94-104)	77 (72-81)	European	77 (72-81)
	1994-2002	1462	1346085	All	109 (103-114)	81 (76-85)	European	81 (76-85)
Martinique, France <sup>15</sup>	1995-1997	591	429264	All	137 (127-149)	67 (61-73); 100 (92-109)	WHO; European	100 (91-109)
	2003-2011	1742	1361911	All	128 (122-134)	89 (84-93)	European	89 (84-93)
	1998-1999	580	362256	All	160 (149-172)	111 (102-120)	WHO	149 (137-161)
Porto, Portugal <sup>13</sup>	2011-2012	544	370854	All	147 (134-159)	77 (70-84)	WHO	101 (99-103)
	1998-2000	462	172046	All	269 (244-293)	173 (153-192); 261 (237-285)	European; Portugal	173 (153-192)
Erlangen, Germany <sup>14</sup>	2009-2011	405	204424	All	198 (179-217)	130 (114-146); 203 (183-223)	European; Portugal	130 (114-146)
	1995-1996	390	NR	All	NR	Female 130; Male 176	European	Female 130; Male 176
South London, UK <sup>29-31</sup>	1997-1998	407	NR	All	NR	Female 130; Male 166	European	Female 130; Male 166
	1999-2000	427	NR	All	NR	Female 126; Male 188	European	Female 126; Male 188
	2001-2002	359	NR	All	NR	Female 93; Male 157	European	Female 93; Male 157
	2003-2004	388	NR	All	NR	Female 124; Male 139	European	Female 124; Male 139
	2005-2006	428	NR	All	NR	Female 126; Male 159	European	Female 126; Male 159
	2007-2008	428	NR	All	NR	Female 106; Male 174	European	Female 106; Male 174
	2009-2010	411	NR	All	NR	Female 117; Male 127	European	Female 117; Male 127
1995-1997	911	703599	15y	129 (121-138)	82 (76-88); 124 (116-132)	WHO; European	124 (116-132)	
1995-1998	1303	NR	15y	NR	247 (234-261)	England & Wales	141 (133-149)	
1999-2002	1072	310026 (2001)	15y	NR	212 (199-225)	England & Wales	119 (111-127)	

Study (country)	Study duration	Number of stroke	Person-years	Age range	Crude incidence	Standardised incidence <sup>‡</sup>	Standardised population <sup>‡</sup>	Standardised to European population <sup>‡</sup>
	2003-2006	994	NR	15y	NR	175 (164-187)	England & Wales	102 (95-109)
	2007-2010	876	357308 (2011)	15y	NR	150 (139-160)	England & Wales	90 (84-95)
<b>Lund, Sweden</b> <sup>17, 27, 28</sup>	1993-1995	1318	678893	All	194 (184-205)	235 (222-248); 158 (150-167)	Swedish; European	158 (149-167)
	2001-2002	456	234505	All	194 (176-213)	230 (209-253); 144 (130-158)	Swedish; European	144 (130-158)
	2015-2016	413	276400	All	149 (135-165)	165 (149-182)	Swedish 2015	98 (88-108)
<b>Örebro, Sweden</b> <sup>18</sup>	1999	388	123503	All	314 (284-347)	344 (312-378)	Swedish 2017	199 (178-220)
	2017	228	150291	All	152 (133-173)	166 (146-188)	Swedish 2017	99 (85-113)
<b>Oxfordshire, UK</b> <sup>19</sup>	1986 <sup>*</sup>	128	86487	All	148 (123-176)	203 (167-239)	England & Wales	143 (118-168)
	2002-2005	430	273402	All	157 (143-173)	186 (168-204)	England & Wales	124 (112-136)
	2005-2008	428	275160	All	156 (141-171)	176 (159-193)	England & Wales	114 (103-125)
	2008-2011	463	283513	All	163 (149-179)	177 (161-193)	England & Wales	119 (108-130)
	2011-2014	471	286313	All	165 (150-180)	169 (154-184)	England & Wales	118 (107-129)
	2014-2017	462	294063	All	157 (143-172)	154 (141-169)	England & Wales	106 (96-116)

<sup>‡</sup> reported in original publication

<sup>‡</sup> Restandardised to European population based on raw numbers provided in original publication.

\* n=557 for incident stroke cases ascertained between 1981-1984/1986



**Table 2**  
**Meta-analysis (random effects) of standardised incidence rate ratio (temporal trend) in population-based studies of first-ever stroke stratified by stroke subtypes and by sex**

	No. of studies	Study duration (mean)	Pooled IRR	95%CI	p	phet
<b>Overall (1990s-2010s)</b>						
<b>Stroke subtypes</b>						
Ischaemic stroke	5	16 years	0.84	0.75-0.93	0.001	0.04
Intracerebral haemorrhage	5	16 years	0.63	0.55-0.73	<0.0001	0.43
Subarachnoid haemorrhage	5	16 years	0.60	0.36-1.00	0.05	0.002
<b>Sex</b>						
Men	8	17 years	0.69	0.61-0.77	<0.0001	0.003
Women	8	17 years	0.66	0.59-0.74	<0.0001	0.001
<b>1990-2000</b>						
<b>Stroke subtypes</b>						
Ischaemic stroke	5	10 years	0.83	0.78-0.88	<0.0001	0.66
Intracerebral haemorrhage	5	10 years	0.68	0.53-0.88	0.003	0.01
Subarachnoid haemorrhage	5	10 years	1.05	0.75-1.47	0.77	0.12
<b>Sex</b>						
Men	8	10 years	0.86	0.80-0.92	<0.0001	0.12
Women	8	10 years	0.80	0.74-0.86	<0.0001	0.11
<b>Post year 2000</b>						
<b>Stroke subtypes</b>						
Ischaemic stroke	7	10 years	0.83	0.76-0.91	<0.0001	0.002
Intracerebral haemorrhage	7	10 years	0.82	0.61-1.09	0.16	<0.001
Subarachnoid haemorrhage	7	10 years	0.67	0.47-0.96	0.03	0.001
<b>Sex</b>						
Men	8	11 years	0.76	0.69-0.83	<0.0001	0.003
Women	8	11 years	0.79	0.68-0.91	0.001	<0.0001

IRR=Incidence Rate Ratio.