Does the incidence, severity, or case fatality of stroke vary in southern England?

C D A Wolfe, N A Taub, J Woodrow, E Richardson, F G Warburton, P G J Burney

Abstract

Study objectives—To determine differences in incidence and case fatality of stroke in district health authorities with differing standardised mortality ratios (SMR) for stroke in residents aged under 65 years in whom death from stroke is considered ‘avoidable’.

Design—Registration of first ever strokes in three district health authorities. Patients were assessed and followed up over one year by one of three observers.


Participants—Patients under the age of 75 years having a first ever in a lifetime stroke between 15 August 1989 and 14 August 1990.

Measurements and main results—Age specific incidence rates and survival time from stroke to death. Severity was assessed in terms of the level of consciousness and the presence of speech, urinary, and motor impairment within the first 24 hours of the stroke.

Altogether 386 strokes were registered. There was a significant difference in the incidence rate between district health authorities in those aged under 65 (p<0.01).

Methods

Stroke registers were set up in three districts of the South East Thames Regional Health Authority—West Lambeth (WL) and Lewisham and North Southwark (LNS) are inner London districts and Tunbridge Wells (TW) is a rural district 40 miles from London. WL has a high SMR (134; based on 366 deaths), LNS a moderate SMR (108; based on 647 deaths), and TW a low SMR (70: based on 264 deaths) for hypertension and cerebrovascular disease in those aged under 65 years. The denominators for the incidence rates were the Office of Population Censuses and Surveys 1988 mid-year estimates. The total study population under the age of 75 years was 621 966, with 148 034 in WL, 293 364 in LNS, and 180 568 in TW. To detect a significant difference in case fatality rates at three weeks at the 5% level with a power of 0.8 when the true rates are 20% and 40%, at least 85 strokes would be required in each
of two districts. Similarly, to detect a difference in crude incidence rates when the true rates are 1-75 and 2-25 per thousand, populations of at least 130 000 are required in each district. Both these conditions were met by studying the three districts over a one year period. The register was piloted in WL for six months before this.

**SOURCES OF NOTIFICATION**

All general practitioners (GPs) with patients living in the three districts were contacted, sent notification booklets to enable them to register a patient with minimum inconvenience, and invited to a seminar to discuss the project. A bi-monthly newsletter was sent to each GP. This helped to maintain cooperation and also asked for further notification of new cases. A letter was sent to the GP after each case was detected and all GPs were phoned and visited once during the study period. A questionnaire was also sent at the end of the year asking about further cases not previously reported. Similar notification systems were arranged with the district nursing and rehabilitation services, which were contacted on a weekly basis, and consultant physicians, who were sent the newsletter. Three fieldworkers identified hospital patients by looking through casualty admission registers and visiting wards on a weekly basis. Hospital pathology and coroners' post mortem reports and district residents' death certificates were all inspected. The fieldworkers visited all hospitals in the districts and also, in London, the hospitals in the surrounding area to which residents of these districts were known to have been admitted in the past after strokes. Consequently, 28 hospitals were involved in the study.

**CRITERIA FOR REGISTRATION**

The WHO definition of a stroke was used for all cases of first ever in a lifetime stroke under the age of 75 years whether or not they survived the event. Details of community cases and sudden deaths were also obtained from GP records. All questionnaires were checked by CW to ensure entry criteria were fulfilled, and if the diagnosis was in doubt there was further discussion with the relevant clinicians before deciding on whether to include a case.

Information was collected from either the patient, the hospital or GP notes, health care workers, or relatives. Records were made of the date of stroke, demographic details, and an assessment of the patient's activities of daily living (ADL) before stroke using the Barthel 20 point scale and the Rankin 6 point scale for handicap. Clinical details of maximal impairment within the first 24 hours included the level of consciousness, paralysis, speech, and urinary or swallowing impairment. For cases of sudden death the level of consciousness was assessed as coma but other variables were coded as 'not known'. Details of the registered cause of death and post mortem examination results were recorded in patients who died.

**STATISTICAL METHODS**

Standardised incidence ratios were calculated as the ratio of the observed number of district registrations to that expected by applying the total study age specific incidence rates to the relevant district populations.

Log linear modelling was used to compare the proportions of deaths in the three districts caused by subarachnoid haemorrhage, adjusting for the different age distributions.

The survival curve was estimated using the Kaplan-Meier product limit technique, survival times being censored on 30 September 1990. The combined effect of baseline prognostic variables on survival time was examined using Cox's proportional hazards regression. This analysis was performed separately on data for the first three weeks after the stroke and for the subsequent period. The patients' age groups and districts of residence were included in the model. Other variables were selected by a backwards stepwise procedure which excluded variables that were not significantly predictive of survival at the 5% level. The variables considered for selection were: whether living alone before the stroke, ethnic group, Rankin and Barthel scores before the stroke, coma, paralysis, incontinence, speech and swallowing impairment, and gender. For each variable any 'not known' responses were considered as a separate category.

**Results**

The sources of notification are given in table I. Thirty-two (8%) of the cases were notified by two or more sources. Forty one (61%) of the strokes notified by GPs were community cases which were not admitted to hospital.

Ninety four per cent of practices responded to a questionnaire sent at the end of the study and 12 extra patients were registered. Overall 78% of patients were treated in hospital, with no significant difference between the districts ($X^2=2.7, p>0.1$).

Table I shows the number of strokes in each age group, the incidence rates per 1000 population, and the 95% confidence intervals. The standardised incidence ratio in the under 65s varied from 1-48 in WL to 0-96 in LNS and 0-73 in

<table>
<thead>
<tr>
<th>Table I</th>
<th>Notifications from each source. Numbers (%)*</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Notification source</strong></td>
<td><strong>West Lambeth</strong></td>
</tr>
<tr>
<td>General practitioner</td>
<td>19 (19)</td>
</tr>
<tr>
<td>District nurses</td>
<td>6 (6)</td>
</tr>
<tr>
<td>Rehabilitation services</td>
<td>21 (21)</td>
</tr>
<tr>
<td>Hospital staff</td>
<td>2 (2)</td>
</tr>
<tr>
<td>Death certificates/coroner</td>
<td>17 (17)</td>
</tr>
<tr>
<td>Register/fieldworker</td>
<td>53 (54)</td>
</tr>
<tr>
<td>Others</td>
<td>4 (4)</td>
</tr>
</tbody>
</table>

*Percentages do not sum to 100 because some patients were notified more than once.

<table>
<thead>
<tr>
<th>Table II</th>
<th>Number of strokes (men and women), incidence rates (1000 population), and 95% confidence intervals (CI) for the incidence rates</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Age group (yr)</strong></td>
<td><strong>West Lambeth</strong></td>
</tr>
<tr>
<td>&lt;45</td>
<td>11</td>
</tr>
<tr>
<td>45-54</td>
<td>9</td>
</tr>
<tr>
<td>55-64</td>
<td>20</td>
</tr>
<tr>
<td>65-74</td>
<td>49</td>
</tr>
<tr>
<td>Total</td>
<td>98</td>
</tr>
</tbody>
</table>
TW. There was a significant difference in incidence between the districts in those patients aged under 65 years ($\chi^2 = 11.4, p < 0.01$). There was no significant difference between districts in the 65-74 year old age group ($p > 0.1$).

The mean (SD) age of patients was 64.2 (10.1) years with a median of 67.5 years. Stroke patients in WL were younger (median ages: WL=65.1, LNS=67.8, and TW=68.4; Kruskall-Wallis test, $p=0.05$). There was an excess of men (218 of 386, (56%)). Overall, 24% of subjects lived alone before the stroke. The proportion of stroke patients who were black varied between districts, WL having 20%, LNS 11%, and TW no black patients (table III).

Table III  Distribution of cases by age, ethnic group (white (W) or black (B), and health district (16 patients of unknown ethnic group are not included)

<table>
<thead>
<tr>
<th>No of patients</th>
<th>West Lambeth</th>
<th>Lewisham &amp; N Southwark</th>
<th>Tunbridge Wells</th>
<th>All districts</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (s)</td>
<td>W  B (%)</td>
<td>W  B (%)</td>
<td>W  B (%)</td>
<td>W  B (%)</td>
</tr>
<tr>
<td>&lt;45</td>
<td>6 3 (33)</td>
<td>9 1 (10)</td>
<td>4 0 (0)</td>
<td>19 4 (17)</td>
</tr>
<tr>
<td>45-54</td>
<td>8 3 (43)</td>
<td>8 3 (27)</td>
<td>5 0 (0)</td>
<td>17 6 (26)</td>
</tr>
<tr>
<td>55-64</td>
<td>20 7 (26)</td>
<td>39 6 (13)</td>
<td>26 0 (0)</td>
<td>85 13 (13)</td>
</tr>
<tr>
<td>65-74</td>
<td>40 5 (11)</td>
<td>99 9 (8)</td>
<td>73 0 (0)</td>
<td>212 14 (6)</td>
</tr>
<tr>
<td>Total</td>
<td>70 18 (20)</td>
<td>155 19 (11)</td>
<td>108 0 (0)</td>
<td>333 37 (10)</td>
</tr>
</tbody>
</table>

The Barthel and Rankin scores before stroke are given in table IV. Over 90% of patients were functionally independent before the stroke.

CASE FATALITY

During the study period, 127 (33% of the 386) stroke patients died. The mean (SD) age at death was 63.5 (10.5) years. The median length of time from stroke to death was 2.5 days in WL, 4 days in LNS, and 6.5 days in TW (Kruskall-Wallis test, $p=0.01$).

Thirty two patients died on the day of their stroke. The overall three week case fatality was 26% (99 out of 386) and there was no significant difference between districts for unadjusted three week case fatality ($\chi^2=0.1, p=0.1$). There was no significant difference in case fatality between districts after standardisation for differences in age structure ($\chi^2=0.3, p=0.1$). Twenty one (17%) deaths were recorded as being a result of subarachnoid haemorrhage, with no difference between districts ($p=0.1$) (WL:5, LNS:11, TW:5).

Thirty per cent (38) of patients who died underwent an autopsy and a diagnosis of stroke was verified in all cases.

SEVERITY INDICATORS

Twenty five (26%) patients in WL, 36 (20%) in LNS, and 16 (15%) in TW were comatose ($\chi^2=3.9, p=0.04$). Twenty one (5%) patients were known to have had a paralysis or weakness before the stroke and 277 (72%) were paralysed as a result of the stroke. Four per cent were not paralysed and for 71 (18%) (55 of whom were in a coma) assessment was not possible. Two hundred and twenty six (59%) patients had a speech impairment, 60 of whom were in coma, and 103 (27%) had a swallowing impairment, 58 of whom were in coma. Sixty three (16%) patients were incontinent and 69 (18%) had a urinary catheter, of whom 48 were in coma.

There was no district difference in the proportions with coma ($\chi^2=0.9, p=0.1$), paralysis ($\chi^2=1.6, p=0.1$), or incontinence ($\chi^2=0.9, p=0.1$). There was a significant difference in initial speech impairment ($\chi^2=9.4, p=0.009$) with TW having the highest proportion (WL 58%, LNS 74%, TW 78%).

SURVIVAL ANALYSIS

The Kaplan–Meier survival curve for all patients is given in the figure.

All comatose patients were assumed to have paralysis, speech, urinary, and swallowing problems. Cox’s proportional hazards regression showed significant combined effects of age, incontinence, swallowing, and coma in the first three weeks and of all these variables except incontinence in the subsequent period (table V).

Discussion

This study has used a stroke register to investigate the variation in SMR in residents aged under 65 years for three district health authority populations and has fulfilled Malmgren’s criteria for a register, except for recording the diagnostic categories of stroke.8 16 This study was large, encompassing a population of over 600 000 district residents and involving 500 GPs and 28 hospitals. No previous studies in the United Kingdom have tried to explain regional variations in mortality from stroke or have used district health authority populations. The methodology used does not involve neurological assessment and computed tomography as part of the protocol but relies on local clinicians to make the diagnosis using investigations they consider appropriate. Comparisons with studies employing more detailed clinical methods, including computed tomograms, should therefore be made cautiously. This study did, however, use the WHO definition of stroke as recommended by Malmgren.8

Although no formal attempt to validate or standardise the diagnosis of stroke between districts was made, each case was assessed by one of the registry team who liaised with the relevant clinicians when fulfillment of the WHO criteria was in doubt. The diagnosis of stroke was verified in all cases that underwent autopsy. This methodology could be reproduced in health districts with a high SMR for stroke, including inner city districts, without the need for protocols necessitating expensive computed tomography and detailed neurological examination, which although contributing to the diagnosis, may not be available for all patients. Although the overall population incidence cannot be assessed in this study, more meaningful age-specific stroke incidence rates can be calculated and compared with other studies.
Kaplan-Meier survival curve for six months after first stroke

An initial hypothesis for this study was that the mortality rates in the under 65 year olds might be explained by differences in the incidence of stroke in the three districts. The standardised incidence ratios for first in a lifetime strokes in the under 65s were 148 in WL, 93 in LNS, and 73 in TW, indicating significantly different incidence rates (p<0.01).

This and the lack of any difference in case fatality suggests that mortality rates in this age group largely reflect the local incidence of strokes. However, these results have to be treated with due caution. We have studied only three health districts in a single region and a broader more representative sample of districts might produce different results.

In residents aged over 65 years there seems to be no significant difference in incidence between the districts. Routine data from the Office of Population Censuses and Surveys shows the age specific mortality rate for stroke in the over 65 year olds is highest in Tunbridge Wells, the converse to the findings in the under 65s.

A large proportion of strokes occurs in young blacks in the two inner city districts. This may be the result of the presence of specific risk factors for stroke in this group and particular problems in managing hypertension have been identified in African and West Indian patients in West Lambeth. The incidence of stroke might then be expected to increase in these districts as the black population ages.

The results do illustrate the use of population registers to assess the prevalence of risk factors among stroke patients which could enable preventive strategies to be more effectively focused.

The incidence rates estimated from this study are lower than those quoted from some other comparable studies. Rates quoted by Malmgren, from studies that she regarded as technically acceptable, range from 1/7 to 3/6/100 per year for 55-64 year olds compared with 1/49 in this study, and from 4/9 to 6/9 for 65-74 year olds compared with 3/97 in this study. Many of the studies quoted by Malmgren, however, were undertaken over two decades ago and the incidence of stroke has been declining, at least in the USA. More recent European studies quote age specific rates which are similar to those in the present study, but all studies suffer from having rates with wide confidence intervals, especially in the young age groups.

The Oxford stroke register was included in Malmgren's studies and was the closest in time as well as place to the current study. The study serves as a comparison with ours, although different aims, methodologies, and populations make such comparisons problematic. It also reported higher incidence rates than the present study. The Oxford study, as part of the protocol, had a higher rate of computed tomography, compared with 88% compared with 33% in the present study. It has been argued by some that computed tomograms may increase the recorded incidence of stroke but stroke is essentially a clinical diagnosis that has been shown to be reliable.

The SMR for stroke in those aged under 65 years has been consistently higher in the Oxford region than in both South East Thames Regional Health Authority overall and in inner London specifically over the past decade. A higher incidence of stroke in Oxford would therefore be consistent with our finding that the incidence seemed to play a major role in determining the mortality from stroke. Oxford is also geographically different to the inner city and rural districts in this study and the study was based on selected general practices rather than health district populations.

If under-reporting of strokes had occurred in the present study, we might have expected those cases that were reported to have been the more severe. Though the 26% case fatality at three weeks is higher than in Oxford and recent studies in Italy and France, it is similar to other studies. Previous studies have not, however, published age specific case fatality rates and it is therefore not possible to compare the case fatality rate of this study of residents aged under 75 years with all age case fatality rates published previously. There was no difference in age standardised case fatality rates, and this suggests that there was no difference in case ascertainment between districts. Similarly the proportion of deaths caused by subarachnoid haemorrhage was identical in the three districts. To assess further whether the relatively low incidence rates are

<table>
<thead>
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<th>Variable</th>
<th>Category</th>
<th>b (95% CI)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
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<td>Age:</td>
<td>≤54</td>
<td>0.00</td>
<td>1.00</td>
</tr>
<tr>
<td></td>
<td>55-64</td>
<td>0.84</td>
<td>0.33</td>
</tr>
<tr>
<td></td>
<td>65-69</td>
<td>0.27</td>
<td>0.36</td>
</tr>
<tr>
<td></td>
<td>70-74</td>
<td>0.15</td>
<td>0.36</td>
</tr>
<tr>
<td>District:</td>
<td>West</td>
<td>0.00</td>
<td>1.00</td>
</tr>
<tr>
<td></td>
<td>Lambeth</td>
<td>0.18</td>
<td>0.26</td>
</tr>
<tr>
<td></td>
<td>L and N</td>
<td>0.32</td>
<td>0.30</td>
</tr>
<tr>
<td></td>
<td>Southwark</td>
<td>0.00</td>
<td>1.00</td>
</tr>
<tr>
<td></td>
<td>Tunbridge</td>
<td>0.00</td>
<td>1.19</td>
</tr>
<tr>
<td></td>
<td>Wells</td>
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<td>1.00</td>
</tr>
<tr>
<td></td>
<td>Yes</td>
<td>1.97</td>
<td>0.56</td>
</tr>
<tr>
<td></td>
<td>No</td>
<td>0.00</td>
<td>1.00</td>
</tr>
<tr>
<td>Swallowing</td>
<td>Yes</td>
<td>1.61</td>
<td>0.43</td>
</tr>
<tr>
<td></td>
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<td>0.00</td>
<td>1.00</td>
</tr>
<tr>
<td>Impairment</td>
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<td>0.56</td>
</tr>
<tr>
<td></td>
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<td>1.00</td>
</tr>
<tr>
<td>Coma:</td>
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<td>1.97</td>
<td>0.56</td>
</tr>
<tr>
<td></td>
<td>No</td>
<td>0.00</td>
<td>1.00</td>
</tr>
</tbody>
</table>

*p value = significance of overall association of variable

Table V Cox proportional hazards regression for survival after first time stroke

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recorded in this study were the result of underreporting. We have undertaken one further study. A survey of GPs and district community rehabilitation and nursing staff at the end of the year elicited a response from 94% of practices and identified only 12 further patients, and this was after a year of intense communication with these groups.

To enable comparison of case fatality with other registers, assessment of the clinical state at the time of maximum disability within the first 24 hours was used. Ratings were allocated along with assessments of activity of daily living before the stroke. These and the age, living conditions before the stroke, ethnic group, and district of residence were used to predict outcome at three weeks and in the subsequent period. Proportional hazards regression analysis of survival allows the joint effects of prognostic factors on survival to be estimated.

The Rankin and Barthel scores before the stroke showed that over 90% of patients were functionally independent, which agrees with previous studies, but the level of disability before the stroke was not a significant independent predictor of survival.

Age had a significant effect on survival in both time periods but in the first three weeks the highest risk was to 55–64 year olds. In the subsequent period there was consistently increasing risk with age. The prediction that a comatose patient has a poor survival was consistent with other studies.

Both coma and the other indicators of severity, including incontinence and swallowing impairment were individually a predictive of poor survival, especially in the first three weeks. However, district of residence and race were not significantly related to survival.

The approach taken in this study uses the minimum of resources and is readily adaptable to urban or rural areas. This study has shown that the SMRs for hypertension and stroke in three districts reflect the incidence rates and that case fatality did not contribute to the variation between districts. This information should alert public health physicians to the importance of assessing the prevalence of risk factors in their districts. The study has shown that by collecting demographic and clinical data on all incident cases of stroke in a defined population, a clear picture of the prevalence of selected risk factors in the patients can be built up. This information will allow the purchasers of health care to target, through contracts, those at highest risk of stroke. For example, in West Lambeth the young black community has a high incidence of stroke which is partly caused by uncontrolled hypertension. The results of this study shows the need for effective health education and other primary preventive strategies to reduce the incidence of hypertension and stroke in this group. Furthermore, these types of initiatives are also the focus of the British government's recent consultative white paper which aims to reduce the number of premature deaths from stroke.

We wish to thank the Chest Heart and Stroke Association and Department of Health for funding. We are most grateful to all those who notified the register of cases, especially those in the community. Our thanks to Ana Chalda for typing the manuscript.

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