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Abstract

Many patients with minor stroke are referred to outpatient clinics and are not scanned immediately. A clinical rule is needed to identify patients who are likely to have intracerebral haemorrhage (ICH) and require urgent brain imaging, and patients who can safely start antiplatelet agents prior to scanning.

Methods: We determined clinical factors associated with ICH in 334 consecutive patients with minor stroke (NIHSS≤3) in the OXVASC Study, and derived a predictive model for ICH which was validated in a cohort of 280 patients presenting to a hospital-stroke clinic. Prognostic value was quantified as the area under the ROC curve (c statistics).

Results: The rate of ICH in minor stroke was 5.1% (95%CI 3.2-8.0%) in OXVASC, and 5.4% (3.3-8.7%) in the clinic cohort. Clinical factors predictive of ICH in OXVASC included BP on initial assessment \geq 180/110mmHg (OR=14.5, 95%CI 1.8-114, p=0.001), vomiting (OR=15.7, 5.4-46, p<0.001), confusion (OR=8.2, 2.9-23, p<0.001), and anticoagulation use (OR=7.8, 2.2-28, p=0.006), and at least one predictive factor was identified in all 17 patients with ICH and in 35% overall; c statistic 0.92 (95% CI 0.88-0.97). Therefore, we derived the SCAN rule to identify ICH if \geq 1 of the following were present: (S) severe hypertension, (C) confusion, (A) anticoagulation, (N) nausea and vomiting. In the clinic validation cohort, \geq 1 predictive factor was identified in 14/15 of patients with ICH and in 24% overall; c statistic 0.87 (0.79-0.95).

Conclusion: The SCAN rule appears to be specific and sensitive at identifying ICH in an independent cohort of patients with minor stroke, although further independent validations are needed.

Introduction

In the UK and elsewhere,^{1,2} many patients with minor stroke are not admitted to hospital but are assessed and investigated in specialist clinics. Although current guidelines recommend that all patients with TIA and minor stroke should be seen in clinic within seven days of symptom onset,³ currently only 35% of patients in the UK are seen within this time and the median delay to assessment is around 12 days,⁴ with many patients waiting substantially longer for brain imaging.

Delays to assessment and brain imaging in particular, can have a number of consequences. Firstly, referring physicians may be reluctant to start antiplatelet therapy until ICH has been excluded radiologically, even though the prompt initiation of secondary stroke prevention including antiplatelet agents is known to reduce early ischaemic stroke recurrence.^{5,6} Secondly, if late presenting patients are only scanned with CT which is insensitive to acute haemorrhage after 7-10 days,⁷⁻⁹ then cases with primary ICH may be missed. An earlier clinic-based study (of patients presenting > 4 days) using concurrent MRI and CT imaging showed that 75% of primary ICH identified on MRI were misdiagnosed as infarcts on CT because of delays to scanning,⁷ and might have been started on inappropriate and potentially harmful long-term prevention therapy with antithrombotic agents as a result. MRI was therefore recommended for all late-presenting patients, but this may be difficult to achieve when access to MRI is limited, and a system for identifying which patients with minor stroke are most likely to have had a recent ICH on clinical grounds would be useful.

Previous scoring systems have been created to identify patients with ICH,^{10,11} but these are weighted heavily on signs of major ICH including coma. These scores have been criticised for not being adequately sensitive in detecting ICH,¹² and are unlikely to be useful in patients presenting with minor stroke. We aimed to identify clinical factors which might predict the presence or absence of ICH in patients with minor stroke, and which could be used to determine which patients could be safely treated with antiplatelet agents before assessment in clinic, and which patients needed urgent brain imaging prior to treatment.

Methods:

Patients with minor stroke were prospectively ascertained from separate derivation and validation cohorts, defining minor stroke as any stroke meeting the World Health Organisation (WHO) criteria¹³ with a National Institute of Health Stroke Scale (NIHSS) score¹⁴ on assessment of \leq 3. The derivation cohort included consecutive patients prospectively ascertained from the first five years of the Oxford Vascular Study (OXVASC). OXVASC is a population-based study of all incident or recurrent acute vascular events including transient ischaemic attack (TIA) and stroke, in an Oxfordshire population of around 91,000 individuals registered with 63 family physicians, the methods of which have been described in detail elsewhere.¹⁵

The validation cohort comprised consecutive patients with minor stroke, who had been referred by their general practitioners from 2000-2006 to a "TIA" clinic in a district general hospital located in a neighbouring county. Nearly all of the patients in this cohort were scanned with MRI on the same day as the clinic assessment. The MRI scanner was a 1.5 Tesla Siemens Symphony system using an axial turbo gradient spin echo sequence (TGSE) (repetition time (TR) 4000ms, echo time (TE) 95 ms, 19 slices, slice thickness 6.0 mm, matrix 256x256, FOV 230x230).

In both cohorts, details of each patient's presenting symptoms and examination findings as well as background vascular risk factors, were recorded by a study neurologist. Stroke subtyping was based on the results of the first scan taken post stroke, and classified as being consistent with ischaemic stroke, ICH, or haemorrhagic transformation of infarction. For the purpose of later comparisons between ischaemic stroke and ICH, cases of haemorrhagic transformation of infarction were included in the group with ischaemic stroke, as most cases of haemorrhagic transformation of infarction only involve petechial haemorrhage into the region of infarction, and this relatively frequent "complication" is often asymptomatic¹⁶ and does not usually influence the decision to use antithrombotic therapy in the longer term. Cases of haemorrhage secondary to trauma, intracranial tumour, and haematopoietic malignancy, as well as cases of subdural and subarachnoid haemorrhage were excluded.

Data from the OXVASC cohort were used to derive a model to predict the presence of ICH on imaging. Because we expected a small number of patients with ICH in this cohort, we tried to minimise the possibility of chance associations, by testing variables which had been identified as predictors of ICH in previous clinical scores, and which appeared to be relevant to a population with minor stroke. These included variables which had featured in both the Allen score¹⁰ and Siriraj score,¹¹ such as headache or vomiting on presentation, and blood pressure at assessment. We also

looked for an association between ICH and a history of an acute confusional state with stroke onset, defined in accordance with DSM IV-R criteria¹⁷ as a history of acute onset reduced attention and disorganised thinking, disorientation or memory disturbance. This had not been included in the above scores, but acute confusional states have been shown to be associated with ICH by previous investigators.^{18,19} Blood pressure values were transformed into categorical variables for ease of use in the model, with cut-offs corresponding to the classification system of the joint European Societies of Cardiology and Hypertension²⁰ for increasing levels of hypertension. When variables were missing, these were assumed to be absent and the frequency of missing values never exceeded 5% of the cohort.

Univariate associations between the presence of ICH and clinical variables were evaluated using Student's t-test for continuous variables and Fisher's exact test for categorical variables. In addition, the usefulness of each clinical variable as a tool for differentiating between ICH and ischaemic stroke was compared using diagnostic odds ratios. There was insufficient statistical power to do multivariate modelling on the derivation cohort alone. Instead we continued to add variables associated with ICH with a significance of p<0.1 to the model in decreasing order of their diagnostic odds ratio, until addition of further variables did not appear to enhance the predictive value of the model.

This model was then internally validated using the OXVASC cohort and externally validated using the hospital clinic cohort. The frequency of ICH in each cohort was stratified according to the number of predictor variables present. Receiver operating characteristic (ROC) curves were then derived by determining the sensitivity and specificity of the model for identifying ICH with increasing numbers of predictor variables. In addition, we pooled data from both cohorts of patients of minor stroke, and analysed the independent predictive effect of the variables used in the model.

Finally, we wished to compare the performance of our model in both cohorts against an established and well-known model - the Siriraj score.¹¹ This score uses a cutoff of \leq -1 to predict infarction, and \geq 1 to predict ICH, so that brain imaging to detect ICH is needed if the score is > -1. As one of the clinical variables used in the Siriraj score is the patient's level of consciousness at assessment, and many patients in these cohorts presented late to outpatient clinics having already recovered, this variable was redefined as a history of drowsiness at the time of the stroke. The Allen score¹⁰ was not derived for either cohort as the score depends on examination findings including plantar responses at 24 hours post stroke, and such data were not complete in the hospital clinic based cohort.

Results

In the OXVASC derivation cohort, we identified 412 consecutive patients with minor stroke. Of these, 78 patients were scanned by CT beyond 10 days or declined brain imaging, and were excluded from further analysis as ICH could not be reliably diagnosed.⁷⁻⁹ Of the remaining 334 patients, who were either scanned with CT within 10 days (median interval (IQR) of 3 (1-5) days) or MRI, there were 17 patients with ICH (5.1%, 95%CI 3.2-8.0%), and 4 with haemorrhagic transformation of infarction (1.2%, 0.5-3.0%).

In the hospital clinic validation cohort, we identified 284 consecutive patients with minor stroke. Of these, four patients had undergone CT at >10 days post event because of contraindications to MRI, and were excluded from further analysis. Of the remaining 280 patients who were scanned with MRI on the day of clinic after a median delay of 15 days (IQR 10-23) following stroke onset, there were 15 patients with ICH (5.4%, 95% CI 3.3-8.7%), and 6 with haemorrhagic transformation of infarction (2.1%, 1.0-4.6%).

Baseline characteristics including risk factors, pre-morbid medication use, and stroke symptoms are shown for both cohorts in table 1. The OXVASC cohort was slightly older than the hospital clinic cohort, but had a similar ratio of male to female patients. The OXVASC cohort had a higher proportion of patients with a past history of TIA (p=0.002), but there were no other significant differences between the cohorts with respect to previously diagnosed comorbidities. Proportions on pre-morbid anticoagulation therapy were also similar, but more patients in OXVASC were taking antiplatelet agents (p=0.03). Headache (p=0.02) and vomiting (p=0.006) were more commonly identified in the OXVASC cohort, although these symptoms were no less frequent in outpatients compared to inpatients within this cohort. A similar proportion of minor strokes in both cohorts were associated with transient confusion at onset. The mean systolic blood pressure at presentation was higher in the OXVASC cohort (p<0.001), although the rates of previously diagnosed hypertension were similar in both cohorts.

Table 2 shows the diagnostic odds ratios for ICH versus ischaemic stroke for each of the variables tested in the OXVASC derivation cohort. Three clinical symptoms at stroke onset were predictive for ICH – vomiting (OR=15.7, 95% CI 5.4-46, p<0.001), confusion (OR=8.2, 2.9-23, p<0.001) and headache (OR=4.9, 1.8-13, p=0.002). Severe hypertension at initial assessment (SBP≥180mmHg or DBP≥110mmHg) was most predictive of ICH (OR=14.5, 1.8-114, p=0.001), and lower cut-off values

for blood pressure were less predictive. Pre-morbid anticoagulant use was the only pre-existing clinical risk factor associated with an increased likelihood of ICH (OR=7.8, 2.2-28, p=0.006). Age was not a discriminatory variable, with no difference in the mean age (SD) of patients with ICH or ischaemic stroke [74 (12) vs 74 (11) years, p=0.88]. The diagnostic odds ratios were derived for the same variables in the validation cohort (also shown in table 2). With the exception of a history of vomiting, all variables that were significantly associated with ICH in the derivation cohort were also significantly associated with ICH in the validation cohort.

A history of headache at stroke onset had the lowest diagnostic odds ratio of all the predictor variables that were significant on univariate testing in the derivation cohort, and adding this variable to our model did not improve the sensitivity, but reduced the specificity. Therefore we only selected the four variables which appeared to be most useful in the prediction of ICH - severe hypertension at initial assessment, pre-morbid anticoagulation, a history at presentation of transient confusion and vomiting at stroke onset. Table 3 shows the percentage risk of having ICH stratified according to the number of predictor variables in the OXVASC and hospital clinic cohort. In OXVASC, at least one clinical predictor was present in 116 (35%) patients with any minor stroke, but in all patients with ICH. If two or more clinical predictors were present approximately 42% of patients had evidence of a recent haemorrhage on scan.

In the hospital clinic validation cohort at least one clinical predictor was present in 66 (24%) patients, but in 14 (93%) patients with ICH. If two or more clinical predictors were present, 25% of patients had evidence of ICH on scan. ROC areas were 0.92 (95% 0.88-0.97) and 0.87 (0.79-0.95) for the derivation and validation cohorts respectively. In the pooled dataset comprising 31 cases of ICH, ICH was only identified in 0.2% of all patients in whom all clinical predictors were absent. In a multivariate analysis of the pooled dataset that included the four predictor variables identified in the derivation cohort, all remained independently predictive of ICH.

Based on these findings we derived the SCAN rule (table 4), in which scanning with expeditious CT, or MRI for late presenting patients, was recommended when one or more of the following variables were present: Severe hypertension at assessment (systolic blood pressure \geq 180mmHg or diastolic blood pressure \geq 110mmHg), Confusion at onset, Anticoagulation, Nausea and vomiting at onset. We then compared this rule to the Siriraj score which predicts absence of ICH in patients with a score of < -1. In the OXVASC cohort use of the Siriraj score would have missed 5 patients (29%) with ICH. However in the hospital TIA clinic cohort, only scanning patients with a Siriraj score of ≥-1, which is the recommended cut-off for identifying any potential cases with ICH, would have missed 13 patients (87%) with ICH.

Discussion:

We have derived and validated a rule in which the identification of at least one of four predictor variables: severe hypertension at assessment, pre-morbid anticoagulation, or a history of vomiting, or confusion at stroke onset, can be used to identify those patients with minor stroke who are most likely to have ICH. One or more of these predictor variables were present in nearly all patients with ICH, but were present in only 30% of all patients with minor stroke. A presentation with two or more predictor variables substantially increased the likelihood of ICH from an overall proportion of 5% to 38%. Furthermore the absence of any one of these variables made the likelihood of a recent ICH virtually negligible.

The consistency of prediction in both the derivation and validation cohorts suggests that the rule should be reliable and generalisable. Furthermore the clinical predictors of ICH in minor stroke have face validity. Some predictors such as a history of vomiting with stroke onset, or an elevated blood pressure at assessment, have been used in previous scores,^{10,11} albeit to predict ICH in patients with more severe stroke. Other predictors used by these scores, such as depressed conscious state, are clearly not relevant to a cohort with minor stroke, but we have shown that milder disturbances of the sensorium such as transient confusional states and ICH has also been shown in previous studies.^{18,19} Finally, anticoagulation use, while not a feature of previous scores, is known to be associated with a 2-4 fold increased risk of ICH²¹ and anticoagulation associated ICH now makes up a substantial proportion of all ICH in recent reports.²²

This rule may be used to guide the management of patients with minor stroke, many of whom are referred to specialist outpatient clinics and consequently face some delay to undergoing investigations including brain imaging. Because there is a lack of evidence that antiplatelet agents are not harmful in acute ICH, some referring physicians may be reluctant to start these agents in patients with minor stroke until ICH has been excluded radiologically. However the SCAN rule can be used to identify those patients in whom the likelihood of a recent ICH is virtually negligible, and who should benefit from the immediate introduction of antiplatelet therapy prior to assessment in the specialist clinic.^{5,6}

Conversely the SCAN rule identifies those patients in whom the likelihood of a recent ICH may be as high as 40%. If such patients present more than a week post stroke onset and are scanned with CT

there is a risk that an ICH will be misdiagnosed as ischaemic stroke⁷⁻⁹ resulting in inappropriate or potentially harmful long-term secondary prevention with antithrombotic therapy. Although the risks of antiplatelet and warfarin use post ICH are not absolutely defined, patients with ICH are three times more likely to have a further haemorrhagic stroke rather than an ischaemic stroke,²³ and if taking aspirin or warfarin prior to ICH the risk of fatality is increased.²⁴⁻²⁶ In particular, warfarin use post-ICH is estimated to reduce quality-adjusted life expectancy by up to two years.²⁷ In view of this risk, some clinicians have advocated using MRI to screen all late presenting patients for ICH.⁷ However this is not always practical in centres with limited access to MRI, and so the SCAN rule may be useful in prioritising the need for MRI. The SCAN rule may also be useful in prioritising the need for early CT imaging in developing countries or rural communities, where access to CT is limited.²⁸

While several clinical scores for predicting ICH already exist,²⁹⁻³³ the best known of which are the Allen and Siriraj scores,^{10,11} these have been developed using hospitalised cohorts of patients with more disabling strokes, and are not necessarily applicable to patients with minor stroke. For example, the Allen score is strongly weighted towards features of severe ICH including a history of loss of consciousness at stroke onset, and decreased consciousness or the presence of bilateral extensor plantar responses 24 hours post stroke. When scores have been independently validated they have often been criticised for being insensitive in detecting ICH,^{12,34,35} possibly because cases with less debilitating stroke were missed. When the Siriraj score was applied to our cohorts with minor stroke, 29% of cases with ICH were missed in the OXVASC cohort, and 87% in the hospital clinic based cohort. In contrast to this, the SCAN rule missed no cases with ICH in OXVASC and only 7% in the hospital clinic cohort. Most importantly the SCAN rule was validated in exactly the sort of population for which it should be most applicable – that is a cohort of late presenting outpatients.

Our study does have some limitations. The number of patients with ICH in the derivation cohort was small. Nevertheless, we were still able to demonstrate that several variables had a sizeable predictive effect as shown by their diagnostic odds ratios. While larger numbers with ICH would have made it possible to perform logistic regression analysis and further refine the model by weighting variables according to their corresponding regression coefficients, this might have produced a model that was well fitted to the derivation cohort but less generalisable to other populations with different exposures to different predictor variables. This approach would also have increased the complexity of the rule, making it less easy to remember and apply at the bedside. The model also appears to work better in the derivation cohort compared to the validation cohort. This is probably partly because patients were

seen earlier in the derivation cohort; and so recall of symptoms was more reliable than in the validation cohort. The one patient with ICH who was missed by the SCAN rule in the validation cohort was seen 23 days after stroke onset. Blood pressure measurements are also more likely to reflect immediate post-stroke measurements when performed earlier. Elevated post stroke systolic blood pressures do appear to discriminate between ischaemic stroke and ICH,^{30,31} and this appears to hold true for patients with smaller haemorrhages and less disabling stroke as shown in the OXVASC cohort. However blood pressure is highly variable and is more likely to be elevated just after stroke onset and fall again over the following days,³⁶ explaining why mean systolic blood pressures were higher in the OXVASC cohort compared to the hospital clinic based cohort. The model might have performed better in the validation cohort if the first ever blood pressure taken by primary care doctors or emergency department physicians had been available for use rather than blood pressure recordings taken in clinic. Finally some of the other predictor variables such as a history of vomiting and confusion might be considered as unlikely symptoms to encounter in an outpatient group. Nevertheless, these symptoms tended to be short-lived when they did occur, and there were as many inpatients as outpatients with a history of vomiting in the OXVASC cohort, and as many patients in the hospital clinic based cohort with a history of confusion as in the whole OXVASC cohort.

In conclusion, we have derived a simple rule – the SCAN rule for identifying patients with minor stroke who are most likely to have had a recent ICH. This rule is sensitive and reasonably specific on validation in an independent cohort of patients with minor stroke. Furthermore the validation cohort was representative of those patients for whom we believe this score will be most useful – i.e. patients referred for investigation and treatment in the outpatient clinic. Ideally all patients with minor stroke should be imaged and start appropriate secondary prevention therapy on the same day as symptom onset, but until major changes take place both in the way stroke care is delivered and in the public's recognition of the need for seeking immediate medical attention for stroke symptoms, a significant number of such patients will continue to present late. The SCAN rule, while still requiring further independent validation, should be more sensitive than previous scores at identifying which patients with minor stroke are very unlikely to have had a recent ICH when starting antiplatelet therapy prior to outpatient brain imaging, and which patients need urgent inpatient investigations or MRI to confirm a suspected ICH.

Ethical Approval: OXVASC and related sub-studies have been approved

by our local ethics review committee

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designed the hospital clinic cohort study. PMR designed the OXVASC study, and edited the paper

Competing Interests: None declared

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References:

- 1. Scholte op Reimer WJ, Dippel DW, Franke CL, et al. Quality of hospital and outpatient care after stroke or transient ischemic attack: insights from a stroke survey in the Netherlands. *Stroke* 2006;**37**(7):1844-9.
- Roebers S, Wagner M, Ritter MA, Dornbach F, Wahle K, Heuschmann PU. Attitudes and current practice of primary care physicians in acute stroke management. *Stroke* 2007;**38**(4):1298-303.
- 3. Intercollegiate working party for stroke. National Clinical Guidelines for Stroke 2004. London: Royal College of Physicians.
- 4. Intercollegiate Working Party for Stroke (2006). National Sentinel Stroke Audit. Phase 1 organizational audit. Clinical Effectiveness and Evaluation Unit, Royal College of Physicians.
- 5. Rothwell PM, Giles MF, Chandratheva A, et al. Effect of urgent treatment of transient ischaemic attack and minor stroke on early recurrent stroke (EXPRESS study): a prospective population-based sequential comparison. *Lancet* 2007;**370**(9596):1432-42.
- 6. Kennedy J, Hill MD, Ryckborst KJ, Eliasziw M, Demchuk AM, Buchan AM. Fast assessment of stroke and transient ischaemic attack to prevent early recurrence (FASTER): a randomised controlled pilot trial. *Lancet Neurol* 2007;6(11):961-9.
- Wardlaw JM, Keir SL, Dennis MS. The impact of delays in computed tomography of the brain on the accuracy of diagnosis and subsequent management in patients with minor stroke. *J Neurol Neurosurg Psychiatry* 2003;**74**(1):77-81.
- 8. Hung TP, Lee KY. Small intracerebral haemorrhage: a study of clinical manifestations and CT findings on 31 cases. *Ann Acad Med Singapore* 1985;**14**(1):22-31.
- Dennis MS, Bamford JM, Molyneux AJ, Warlow CP. Rapid resolution of signs of primary intracerebral haemorrhage in computed tomograms of the brain. *Br Med J* (*Clin Res Ed*) 1987;295(6594):379-81.
- 10. Allen CM. Clinical diagnosis of the acute stroke syndrome. *Q J Med* 1983;**52**(208):515-23.
- Poungvarin N, Viriyavejakul A, Komontri C. Siriraj stroke score and validation study to distinguish supratentorial intracerebral haemorrhage from infarction. *Bmj* 1991;**302**(6792):1565-7.
- Hawkins GC, Bonita R, Broad JB, Anderson NE. Inadequacy of clinical scoring systems to differentiate stroke subtypes in population-based studies. *Stroke* 1995;**26**(8):1338-42.
- 13. Cerebrovascular diseases: prevention, treatment, and rehabilitation. Report of a WHO meeting. *World Health Organ Tech Rep Ser* 1971;**469**:1-57.
- 14. Goldstein LB, Bertels C, Davis JN. Interrater reliability of the NIH stroke scale. *Arch Neurol* 1989;**46**(6):660-2.
- 15. Rothwell PM, Coull AJ, Giles MF, et al. Change in stroke incidence, mortality, casefatality, severity, and risk factors in Oxfordshire, UK from 1981 to 2004 (Oxford Vascular Study). *Lancet* 2004;**363**(9425):1925-33.
- 16. Mayer TE, Schulte-Altedorneburg G, Droste DW, Bruckmann H. Serial CT and MRI of ischaemic cerebral infarcts: frequency and clinical impact of haemorrhagic transformation. *Neuroradiology* 2000;**42**(4):233-9.
- 17. Diagnostic and Statistical Manual of Mental Disorders, 4th Ed, text reviewed, Washington DC: American Psychiatric Association, May 2000.
- 18. Sheng AZ, Shen Q, Cordato D, Zhang YY, Yin Chan DK. Delirium within three days of stroke in a cohort of elderly patients. *J Am Geriatr Soc* 2006;**54**(8):1192-8.

- 19. Caeiro L, Ferro JM, Albuquerque R, Figueira ML. Delirium in the first days of acute stroke. *J Neurol* 2004;**251**(2):171-8.
- 20. 2003 European Society of Hypertension-European Society of Cardiology guidelines for the management of arterial hypertension. *J Hypertens* 2003;**21**(6):1011-53.
- 21. Hart RG, Tonarelli SB, Pearce LA. Avoiding central nervous system bleeding during antithrombotic therapy: recent data and ideas. *Stroke* 2005;**36**(7):1588-93.
- 22. Flaherty ML, Kissela B, Woo D, et al. The increasing incidence of anticoagulantassociated intracerebral hemorrhage. *Neurology* 2007;**68**(2):116-21.
- Bailey RD, Hart RG, Benavente O, Pearce LA. Recurrent brain hemorrhage is more frequent than ischemic stroke after intracranial hemorrhage. *Neurology* 2001;56(6):773-7.
- Roquer J, Rodriguez Campello A, Gomis M, Ois A, Puente V, Munteis E. Previous antiplatelet therapy is an independent predictor of 30-day mortality after spontaneous supratentorial intracerebral hemorrhage. *J Neurol* 2005;252(4):412-6.
- 25. Saloheimo P, Ahonen M, Juvela S, Pyhtinen J, Savolainen ER, Hillbom M. Regular aspirin-use preceding the onset of primary intracerebral hemorrhage is an independent predictor for death. *Stroke* 2006;**37**(1):129-33.
- Rosand J, Eckman MH, Knudsen KA, Singer DE, Greenberg SM. The effect of warfarin and intensity of anticoagulation on outcome of intracerebral hemorrhage. *Arch Intern Med* 2004;**164**(8):880-4.
- Eckman MH, Rosand J, Knudsen KA, Singer DE, Greenberg SM. Can patients be anticoagulated after intracerebral hemorrhage? A decision analysis. *Stroke* 2003;**34**(7):1710-6.
- 28. Joubert J, Prentice LF, Moulin T, et al. Stroke in rural areas and small communities. *Stroke* 2008;**39**(6):1920-8.
- 29. Besson G, Robert C, Hommel M, Perret J. Is it clinically possible to distinguish nonhemorrhagic infarct from hemorrhagic stroke? *Stroke* 1995;**26**(7):1205-9.
- 30. Massaro AR, Sacco RL, Scaff M, Mohr JP. Clinical discriminators between acute brain hemorrhage and infarction: a practical score for early patient identification. *Arq Neuropsiquiatr* 2002;**60**(2-A):185-91.
- 31. Sturmer T, Schlindwein G, Kleiser B, Roempp A, Brenner H. Clinical diagnosis of ischemic versus hemorrhagic stroke: applicability of existing scores in the emergency situation and proposal of a new score. *Neuroepidemiology* 2002;**21**(1):8-17.
- Efstathiou SP, Tsioulos DI, Zacharos ID, et al. A new classification tool for clinical differentiation between haemorrhagic and ischaemic stroke. *J Intern Med* 2002;**252**(2):121-9.
- Woisetschlager C, Kittler H, Oschatz E, et al. Out-of-hospital diagnosis of cerebral infarction versus intracranial hemorrhage. *Intensive Care Med* 2000;26(10):1561-5.
- 34. Weir CJ, Murray GD, Adams FG, Muir KW, Grosset DG, Lees KR. Poor accuracy of stroke scoring systems for differential clinical diagnosis of intracranial haemorrhage and infarction. *Lancet* 1994;**344**(8928):999-1002.
- 35. Mader TJ, Mandel A. A new clinical scoring system fails to differentiate hemorrhagic from ischemic stroke when used in the acute care setting. *J Emerg Med* 1998;16(1):9-13.
- 36. Britton M, Carlsson A, de Faire U. Blood pressure course in patients with acute stroke and matched controls. *Stroke* 1986;**17**(5):861-4.

	(DXVASC	Hospital	l Clinic Cohort	р
		N= 334	1	N=280	
Mean age (SD)	74	(12)	72	(10)	0.04
Males (%)	195	(58)	166	(59)	0.87
Mean systolic BP (SD)	158	(30)	150	(23)	< 0.001
Mean diastolic BP (SD)	85	(15)	83	(12)	0.17
Premorbid risk factors (%)					
Hypertension	209	(63)	172	(61)	0.80
Myocardial infarction	39	(12)	49	(18)	0.05
Diabetes	33	(10)	41	(15)	0.08
Current smoker	54	(16)	51	(18)	0.52
Previous stroke	50	(15)	31	(11)	0.19
Previous TIA	58	(17)	24	(9)	0.002
Hyperlipidemia	91	(27)	64	(23)	0.23
Premorbid medications (%)					
Antiplatelet	148	(44)	99	(35)	0.03
Anticoagulation	16	(5)	12	(4)	0.85
Symptoms at onset					
Confusion	32	(10)	20	(7)	0.31
Vomiting	25*	(8)	7	(3)	0.006
Headache	68**	^c (20)	37	(13)	0.02

Table 1 Characteristics of patients with minor stroke in the OXVASC derivation cohort and hospital clinic validation cohorts.

* 12 of these were clinic patients, ** 35 of these were clinic patients

Table 2 Diagnostic odds ratios and 95% confidence interval for	predicting ICH versus ischaemic stroke in relation to potential clinical predictors in the OXVASC
derivation cohort and the hospital-clinic based validation cohort	

		Derivati	on Cohort			Validat	tion cohort	
	Infarct (n/%)	ICH (n/%)	Diagnostic OR	р	Infarct (n/%)	ICH (n/%)	Diagnostic OR	p
	N=317	N=17	(95%CI)	_	<mark>N=265</mark>	N=15	(95%CI)	
Demographic data								
Male	184 (58)	11 (65)	1.3 (0.5-3.7)	0.63	158 (60)	8 (53)	0.6 (0.2-1.7)	0.79
Mean age (sd)	74 (12)	74 (11)	-	0.88	72 11)	73 (5)	-	0.59
Symptoms								
Confusion	25 (8)	7 (41)	8.2 (2.9-23)	< 0.001	14 (5)	6 (40)	12.0 (3.7-38)	<mark><0.001</mark>
Vomiting	17 (5)	8 (47)	15.7 (5.4-46)	< 0.001	<u>6 (2)</u>	1 (7)	3.1 (0.3-27)	0.32
Headache	59 (19)	9 (53)	4.9 (1.8-13)	0.002	34 (13)	3 (20)	1.7 (0.5-6.3)	0.43
Vascular risk factors								
Hyperlipidemia	86 (27)	5 (29)	1.1 (0.4-3.3)	0.79	60 (23)	4 (27)	1.2 (0.4-4.0)	0.75
Previous MI	38 (12)	1 (6)	0.5 (0.1-3.6)	0.70	47 (18)	2 (13)	0.7 (0.2-3.3)	1.00
Previous stroke	46 (15)	4 (24)	1.8 (0.6-5.8)	0.30	28 (11)	3 (20)	2.1 (0.6-8.0)	0.22
Previous TIA	56 (18)	2 (12)	0.6 (0.1-2.8)	0.75	23 (9)	1 (7)	0.8 (0.1-6.0)	1.00
Current smoker	51 (16)	3 (18)	1.1 (0.3-4.0)	0.74	50 (19)	1 (7)	0.3 (0.0-2.4)	0.32
Hypertension	198 (63)	11 (65)	1.1 (0.4-3.1)	1.0	162 (61)	10 (67)	1.3 (0.4-3.8)	0.79
Pre-morbid medications								
antiplatelet ¹	141 (46)	3 (23)	0.4 (0.1-1.3)	0.15	95 (37)	4 (36)	0.9 (0.3-3.4)	1.00
anticoagulation	12 (4)	4 (24)	7.8 (2.2-28)	0.006	8 (3)	4 (27)	11.7 (3.0-45)	0.002
BP at assessment								
<140/90	81 (26)	1 (6)	1.0 (ref)		151 (58)	7 (47)	1.0 (ref)	
140/90 – 159/99 mmHg	95 (30)	2 (12)	1.7 (0.2-19)	1.00	14 (5)	0 (0)	-	1.00
160/100 – 179/109 mmHg	74 (23)	2 (12)	2.2 (0.2-25)	0.61	70 (26)	3 (20)	0.9 (0.2-3.7)	1.00
≥ 180/110 mmHg	67 (21)	12 (70)	14.5 (1.8-114)	0.001	30 (11)	5 (33)	3.6 (1.1-12)	0.04

¹ excludes individuals on warfarin

Number of predictor variables	Patients (%)	ICH(%)	% probability of ICH (95% CI)	
OXVASC				
0	218 (65%)	0 (0%)	0 % (0-2%)	
1	90 (27%)	6 (35%)	7 % (3-14%)	
<mark>≥1</mark>	116 (35%)	17 (100%)	15 % (9-22%)	
≥ 2	26 (8%)	11 (65%)	42 % (26-61%)	
Total	334 (100%)	17 (100%)	5 % (3-8%)	
Hospital clinic				
0	214 (76%)	1 (7%)	0.5 % (0-3%)	
1	58 (21%)	12 (80%)	21 % (12-33%)	
<mark>≥1</mark>	<u>66 (24%)</u>	14 (93%)	21 % (13-33%)	
≥2	8 (3 %)	2 (13%)	25 % (7-59%)	
Total	280 (100%)	15 (100%)	5 % (3-9%)	
Decled deterest				
Pooled dataset	100 (700)	1 (20)		
0	432 (70%)	1 (3%)	0.2 % (0-1%)	
1	148 (24%)	18 (56%)	12 % (8-18%)	
<u>≥1</u>	182 (30%)	31 (97%)	17 % (12-23%)	
≥2	34 (6%)	13 (41%)	38 % (24-55%)	
Total	614 (100%)	32 (100%)	5 % (4-7%)	

Table 3 Probability of ICH stratified by number of predictor variables in the OXVASC derivation and the hospital clinic validation cohorts

Table 4. The SCAN rule

Expedite CT brain imaging or arrange MRI for late-presenting patients with minor stroke if one or more of the following are present on history taking or assessment:

(S) Severe hypertension at presentation: SBP \geq 180 mmHg or DBP \geq 110mmHg

(C) Confusion

(A) Anticoagulation use prior to stroke onset

(N) Nausea and vomiting