

1 **Why do electronic health records reveal oral anticoagulant prescription after**  
2 **haemorrhagic stroke?**

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4 **Running head: Warfarin and Haemorrhagic stroke**

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46 Oral anticoagulation (OAC) is used to prevent stroke in patients with atrial fibrillation or heart  
47 valve disorders. However, OAC may cause haemorrhagic stroke. We completed a randomised  
48 trial of stroke secondary prevention [1], using primary care electronic health records (EHRs)  
49 collected into the UK Clinical Practice Research Datalink (CPRD). A small proportion of  
50 participants were prescribed warfarin following a haemorrhagic stroke diagnosis. The present  
51 research aimed to explain this observation.

52

53 Participants were selected from all 11,391 patients with prevalent stroke, from 106 family  
54 practices in England and Scotland, included in the trial [1]. Patients were included if they had a  
55 record of haemorrhagic stroke before the trial and were treated with warfarin during the trial.  
56 Family physicians were sent a postal questionnaire asking respondents about the nature of the  
57 stroke suffered by the patient and indications for warfarin therapy. Type of stroke was coded  
58 using the International Classification of Diseases 10<sup>th</sup> Revision [2]. Warfarin accounts for 99% of  
59 OAC prescriptions in CPRD. The project was approved by external scientific and ethical review  
60 committee (ISAC protocol 08\_083A2). GPs consented to their practice's participation. Fully  
61 anonymised data were used and no patient consent was sought.

62

63 There were 134 (1.2%) participants with EHRs documenting previous haemorrhagic stroke who  
64 were prescribed warfarin during the trial. The index stroke event was coded into the EHR as a  
65 sub-arachnoid haemorrhage for 25 (24%) of cases and as intracerebral haemorrhage (ICH) for 78  
66 (76%) of cases. There were 40 (47%) women and the mean age was 75 years (range 40 to 94  
67 years). Questionnaire responses were obtained for 103 (77%) patients with sufficient information  
68 to confirm the type of stroke obtained for 86 (64%) (Figure 1).

69

70 Questionnaire responses revealed that there were 41 (48%) patients with a physician-confirmed  
71 diagnosis of cerebral infarction (n=39) or other non-haemorrhagic stroke (n=2). The diagnosis  
72 was confirmed by imaging reports recorded for 36 (88%) of patients. There were five patients  
73 with a code for ischaemic stroke recorded after the initial haemorrhagic stroke but only two of  
74 these were within one year of the index stroke, possibly representing a clarification of the initial  
75 diagnosis. Indications for OAC in this group of participants included atrial fibrillation (n=27),  
76 disease of pre-cerebral and cerebral arteries (n=4), venous thromboembolism and other and not  
77 known (n=10).

78

79 There were 45 patients with a physician-confirmed diagnosis of haemorrhagic stroke including  
80 sub-arachnoid haemorrhage (n=17) and ICH (n=28). The diagnosis was supported by imaging  
81 results for 38 (84%) patients. Indications for OAC therapy included atrial fibrillation (n=20),  
82 venous thrombosis (n=10), pulmonary embolism (n=5), aortic valve replacement (n=3),  
83 percutaneous coronary intervention, vasculitis and not known (n=7).

84

85 We conclude that in the EHRs of a large population of stroke patients, there may be appreciable  
86 numbers with a diagnosis of haemorrhagic stroke who are treated with OAC therapy. We  
87 reported previously on the clinical coding of stroke in EHRs [3] and the potential for  
88 misclassification of stroke diagnoses. The present results should not be considered to evaluate  
89 the overall reliability of stroke coding in EHRs, because the sample was one in which data  
90 discrepancies were expected.

91

92 In about half of patients, the stroke event was mis-coded as a haemorrhagic stroke when the  
93 physician-reported diagnosis for the same event date was one of cerebral infarction.

94 Misclassification might sometimes result from haemorrhagic transformation, which is not  
95 necessarily a contra-indication to OAC therapy. Physicians may draw on sources of data from  
96 outside the EHR. Nevertheless, it is concerning that appreciable numbers of stroke patients are  
97 prescribed OAC therapy with incorrect diagnostic information coded into their EHRs, with  
98 potential medico-legal implications. Improved information sharing between providers would be  
99 of benefit to physicians.

100

101 A second group comprises patients with a confirmed diagnosis of haemorrhagic stroke who were  
102 treated with OAC therapy for co-existing thromboembolic disorders. This highlights the difficult  
103 balance of risks and benefits that may sometimes be required in clinical practice. However, the  
104 use of OAC therapy in patients with previous ICH remains controversial [4]. Improved evidence  
105 and decision support are required for these high-risk patients.

106

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121

122 **Figure 1 Legend: Flow chart showing physician-reported data for confirmed stroke type**  
123 **and indication for OAC therapy in 103 participants with electronic records for**  
124 **haemorrhagic stroke.**

125

Figure 1.

