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Original Research Article

Balance and Mobility as Predictors of Post-Stroke Cognitive Impairment

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Key Words

 $Stroke \cdot Post\text{-}stroke \ cognitive \ impairment \cdot Dementia \cdot Mild \ cognitive \ impairment \cdot Balance \cdot Mobility$

Abstract

Background: The number of patients with cognitive impairment following stroke is increasing due to the rise in the number of stroke survivors. Health authorities highlight the need for prediction and early diagnostics. The aims of this study were to investigate if balance and mobility may predict cognitive impairment 1 year after stroke. *Methods:* The participants were patients with first-ever stroke or transient ischaemic attack (TIA). The exclusion criteria were pre-stroke cognitive impairment and dementia. Measurements of balance comprised the Berg Balance Scale (BBS) and the Figure of Eight test (Fig8). Mobility was measured by maximum walking speed and the Timed Up and Go test. Dementia and mild cognitive impairment were merged into a main outcome: cognitive impairment. Unadjusted and adjusted multivariate logistic regression models were performed. Results: One hundred and eighty subjects performed balance and mobility measures at baseline, and 158 participated in the follow-up; 13 died and 9 did not complete the follow-up. Two variables made a significant contribution in the adjusted analyses (Fig8, BBS). The strongest predictor of cognitive impairment was Fig8 with an odds ratio of 1.06. Conclusion: The results of Fig8 and BBS measured in the acute phase of stroke were predictors of cognitive impairment 1 year later in this cohort of firstever stroke or TIA. © 2015 S. Karger AG, Basel

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Introduction

Every year, about 16 million people worldwide have a first-ever stroke [1]. Stroke considerably increases the risk of dementia, and the majority of patients suffering a stroke experience some degree of cognitive impairment [2, 3]. Cognitive impairment affects secondary prevention, rehabilitation, prognosis and quality of life after stroke [4]. The mechanisms involved in post-stroke cognitive impairment include vascular cognitive disease, degenerative disease and mixed pathologies [5]. Stroke is a heterogeneous disease, with substantial variability between patients in response to treatment after stroke depending on both neural injury as well as physical function [6]. Predictors of cognitive impairment due to cerebrovascular diseases are needed in order to improve the understanding of the aetiology, cognitive decline and the patients' overall prognosis [7]. Risk factors for post-stroke cognitive decline include stroke-related factors, non-modifiable factors such as ageing, and modifiable factors such as presence of chronic vascular disease, elevated blood pressure, diabetes, obesity, smoking, alcohol and dyslipidaemia [4].

There is a growing body of evidence showing an association between physical performance and cognition [8, 9]. The association between balance and gait and higher-level cognitive functioning can shed light on new ways to approach and understand treatment and early diagnosis of dementia [10]. Individuals with mild dementia have worse balance performance and decreased walking speed [11, 12]. Disturbances in gait are suggested as one of the earliest predictors of dementia [13–15]. Although walking speed is a relevant predictor of health outcomes [16], there is limited research of the possible prediction of cognitive impairment. Recently, a multicentre study suggested that the motoric cognitive risk syndrome, a combined assessment of walking speed and memory, identify those at risk of developing cognitive impairment [17].

Our understanding of how to predict and prevent cognitive impairment and dementia following a stroke is limited. A systematic review confirmed the importance of early diagnostics to provide customized interventions to vulnerable persons belonging to a risk group [18]. The trend with less severe strokes [19] provides less time for the patients in the stroke unit and thereby less opportunity for the staff to uncover cognitive impairment during hospital stay [20]. Hence it is of great interest to develop easily available clinical methods that may predict cognitive impairment at an early stage after stroke in order to be able to provide appropriate recommendations on medical care before discharge.

Low-cost disease prevention and early detection have been requested by the WHO [21] to form a basis for healthy ageing. The included measures of balance and mobility represent potential tools that may be valuable in detecting vulnerable subjects. Thus, finding the strongest motor predictor of cognitive impairment is requested in order to develop potential interventions. To our knowledge, no longitudinal study has examined if balance or mobility are predictors of cognitive impairment in a sample of acute stroke patients. The aims of this study were to investigate if measures of balance and mobility in the acute phase of stroke may predict cognitive function 1 year after stroke onset.

Methods

All patients with first-ever stroke or transient ischaemic attack (TIA) admitted to the stroke unit between March 2007 and July 2008 were invited to participate in this study. The exclusion criteria were pre-stroke cognitive impairment and dementia. We excluded patients with cognitive decline as indicated by a score \geq 3.7 on the informant Questionnaire on Cognitive Decline in the Elderly (IQCODE) [22] and patients with a life expectancy of less than





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a year as estimated by the treating physician. In addition, all participants in this study needed results on one of the balance and mobility measures at baseline. Sociodemographic characteristics reported were gender, age, education (more or less than 9 years) and vascular risk factors. Body weight and height were measured and body mass index calculated. Blood samples were analysed for the presence of the apolipoprotein e4 allele (ApoE4) [23].

Neurological impairment was measured using the National Institutes of Health Stroke Scale (NIHSS) on the 1st day after admittance. In addition, an experienced stroke physician classified patients with ischaemic stroke according to the Trial of Org 10172 in Acute Stroke Treatment (TOAST) classification [24] and the Oxfordshire Community Stroke Project (OCSP) classification [25].

The measurements assessing balance and mobility were performed within 1 week after stroke by dedicated physiotherapists working in the stroke unit. Balance was assessed using the Berg Balance Scale (BBS) and the Figure of Eight test (Fig8). BBS rates performance on a 5-level scale from 0 (cannot perform) to 4 (normal performance) on 14 different tasks involving functional balance control, including transfer, turning and stepping [26]. The total score ranges from 0 to 56. In Fig8, the subjects were asked to walk 'the figure of eight' twice at a given speed in their normal shoes. The number of steps made outside 'the figure of eight' was recorded [27]. Mobility was measured by maximum walking speed and the Timed Up and Go test (TUG). For the maximum walking speed test, subjects walked 10 m from a standing still position, and the time in seconds was registered [28]. For TUG, the patient was observed and timed while rising from an armchair, walking 3 m, turning, walking back and sitting down again [29]. BBS, TUG and maximum walking speed are commonly used to assess balance and mobility in patients who have suffered stroke, and have been tested according to reliability and validity [26, 29–32]. Fig8 has been tested for reliability [33] and has been used in other populations [27, 34].

At the 1-year follow-up, mild cognitive impairment (MCI) and dementia were diagnosed in accordance with the Guidelines at the Memory Clinic in our hospital. For dementia, we used the International Classification of Diseases 10th revision criteria [35], and for MCI the criteria outlined by Winblad et al. [36]. The diagnoses were based on the following information: the patient's medical history, the IQCODE, results of the cognitive assessments and information regarding the patient's daily functioning – all obtained at 12 months after stroke. The cognitive assessments included Mini-Mental State Examination [37], the Clock Drawing test [38], the Trail Making test A and B [39] and the 10-word test (max score 40), including delayed recall from the Repeatable Battery from the Assessment of Neuropsychological Status [40] and figures from the Alzheimer's Disease Assessment Scale-Cognitive subscale [41]. The incidence and subtypes of MCI and dementia have been reported elsewhere [5]. MCI and dementia were merged into a main outcome named cognitive impairment in the analyses of the data.

Statistical Analysis

Data were described with means and standard deviation for normally distributed variables and with proportions and percentages for categorical variables. Correlation analyses (Pearson's r) were conducted to examine the correlations between the physical performance variables. Univariate and multivariate logistic regression models were performed with the dichotomized cognitive impairment as the dependent variable and the physical performance measurements (BBS, Fig8, TUG and maximum walking speed) and covariates as independent variables. Logistic regression was fitted for each of the physical performance measurements and controlled for all significant covariates (p < 0.2) except independent variables correlating >0.60, which might interfere with the result, as inter-correlating variables [42]. We used odds ratios with 95% confidence intervals to compare the strength of the association between the various possible predictors and the main outcome cognitive impairment after stroke. The Hosmer-Lemeshow test for logistic regression was used to check if the model fitted the data [43].





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Table 1. Baseline characteristics (n = 180)

Demographics		
Age, years	72.1±12.2	
Females	88 (48.1)	
Living arrangement, cohabitant	122 (66.7)	
Education >9 years	140 (76.5)	
Stroke characteristics		
NIHSS, points	4.3 ± 6.5	
TOAST ¹	47 (26.1)	
OCSP ²	42 (23.3)	
Lesion, right	84 (46.7)	
Risk factors		
BMI	25.6±4.2	
Currently smoking	39 (21.3)	
Hypertension	110 (60.1)	
ApoE4	3 (1.7)	
Measures of balance and mobility		
MWS $(n = 155)$, s	8.8 ± 2.0	
TUG ($n = 152$), s	11.8±7.5	
Fig8 (n = 121), missteps	18.1±17.8	
BBS (n = 178), points	44.5±17.6	

Values are presented as n (%) or mean ± SD. BMI = Body mass index; MWS = maximum walking speed.

Results

A sample of 180 participants admitted to the hospital with first-ever stroke or TIA were included in this study. A total of 158 of these participated in the registrations 1 year later; 13 died and 9 refused or were not able to complete. The baseline characteristics are given in table 1.

The results showed that 93 (58.9%) participants were diagnosed with cognitive impairment at the follow-up. Correlation analyses showed a high correlation between the NIHSS and the physical performance variable BBS (r = 0.75) and between the inter-related variables BBS, maximum walking speed, TUG and Fig8 (r > 0.60). Table 2 shows the results of the logistic regression analyses, unadjusted and adjusted for significant variables except intercorrelating variables.

A full model containing all the predictors was statistically significant (χ^2 = 44.00, p < 0.01). As shown in table 2, two independent variables made a unique statistically significant contribution in the adjusted analyses (Fig8, BBS). The strongest predictor of cognitive impairment was Fig8 (odds ratio 1.06), indicating that for every additional misstep, participants were 1.06 times likely to have cognitive impairment at the follow-up.

Discussion

The present study shows that balance in the acute phase of stroke predicted cognitive impairment 1 year after stroke. Participants that performed well on Fig8 and BBS had a significantly lower risk of cognitive impairment 1 year after stroke. The results were consistent when adjusted for stroke-related factors such as neurological impairment and sociodemographic variables. Even though walking speed has been suggested as a vital sign [44] and as



 $^{^{1}}$ Small vessel disease or other pathologies. 2 Lacunar syndromes or other syndromes.



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Table 2. Results of unadjusted and adjusted logistic regression of the performance measurements and covariates predicting cognitive impairment after stroke (n = 158)

Variable, baseline	Unadjusted		Adjusted ¹	Adjusted ¹		
	OR (95% CI)	р	OR (95% CI)	р		
Age (years)	1.06 (1.03-1.09)	< 0.001	1.01 (0.96-1.01)	0.670		
Gender (male = 0)	2.12 (1.11-4.05)	0.024	1.22(0.44-3.40)	0.707		
Education (>9 years = 0)	3.34 (0.135-0.66)	0.009	3.35 (0.93-12.11)	0.065		
BMI $(<25 = 0)$	0.94(0.49-1.77)	0.839				
Living arrangements (living alone = 0)	0.30 (0.14-0.66)	0.003	0.86(0.27-2.78)	0.802		
Smoking (yes = 0)	0.94(0.49-1.77)	0.092	0.75(0.21-2.68)	0.653		
Hypertension (yes = 0)	1.61 (0.85-3.07)	0.148	1.68 (0.61-4.60)	0.313		
TOAST (small vessel = 0)	1.58(0.78-3.21)	0.201				
OCSP (lacunar syndromes = 0)	0.94(0.45-1.98)	0.869				
Lesion (right = 0)	1.68 (0.89-3.17)	0.112	2.40 (0.89-6.48)	0.085		
ApoE4 (not having = 0)	0.34(0.03-3.80)	0.379				
NIHSS (points)	1.23 (1.09 – 1.39)	0.001	$0.50(0.82-1.51)^2$	0.503		
TUG (s)	1.05 (1.00-1.11)	0.065	$1.01(0.94-1.09)^2$	0.730		
MWS (s)	1.09 (0.99-1.19)	0.083	$1.05(0.97-1.14)^2$	0.251		
Fig8 (missteps)	1.06 (1.03-1.10)	< 0.001	$1.06(1.02-1.10)^2$	0.001		
BBS (points)	0.94 (0.91-0.98)	0.001	$0.95 (0.92 - 0.98)^2$	0.004		

OR = Odds ratio; CI = confidence interval; BMI = body mass index; lesion = affected side of the brain; MWS = maximum walking speed.

a predictor of cognition in older people [13], maximum walking speed did not have predicting ability in this population assessed during the 1st week after stroke. TUG, the other measure of mobility, did not have the ability to predict cognitive impairment.

Executive function is associated with balance in older adults after mild stroke [45] and with falls among older people [46], which might correspond well with our results. Performance of balance such as in Fig8 and BBS involves executive functioning often linked to vascular disease and partly regulated by the prefrontal areas of the frontal lobe [47]. It has been suggested that executive functions as opposed to global cognition or memory, are important for gait and balance [48, 49]. Executive function is regarded as higher-level cognitive function, implying a control function of other cognitive functions [50]. To achieve effective goal-directed behaviour, it is not only important to be able to initiate and plan actions, but also to be flexible, to be able to shift plan and to inhibit irrelevant information or responses during action. The inherent diversity of abilities involved in executive function is naturally difficult to grasp in single assessments and entails a lack of gold standard measure. Possibly, the individual's cognitive flexibility also described as a set-shifting ability [47] is challenged when performing Fig8 and BBS.

The data did not show an association between lesions of the right side of the brain and cognitive impairment after stroke (p = 0.08). Persons having involvement of the right side of the brain may experience an impact on visuospatial ability affecting the ability to perform the balance measures included in the study. Fig8 requires a brain function in the visuospatial networks, sensorimotor area and cerebellar functions, and it has been suggested earlier that a decrease in visuospatial performance is an indicator of cognitive decline [51]. Recent studies have assessed visuospatial memory in relation to cognitive impairment and found that such tests were able to differentiate subjects with MCI from healthy elderly subjects, but there is a

 $^{^{1}}$ Adjusted for significant variables with p < 0.2. 2 Variables correlating >0.6 were excluded from the analyses.



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lack of results on the potential predictive value [52]. The fact that Fig8 and BBS are associated with cognitive impairment 1 year after stroke suggests that these balance measures might be some of the earliest predictors of cognitive impairment. Fig8 and BBS are easily applicable low-cost tools that should be studied further to reveal their predictive ability.

Most other comparable studies include subjects who already are diagnosed with MCI or dementia, and there is a lack of studies on the predictive value of balance and mobility in stroke. Predictors of cognitive impairment after stroke are requested for better understanding of the mechanisms involved, prevention, prognosis and quality of life. In the general population, a high level of physical activity increases physical performance such as gait, mobility and balance [53]. In addition, there is a positive relationship between physical performance and cognition such as cognitive flexibility or global cognition in healthy elderly people [54]. Cognitive impairment is a risk factor for decline in physical function [55] and increased aerobic endurance may increase hippocampal volume and improve cognitive capacity in both older people with MCI and people without known cognitive impairment [36, 56-58]. Many forms of dementia will demonstrate motor impairment as the disease progresses due to further damage to brain tissue. Whether this applies to people who have suffered a stroke is unknown. However, balance assessed by the one-leg stance test has been shown to be an independent predictor of progression to further cognitive impairment in the early phase of dementia [59]. This is in line with the study by Pettersson et al. [12] showing that persons with mild Alzheimer's disease had a lower score on BBS and significantly more missteps in Fig8, and needed more time on TUG than healthy controls. When comparing the test results of our population with this specific study, our population in the acute phase of stroke showed similar results on TUG with a mean value of 11 s, but worse mean scores on BBS (44.5 vs. 53 points) and Fig8 (18 vs. 13 missteps).

In the present study, the measures of maximum walking speed and TUG did not have the ability to predict cognitive impairment. Still, both maximal walking speed (p = 0.08) and TUG (p = 0.07) showed borderline significant results in the unadjusted analyses. Since this is the first study seeking to predict cognitive impairment after stroke by assessing balance and mobility in the acute phase of stroke and TIA, we are not able to compare our participants with similar populations. Our findings indicate that in the acute phase of stroke, measures including gait such as TUG and maximum walking speed are not sensitive predictors of cognitive impairment. Further knowledge of the associations between physical performance and cognition will be a strength when implementing individualized rehabilitation plans for persons with a first-ever stroke or TIA in the early rehabilitation setting. Fig8 has earlier been described as having the ability to distinguish those with the best motor functioning in the acute phase of stroke [60] regarding balance performance. These persons without severe disabilities in acute stroke will not routinely receive specific supervision after discharge, which underlines the importance of finding in this group those who are vulnerable to developing cognitive impairment. Those at high risk and their families will benefit from customized information and guidance on amendable factors [18]. The best-established protective factors include physical activity, exercise and education [61, 62]. Further studies seeking greater knowledge of targeted treatment to individuals at high risk of developing cognitive impairment would be a strength.

This study has some limitations; the study includes a quite healthy population with first-ever stroke or TIA, and the results may not necessarily be transferable to other populations with more severely affected participants. Furthermore, patients suffering major strokes who were not able to complete the follow-up were excluded due to the longitudinal design. Our study includes maximal walking speed, which is considered to be a valid and reliable measure but, still it would have been a strength if we had included self-selected walking speed, which has been preferred in recent reviews [63]. The strengths of the study include the thoroughness





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of the diagnostics of cognitive impairment and the use of standardized performance-based measures at baseline, in addition to the information on ApoE4 and the affected side of the brain.

Conclusion

The performance on Fig8 and BBS in the acute phase of first-ever stroke or TIA was a predictor of cognitive impairment 1 year after stroke. These performance-based tests require brain function in the visuospatial networks, sensorimotor areas and cerebellar functions. Still, the mechanisms explaining this association between balance and cognitive functioning are not fully understood. Several issues remain to be further explicated, and further research is needed to reproduce these findings in other samples of patients with stroke. In addition, an interventional study of the potential effect of balance training on cognitive functioning would be useful for future patients.

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