## **ORIGINAL REPORT**

# FACTORS PREDICTING THE TOTAL MEDICAL COSTS ASSOCIATED WITH FIRST-EVER ISCHAEMIC STROKE PATIENTS TRANSFERRED TO THE REHABILITATION WARD

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*Objective:* To investigate the predictors of total medical costs for first-ever ischaemic stroke patients transferred to the rehabilitation ward from the acute ward.

*Patients:* A total of 311 first-ever ischaemic stroke patients (mean age 68.9 (standard deviation (SD) 12.2) years).

*Methods:* Data, including common complications and medical events, from July 2002 to June 2012 were collected retrospectively from a regional hospital in Taiwan in order to study the potential predictors for medical costs. Significant variables from univariate analysis were included in a stepwise multivariate linear regression analysis.

*Results:* The mean total medical cost per patient was USD 4,606.80 (SD 2,926.10). The significant predictors for cost were days of total stay (coefficient: 70.3; 95% confidence interval (CI)=56.4–84.3), impaired consciousness (coefficient: 1,031.3; 95% CI=490.8–1,571.8), hypoalbuminaemia in the acute ward (coefficient: 2,045.1; 95% CI=1,054.6–3,035.7), fever (coefficient: 927.0; 95% CI=193.3–1,660.7), hypokalaemia (coefficient: 2,698.4; 95% CI=660.5–4,736.4), and hyponatraemia (coefficient: 1,123.3; 95% CI=72.2–2,174.5) in the rehabilitation ward ( $R^2$ =0.416).

*Conclusion:* These findings can help clinicians to identify risk factors for total medical costs in these patients and reduce costs by minimizing some complications (hypoalbuminaemia, fever, hypokalaemia, and hyponatraemia).

Key words: medical cost; ischaemic stroke; rehabilitation.

J Rehabil Med 2015; 47: 120-125

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Accepted Jun 23, 2014; Epub ahead of print Sep 29, 2014

#### INTRODUCTION

Stroke is a major health problem worldwide. Its consequences include decreased physical activities (1) and social participation (2). The imperative need for effective preventive therapy,

early critical care, and rehabilitation after an acute stroke episode often result in high levels of medical expenditure (3). As the costs for the care of acute stroke patients rise (4, 5), identifying the factors that could significantly predict the overall medical costs of treating acute stroke has become increasingly urgent. The predictors for the cost of acute stroke have been discussed in several reports in the past (6–9). Total hospital costs for acute stroke patients were then correlated mainly with length of stay (6, 8–10), stroke severity (6, 8, 9), male gender (9), female gender (4), age  $\geq 65$  years (7), atrial fibrillation (7, 9), dysphagia (11), and experienced  $\geq 1$ in-hospital complications (6).

Inpatient rehabilitation at the acute stage has been proven to improve independence after treatment (12), to decrease disability at discharge (13), and to decrease long-term costs (14) for stroke patients. Acute stroke patients using inpatient rehabilitation services include those receiving rehabilitation programmes in the acute ward (and eventually discharged from the acute ward), and those transferred to the rehabilitation ward to continue a rehabilitation programme for a prolonged period of time. Although only 34% of patients with acute stroke in Taiwan have utilized in-patient rehabilitation services (15), Chang et al. (16) used cost-effectiveness analysis for stroke management and found that the neurology/rehabilitation wards formed the optimal care model for the management of first-ever ischaemic stroke patients. In Taiwan, the Bureau of National Health Insurance (BNHI) manages a single-payer National Health Insurance (NHI) programme. However, higher-cost NHI claims for single-time hospitalization were reviewed in detail by BNHI to determine whether they were appropriate, and some claims (which BNHI might not have paid for) were sent back to the medical institution if BNHI considered them inappropriate. In case of higher medical costs for acute stroke patients transferred to the rehabilitation ward, the manager of the institution, or even the clinicians of these patients, may also face the problem of claim rejection by NHI. Previous studies discussing the factors affecting medical costs of acute stroke have not focused on the patients transferred to the rehabilitation ward from the acute ward. Considering that ischaemic

stroke patients were the majority of stroke cases, we aimed to design a study that could provide clinicians with information to identify risk factors for medical costs and predict medical costs for first-ever ischaemic stroke patients transferred to the rehabilitation ward from the acute ward. The objective of this study was to retrospectively investigate the predictors of total medical costs for these patients.

Stroke complications frequently occur during a period of stay in the acute ward (17) and during transfer to the rehabilitation ward (18) in the acute stage. As mentioned above, experienc $ing \ge 1$  in-hospital complications (6) is one of the main factors known to increase the costs associated with treating acute ischaemic stroke patients. A previous study showed that the costs of hospitalization for patients with acute stroke with pneumonia were significantly higher than for those without pneumonia (19). On the other hand, the effects of various other common complications or medical events on the total medical costs of acute stroke patients remain unclear. For stroke patients admitted to the rehabilitation ward, medical complications are known to significantly correlate with length of stay, which is also associated with medical costs (20). We hypothesized that particular complications or medical events significantly influence the total medical costs, and these were also selected as potential factors for analysis.

## MATERIAL AND METHODS

#### Enrolment of patients

The electronic medical records of patients admitted to the rehabilitation department at a regional hospital in south-central Taiwan between July 2002 and June 2012 were retrospectively reviewed. The patients considered for this study met the following inclusion criteria: (*i*) being diagnosed with acute first-ever ischaemic stroke according to the World Health Organization's (WHO) criteria definition of stroke (21); (*ii*) being transferred to the rehabilitation ward from the acute ward during the first-time stroke hospitalization; (*iii*) being eventually discharged from the rehabilitation ward after transfer to the rehabilitation ward. The study protocol (number: 100-3662B) was approved by the Institutional Review Board for Human Studies of the regional hospital.

#### Demographic and expense data

The demographic data, including gender (4, 9) and age (7), were analysed. Medical costs incurred during hospitalization generally included diagnosis fees, ward fees, tube feeding fees, laboratory fees, X-ray fees, therapeutic procedure fees, surgical fees, rehabilitation fees, haemodialysis fees, blood/plasma fees, anaesthesia fees, special materials fees, drugs fees, dispensing service fees, psychiatric treatment fees, and injection service fees. In the present study, the total medical costs were defined as the sum of both the NHI claims and co-payment for first-time stroke hospitalization (both in the acute ward and the rehabilitation ward), and this information for each patient was obtained from the hospital's management information system. The NHI claims are the costs that the hospital asks the BNHI to pay; the co-payment is the costs that the patient should pay. The costs were expressed as USD. The New Taiwan Dollar to USD exchange rate was calculated according to the exchange rate on the day (2 December 2013) that the statistical analysis was performed.

#### Medical data and rehabilitation information

In terms of the medical factors potentially affecting the costs associated with treatment, the medical history of the patient was considered, including hypertension (7), diabetes mellitus (DM) (7), hyperlipidaemia (7), atrial fibrillation (7, 9), sites of brain lesion (7) (left, right, or bilateral), stroke-induced impaired consciousness (6, 8, 9), stroke-induced dysphagia (11), and length of total stay (days) (6, 8–10) (including days in both the acute and rehabilitation wards). Hypertension was defined as the hypertension diagnosed by a clinician prior to the stroke, or systemic blood pressure of >160 mmHg and/or diastolic blood pressure of >95 mmHg on 2 separate occasions. DM was defined as fasting plasma glucose of  $\geq$  126 mg/dl or random plasma glucose of  $\geq 200$  mg/dl. Hyperlipidaemia was defined as a fasting cholesterol level of  $\geq$  200 mg/dl or a fasting triglyceride level of  $\geq$  150 mg/dl. Atrial fibrillation was defined as a continuous or paroxysmal arrhythmia and shown as the absence of P waves, with disorganized electrical activity in their place, and irregular R-R intervals due to irregular conduction of impulses to the ventricles on electrocardiogram. Stroke-induced impaired consciousness was defined as a Glasgow Coma Scale (GCS) score <15 after stroke during the entire course of hospitalization. Stroke-induced dysphagia was defined as difficulty in swallowing food or liquids, and was diagnosed after examinations or bedside tests were performed by a physiatrist.

Frequently occurring complications or major medical events during the acute stroke phase (18), including upper gastrointestinal bleeding (UGIB), fever, depression, seizure, shoulder pain, hyponatraemia, hypokalaemia, and hypoalbuminaemia, were also recorded. Complications or major medical events in the acute and rehabilitation wards were considered independent events, because patients with complications or experiencing medical events in the acute ward were treated prior to transfer to the rehabilitation ward. UGIB was defined as either episodes of bloody nasogastric aspirate or ulcers, or erosions or bleeding source proven by oesophagogastroduodenoscopy. Fever was defined as any episode of body temperature >38°C. Depression was diagnosed by a psychiatrist when the symptoms met the fourth edition of Diagnostic and Statistical Manual of Mental Disorders criteria for depression. Seizure was defined as any episode of partial or generalized convulsion and was diagnosed by a neurologist. Shoulder pain was defined as any cause, such as capsulitis, shoulder subluxation, impingement syndrome, rotator cuff injury, and shoulder-hand syndrome, which may induce shoulder pain. Hyponatraemia was defined as any record of serum sodium levels <134 mEq/l, hypokalaemia as any record of serum potassium levels < 3.6 mEq/l, and hypoalbuminaemia as any record of serum albumin levels < 3.5 g/dl.

The rehabilitation programme offered services provided by both physical and occupational therapists. Speech therapists were also available if required. Each of the daily physical and occupational therapy sessions usually lasted more than 50 min (approximately 1 h) during weekdays. Speech therapy was carried out twice a week, and the therapy session was approximately 1 h each time.

#### Statistical analysis

SPSS 12.0 for Windows was used for statistical analyses. Pearson's correlation was used to evaluate the correlations between total medical costs and the continuous variables. A *t*-test was performed to evaluate the differences in total medical costs for patients with and without categorical variables. A *p*-value of < 0.05 was considered statistically significant. Significant independent variables were selected using univariate analysis, and these were included in a stepwise multivariate linear regression analysis in order to identify the most important risk factors for total medical costs. The collinearity of independent variables was also checked.

#### RESULTS

Of the 719 patients analysed, 568 were first-ever stroke patients transferred to the rehabilitation ward from the acute ward and eventually discharged from the rehabilitation ward. After

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Table I. Clinical characteristics of all patients

	Patients
Total, <i>n</i> (%)	311 (100)
Male, <i>n</i> (%)	137 (44.1)
Age, years, mean (SD)	68.9 (12.2)
Acute ward stay, days, mean (SD)	23.3 (11.9)
Rehabilitation ward stay, days, mean (SD)	25.3 (12.0)
Total stay, days, mean (SD)	48.6 (19.2)
Total medical cost, USD, mean (SD)	4,606.8 (2,926.1)
Impaired consciousness, $n$ (%)	118 (37.9)
Dysphagia, n (%)	205 (65.9)
Hypertension, <i>n</i> (%)	240 (77.2)
Diabetes mellitus, $n$ (%)	144 (46.3)
Hyperlipidaemia, n (%)	120 (38.6)
Atrial fibrillation, <i>n</i> (%)	47 (15.1)
Brain lesion, <i>n</i> (%)	
Left	151 (48.6)
Right	152 (48.9)
Bilateral	8 (2.5)

SD: standard deviation.

excluding 257 patients with first-ever haemorrhagic stroke, a total of 311 first-ever ischaemic stroke patients were enrolled in the study, as shown in Table I.

Table II shows the results of the correlations between the total cost and the continuous variables (age and days of total stay). Twelve variables (total number of days of stay, impaired consciousness, dysphagia, UGIB in the acute ward, fever in the acute ward, hyponatraemia in the acute ward, hypoalbuminaemia in the acute ward, UGIB in the rehabilitation ward, fever in the rehabilitation ward, fever in the rehabilitation ward, in the rehabilitation ward) identified from univariate analysis were used for multivariate regression analysis. Six of these (total number of days of stay, impaired consciousness, hypoalbuminaemia in the rehabilitation ward, fever in the rehabilitation ward, hypokalaemia in the rehabilitation ward, fever in the rehabilitation ward, hypokalaemia in the rehabilitation ward, fever in the rehabilitation ward, hypokalaemia in the rehabilitation ward, and hyponatraemia in the rehabilitation ward, hyponatraemia hyponatraemia in the rehabilitation ward, hyponatraemia in the rehabilitation ward, hyponatra

#### DISCUSSION

Six predictors of the total medical costs for the patients in our study were identified and, except for impaired consciousness, they are all adjustable, treatable, or correctable. The total number of days of stay can significantly predict the total medical costs. Length of stay is also a solid factor influencing the total hospital

Table II. Potential factors for total medical cost by univariate analysis

	Cost, USD/r				
Predictive factors	Mean (SD)	<i>p</i> -value			
Demographic data					
Male $(n=137)$	4,386.7 (2,807.2)	0.240			
Age	-0.015	0.790			
Total stay	0.556	< 0.001			
Medical history					
Impaired consciousness $(n=118)$	5,743.6 (3,438.7)	< 0.001			
Dysphagia ( $n=205$ )	5,134.5 (3,064.6)	< 0.001			
Hypertension $(n=240)$	4,683.9 (3,034.1)	0.394			
Diabetes mellitus $(n=144)$	4,473.3 (2,991.7)	0.456			
Hyperlipidaemia $(n=120)$	4,679.4 (2,872.8)	0.730			
Atrial fibrillation $(n=47)$	4,847.7 (2,338.8)	0.541			
Brain lesion					
Left brain $(n=151)$	4,564.2 (2,849.9)	0.803			
Right brain $(n=152)$	4,602.3 (3,018.6)	0.979			
Bilateral brain $(n=8)$	5,499.4 (2,762.2)	0.383			
Complications or medical events occu	urring in the acute ward				
UGIB $(n=35)$	6,350.1 (3,186.0)	< 0.001			
Fever $(n=98)$	5,943.9 (3,371.2)	< 0.001			
Depression $(n=47)$	4,410.2 (2,199.1)	0.618			
Seizure $(n=8)$	6,138.0 (3,580.4)	0.134			
Shoulder pain $(n=14)$	5,102.8 (2,250.5)	0.517			
Hyponatraemia $(n=25)$	6,278.0 (4,482.1)	0.003			
Hypokalaemia $(n=67)$	5,402.8 (3,553.2)	0.012			
Hypoalbuminaemia $(n=23)$	7,294.6 (4,502.5)	< 0.001			
Complications or medical events occurring in the rehabilitation ward					
UGIB $(n=50)$	5,986.3 (3,656.2)	< 0.001			
Fever $(n=46)$	6,100.0 (3,008.6)	< 0.001			
Depression $(n=24)$	5,190.0 (2,251.0)	0.310			
Seizure $(n=3)$	6,743.8 (5,653.2)	0.204			
Shoulder pain $(n=81)$	4,967.7 (2,561.8)	0.197			
Hyponatraemia $(n=20)$	6,850.6 (2,863.3)	< 0.001			
Hypokalaemia $(n=5)$	8,059.5 (6,818.2)	0.008			
Hypoalbuminaemia $(n=7)$	6,584.2 (3,207.3)	0.070			

r: Pearson correlation coefficient; SD: standard deviation; UGIB: upper gastrointestinal bleeding.

cost for acute stroke patients who were not transferred to the rehabilitation ward (6, 8–10), and for those post-stroke patients admitted to hospitals to receive rehabilitation treatment (20, 22). The mean length of stay varies from 6 (9) to 33 (8) days in the acute ward. For acute stroke patients admitted to the rehabilitation ward, the mean length of stay ranged from 24 (23) to 34.7 (24) days. In our study, the mean length of stay was 23.3 days in the acute ward and 25.3 days in the rehabilitation ward; in both cases, the length of stay was within the previously described range. Using the total number of days of stay as a predictor for total medical costs appears to be sort of a circular reasoning, as

Table III. Results of multivariate linear regression analysis for predictors related to the total medical cost ( $R^2 = 0.416$ )

Dependent variable	Independent variable	Coefficient (B)	95% CI	<i>p</i> -value		
Total cost (USD)	Total stay, days	70.3	56.4-84.3	< 0.001		
	Impaired consciousness	1,031.3	490.8-1571.8	< 0.001		
	Hypoalbuminaemia in the acute ward	2,045.1	1,054.6-3035.7	< 0.001		
	Fever in the rehabilitation ward	927.0	193.3-1660.7	0.013		
	Hypokalaemia in the rehabilitation ward	2,698.4	660.5-4736.4	0.010		
	Hyponatraemia in the rehabilitation ward	1,123.3	72.2-2174.5	0.036		

CI: confidence interval.

a significant part of the medical costs consists of regular costs incurred during the hospital stay; the total number of days of stay therefore becomes a significant predictor.

Under NHI, clinicians need to set a reasonable hospital stay for inpatients because the length of stay cannot be extended indefinitely without reason. However, unpredictable causes might prolong the total hospital stay, and hence the cost associated with it (e.g. waiting in the acute ward for transfer to an available rehabilitation bed, or waiting for a transfer to a destination after discharge (nursing home, home, or another medical institution)). A previous study showed that approximately 35.6% of stroke patients admitted to the rehabilitation ward experienced delays during discharge (25). The main reasons for these delays were caregiver-related factors or organizational factors (e.g. waiting for nursing home transfer) (25). Constructing an efficient discharge planning programme for patients may solve the problems related to discharge delays and would help decrease the associated expense.

The rehabilitation fees could not be separated from the total costs in our study. According to a study by Tang et al. (10) in Taiwan, rehabilitation fees spent on physical therapy, occupational therapy, and speech therapy in the rehabilitation ward at a regional hospital was approximately USD 32.5 per day, accounting for 36.7% of the total daily cost (USD 86.4) (10). In the Kuptniratsaikul's multicentre study (23), the cost related to rehabilitation procedures was estimated to be 33.2% of the total cost for stroke patients admitted to the rehabilitation ward. Previous articles illustrated that the rehabilitation fees charged in the acute ward for stroke inpatients represented only approximately 7% (8, 9) of the total acute hospital costs. It seemed that the percentage of rehabilitation fees were higher in the rehabilitation ward (compared with total costs incurred in the rehabilitation ward) than those in the acute ward (compared with total costs incurred in the acute ward). In addition to more intensive care, medical procedures or examinations may be performed in the acute ward, which should be expected to significantly increase the total costs. We believe that future studies should focus on individually identifying predictors for medical costs and rehabilitation fees in acute stroke patients who received rehabilitation in the acute ward and the rehabilitation ward.

According to Wei et al. (6), severe GCS score (defined as 3-8) on admission was a predictor for the cost in acute stroke patients. Following Wei et al., GCS was used as a marker for stroke severity in our study. We provide evidence that the stroke severity with any impaired consciousness, i.e. without the highest GCS score, after stroke could be a predictor for total cost. Diringer et al. (9) used the National Institutes of Health Stroke Scale score (NIHSS) as a measurement for stroke severity and found that total hospital costs for patients with acute severe stroke (NIHSS score > 20) was more than twice that of patients with mild stroke. Yoneda et al. (8) also used HIHSS as a measurement of stroke severity. Total hospital costs were moderately correlated with the initial NIHSS. Because of the retrospective method, only limited GCS data for the measurement of stroke severity were available for the present

study. For future studies, we aim to use more measurements, including a functional survey on admission to the acute ward, in a prospective manner.

After a thorough literature search, we found little evidence illustrating that fever can be associated with increasing total medical costs in stroke patients. One previous study (26) illustrated that in-hospital infection is a predictor of prolonged hospital stay in acute ischaemic stroke patients. Another study (27) illustrated that a history of pneumonia could predict higher hospital charge per day for patients admitted for inpatient rehabilitation. In our study, fever in the rehabilitation ward increased medical costs for acute stroke patients transferred to the rehabilitation ward. Fever is usually associated with certain infections, such as pneumonia or urinary tract infection, which also occur frequently in acute stroke patients. Antibiotic usage and/or delayed discharge due to infection treatment may thus increase medical costs. There are no references considering hypoalbuminaemia, hyponatraemia, or hypokalaemia as a predictor for total costs for stroke patients. However, hypoalbuminaemia has been associated with increased risks for medical complications in acute stroke patients (28) and stroke patients admitted to the rehabilitation ward (27, 29), although it has not been related to length of stay (27, 29). Post-stroke hyponatraemia (30) and hypokalaemia (31) are associated with poor outcomes (increased chance of death). We believe that hypoalbuminaemia, hyponatraemia, and hypokalaemia are somewhat related to certain medical complications, which might increase costs because of the need for management of complications.

No haemorrhagic stroke inpatients were enrolled in the present study from the beginning. Gioldasis et al. showed that the costs for patients with acute haemorrhagic strokes were significantly higher than those for patients with acute ischaemic strokes in a tertiary hospital (32) in Greece. Wei et al. (6) illustrated that acute intracerebral haemorrhage (ICH) was associated with a 19% greater cost than acute ischaemic stroke in a Level 2 hospital (a hospital with at least 100 inpatient beds providing acute medical care and preventative care services to populations of at least 100,000) in China. In the USA, the mean hospital cost for patients with ischaemic stroke was lower than that for patients with ICH and subarachnoid haemorrhage (33, 34). In Japan, patients with acute ICH had higher costs than those with acute ischaemic stroke (35). In Bottacchi's 3-year longitudinal study in Italy (36), the mean annual cost was significantly higher for patients with hemorrhagic stroke than for those with ischaemic stroke. In a study conducted in Taiwan, Tang et al. (10) included acute and chronic stroke patients (4 hospitals with different levels) admitted to the rehabilitation ward, providing evidence that haemorrhagic strokes had a higher total cost than ischaemic strokes. We considered that in the acute stage, more neurosurgical procedures and intensive care, greater disease severity, or longer length of hospital stay will be required for patients with haemorrhagic stroke than for those with ischaemic stroke.

To our knowledge, this is the first study analysing the predictors of total cost for first-ever ischaemic stroke patients

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transferred to the rehabilitation ward and identifying that some complications or medical events as the predictors. However, the design of the study limited the extent of applications of the results, e.g. the data could not be applied to acute ischaemic stroke patients who received rehabilitation programmes but were not transferred to the rehabilitation ward, and to those who were readmitted to the rehabilitation ward for rehabilitation after discharge from the hospital following acute ischaemic stroke. The main limitations of the current study are that it is hospital-based only, and that we obtained only the total medical costs; no medical expense details were included in the analyses. Further studies should focus on analysing in more detail the costs of various factors using nationwide data for stroke patients who subsequently receive rehabilitation.

In conclusion, length of stay is a robust predictor for the total hospital cost for acute stroke patients transferred to the rehabilitation ward. Hypoalbuminaemia in the acute ward, fever, hypokalaemia, and hyponatraemia in the rehabilitation ward are identified as impact factors on the total hospital cost. These findings may be useful for clinicians to identify the risks of increasing total hospital costs in acute ischaemic stroke patients transferring to the rehabilitation ward. Furthermore, we hope that this information could help clinicians to understand that a lower occurrence of these identified complications might reduce the total medical costs.

### ACKNOWLEDGEMENTS

The Chang Gung Memorial Hospital Research Project Grant financially supported this research under Contract No. CMRPG 6B0241.

The authors declare no conflicts of interest.

### REFERENCES

- 1. Askim T, Bernhardt J, Churilov L, Fredriksen KR, Indredavik B. Changes in physical activity and related functional and disability levels in the first six months after stroke: a longitudinal follow-up study. J Rehabil Med 2013; 45: 423–428.
- Fallahpour M, Jonsson H, Joghataei MT, Nasrabadi AN, Tham K. "I am not living my life": lived experience of participation in everyday occupations after stroke in Tehran. J Rehabil Med 2013; 45: 528–534.
- Demaerschalk BM, Hwang HM, Leung G. US cost burden of ischemic stroke: a systematic literature review. Am J Manag Care 2010; 16: 525–533.
- Smurawska LT, Alexandrov AV, Bladin CF, Norris JW. Cost of acute stroke care in Toronto, Canada. Stroke 1994; 25: 1628–1631.
- 5. Evers SM, Ament AJ, Blaauw G. Economic evaluation in stroke research: a systematic review. Stroke 2000; 31: 1046–1053.
- Wei JW, Heeley EL, Jan S, Huang Y, Huang Q, Wang JG, et al. Variations and determinants of hospital costs for acute stroke in China. PLoS One 2010; 5: e13041.
- Huang YC, Hu CJ, Lee TH, Yang JT, Weng HH, Lin LC, et al. The impact factors on the cost and length of stay among acute ischemic stroke. J Stroke Cerebrovasc Dis 2013; 22: e152–e158.
- Yoneda Y, Uehara T, Yamasaki H, Kita Y, Tabuchi M, Mori E. Hospital-based study of the care and cost of acute ischemic stroke in Japan. Stroke 2003; 34: 718–724.
- 9. Diringer MN, Edwards DF, Mattson DT, Akins PT, Sheedy CW,

Hsu CY, et al. Predictors of acute hospital costs for treatment of ischemic stroke in an academic center. Stroke 1999; 30: 724–728.

- Tang FT, Liu DR, Chen PC, Pai JY. [The study of cost payment system for hospitalized rehabilitation patients.] J Reh Med Assoc ROC 2000; 28: 77–85 (in Chinese).
- Wojner AW, Alexandrov AV. Predictors of tube feeding in acute stroke patients with dysphagia. AACN Clin Issues 2000; 11: 531–540.
- Balaban B, Tok F, Yavuz F, Yasar E, Alaca R. Early rehabilitation outcome in patients with middle cerebral artery stroke. Neurosci Lett 2011; 498: 204–207.
- Matsui H, Hashimoto H, Horiguchi H, Yasunaga H, Matsuda S. An exploration of the association between very early rehabilitation and outcome for the patients with acute ischaemic stroke in Japan: a nationwide retrospective cohort survey. BMC Health Serv Res 2010; 10: 213.
- Mahler MP, Zuger K, Kaspar K, Haefeli A, Jenni W, Leniger T, et al. A cost analysis of the first year after stroke – early triage and inpatient rehabilitation may reduce long term costs. Swiss Med Wkly 2008; 138: 459–465.
- Lee HC, Chang KC, Huang YC, Lan CF, Chen JJ, Wei SH. Inpatient rehabilitation utilization for acute stroke under a universal health insurance system. Am J Manag Care 2010; 16: e67–e74.
- Chang KC, Lee HC, Huang YC, Hung JW, Chiu HE, Chen JJ, et al. Cost-effectiveness analysis of stroke management under a universal health insurance system. J Neurol Sci 2012; 323: 205–215.
- Johnston KC, Li JY, Lyden PD, Hanson SK, Feasby TE, Adams RJ, et al. Medical and neurological complications of ischemic stroke: experience from the RANTTAS trial. RANTTAS Investigators. Stroke 1998; 29: 447–453.
- Chen CM, Hsu HC, Chang CH, Lin CH, Chen KH, Hsieh WC, et al. Age-based prediction of incidence of complications during inpatient stroke rehabilitation: a retrospective longitudinal cohort study. BMC Geriatr 2014; 14: 41.
- Katzan IL, Dawson NV, Thomas CL, Votruba ME, Cebul RD. The cost of pneumonia after acute stroke. Neurology 2007; 68: 1938–1943.
- Saxena SK, Ng TP, Yong D, Fong NP, Gerald K. Total direct cost, length of hospital stay, institutional discharges and their determinants from rehabilitation settings in stroke patients. Acta Neurol Scand 2006; 114: 307–314.
- World Health Organization. WHO STEPS Stroke Manual: The WHO STEPwise approach to stroke surveillance. Geneva: World Health Organization; 2006.
- Saxena SK, Koh GC, Ng TP, Fong NP, Yong D. Determinants of length of stay during post-stroke rehabilitation in community hospitals. Singapore Med J 2007; 48: 400–407.
- Kuptniratsaikul V, Kovindha A, Massakulpan P, Permsirivanich W, Kuptniratsaikul PS. Inpatient rehabilitation services for patients after stroke in Thailand: a multi-centre study. J Rehabil Med 2009; 41: 684–686.
- 24. Lin JH, Hsiao SF, Liu CK, Lin YT. Rehabilitation fees, length of stay and efficiency for hospitalized stroke patients: a preliminary study based on function-related groups. Kaohsiung J Med Sci 2001; 17: 475–483.
- Tan WS, Chong WF, Chua KS, Heng BH, Chan KF. Factors associated with delayed discharges after inpatient stroke rehabilitation in Singapore. Ann Acad Med Singapore 2010; 39: 435–441.
- Spratt N, Wang Y, Levi C, Ng K, Evans M, Fisher J. A prospective study of predictors of prolonged hospital stay and disability after stroke. J Clin Neurosci 2003; 10: 665–669.
- Harvey RL, Roth EJ, Heinemann AW, Lovell LL, McGuire JR, Diaz S. Stroke rehabilitation: clinical predictors of resource utilization. Arch Phys Med Rehabil 1998; 79: 1349–1355.
- Gariballa SE, Parker SG, Taub N, Castleden CM. Influence of nutritional status on clinical outcome after acute stroke. Am J Clin Nutr 1998; 68: 275–281.
- 29. Finestone HM, Greene-Finestone LS, Wilson ES, Teasell RW.

Prolonged length of stay and reduced functional improvement rate in malnourished stroke rehabilitation patients. Arch Phys Med Rehabil 1996; 77: 340–345.

- Rodrigues B, Staff I, Fortunato G, McCullough LD. Hyponatremia in the prognosis of acute ischemic stroke. J Stroke Cerebrovasc Dis 2014; 23: 850–854.
- Gariballa SE, Robinson TG, Fotherby MD. Hypokalemia and potassium excretion in stroke patients. J Am Geriatr Soc 1997; 45: 1454–1458.
- 32. Gioldasis G, Talelli P, Chroni E, Daouli J, Papapetropoulos T, Ellul J. In-hospital direct cost of acute ischemic and hemorrhagic stroke in Greece. Acta Neurol Scand 2008; 118: 268–274.
- 33. Qureshi AI, Suri MF, Nasar A, Kirmani JF, Ezzeddine MA, Divani

AA, et al. Changes in cost and outcome among US patients with stroke hospitalized in 1990 to 1991 and those hospitalized in 2000 to 2001. Stroke 2007; 38: 2180–2184.

- Lee WC, Christensen MC, Joshi AV, Pashos CL. Long-term cost of stroke subtypes among Medicare beneficiaries. Cerebrovasc Dis 2007; 23: 57–65.
- 35. Yoneda Y, Okuda S, Hamada R, Toyota A, Gotoh J, Watanabe M, et al. Hospital cost of ischemic stroke and intracerebral hemorrhage in Japanese stroke centers. Health Policy 2005; 73: 202–211.
- 36. Bottacchi E, Corso G, Tosi P, Morosini MV, De Filippis G, Santoni L, et al. The cost of first-ever stroke in Valle d'Aosta, Italy: link-ing clinical registries and administrative data. BMC Health Serv Res 2012; 12: 372.