Determinants of quality of life in patients with hemorrhagic stroke
A path analysis

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Abstract
Identifying the determinants of health-related quality of life (HRQOL) improved assessment and decision-making in clinical practice. A few studies have focused on the determinants of HRQOL and their interrelationships in patients with hemorrhagic stroke. The aim of this study was to identify the factors contributing to HRQOL and examine their interrelationships.

A total of 202 patients with hemorrhagic stroke who were discharged from the neurological unit participated in this study. Stroke-specific quality of life was used to assess HRQOL. The Hamilton Rating Scale for Anxiety, the Hamilton Rating Scale for Depression, the Scandinavian Stroke Scale and the Barthel Index were collected as potential predictors as well as social-demographic data. A path analysis was used to explore the potential interrelationships between various factors based on the International Classification of Functioning model.

The final model reasonably fitted the data. The activities of daily living, neurological function and anxiety had direct effects on quality of life. Age, comorbidities, hemorrhage type, financial status, anxiety, and neurological function also had indirect influences on quality of life. All these factors explained 82.0% of all variance in quality of life.

HRQOL in patients with stroke can be predicted by anxiety, neurological function, activities of daily living and other personal and environmental factors. These identified predictors and their interrelationships may assist clinical professions focusing their assessments and developing strategies for modifiable factors to improve HRQOL.

Abbreviations: AGFI = adjusted goodness-of-fit index, BI = Barthel Index, GFI = goodness-of-fit index, HAMA = Hamilton Rating Scale for Anxiety, HAMD = Hamilton Rating Scale for Depression, HRQOL = health-related quality of life, ICF = International Classification of Functioning, Disability and Health, RMSEA = root mean square error of approximation, SRMR = standardized root mean square residual, SS-QOL = stroke-specific quality of life, SSS = Scandinavian Stroke Scale.

Keywords: determinants, path analysis, quality of life, stroke

1. Introduction
Stroke is the second leading cause of death and is also the main reason for long-term disability worldwide.[1–3] Furthermore, stroke, especially hemorrhagic stroke, is a major health concern in China and is the first leading cause of death and disability.[4] According to a report from The Stroke Control Project Committee of National Health and Family Planning Commission of China (2015), the prevalence rate of hemorrhagic stroke was 125.78/100,000 in urban areas and 159.91/100,000 in rural areas. The morbidity of hemorrhagic stroke in China was nearly 3 times that of the world level (60–80/100,000 vs 24.6/100,000) and increased approximately 8.7% per year.[5,6] The burden of hemorrhagic stroke is expected to worsen in the coming years. Much progress has been made in treatments and rehabilitations for stroke survivors to increase survival rates, and quality of life has been an important outcome of medical care.[7] Health-related quality of life (HRQOL) reflects the impact of the health of an individual on functional and perceived wellness in different life domains.[8] Identifying the predictors for HRQOL can provide deep insights into the treatment and rehabilitation intervention strategies in clinical practice.[9]

Several studies have examined the predictors of HRQOL in patients with stroke. According to a literature review, age,[10–12] sex,[10,11] education level,[9] financial status,[13] physical function,[10,13,14] cognitive function,[12,15] comorbidities,[11,16] anxiety, and depression[17,18] have been reported to be predictors of HRQOL. Despite the large number of studies examining the predictors of HRQOL, little is known about the factors that predict quality of life in hemorrhagic stroke patients. Additionally, most studies used multiple regression analysis to identify predictors of HRQOL, which could not take the potential interrelationships between various variables into account. Thus, path analysis is an alternative method to identify the direct and indirect pathways through which potential interrelationships affect HRQOL.[19] To account for the interrelationships between predictor variables using path analysis, a theoretical framework is needed.

The International Classification of Functioning, Disability and Health (ICF) framework[20] was selected to examine the relationships between the predictor variables of HRQOL. The...
ICF is a universal framework, that covers the spectrum of problems involved in patients with acute and chronic health conditions. The 3 key components of ICF are body function and structure, activity and participation. Loss or deviations from normal body functions and structures are defined as body function or structure impairments; difficulties in daily activities are referred to as activity limitation; problems involving life situations are participation restrictions. Moreover, the ICF framework takes personal and environmental factors into consideration. The multi-perspective domains of the ICF were adopted to describe the health conditions of hemorrhagic stroke (Fig. 1). According to the domains of the ICF framework, the potential predictor variables of HRQOL were divided into clearly defined groups. Thus, a theoretical model can be established based on the hypothesis that the HRQOL of patients with hemorrhagic stroke depends on personal and environmental factors, body function or structure impairments and activity limitations (Fig. 2).

The aim of this study was to identify the predictors of HRQOL in patients with hemorrhagic stroke and explore the direct and indirect relationships between the identified determinants under the framework of ICF.
2. Methods

2.1. Study design and participants

This study was a cross-sectional study performed in the neurosurgery unit of a teaching hospital in Chengdu, China from January 2016 to September 2017. A total of 2437 consecutive patients who were discharged from the ward were screened, and 202 agreed to take part in and complete this study. The inclusion criteria of the stroke survivors were:

1. a diagnosis of first-ever hemorrhagic stroke by CT/MRI;
2. a Glasgow Coma Scale score of 9 or above; and
3. age older than 18 years.

The exclusion criteria were:

1. patients in the acute stroke phase (less than 1 month after onset); and
2. unable to read or understand the questionnaire.

The study was approved by the Ethics Committee of West China Hospital, Sichuan University. Written informed consent was obtained from the participants before the questionnaires were distributed.

2.2. Data collection

2.2.1. Health-related quality of life. The stroke-specific quality of life (SS-QOL) questionnaire is specifically designed for patients with stroke, which is significantly more valid and sensitive compared to traditional instruments.[25,26] The questionnaire consists of 12 domains encompassing 49 items, including the social role, mobility, energy, language, self-care, mood, personality, thinking, upper extremity function, family role, vision, and work/productivity. Each item is scored on a 5-point Likert scale in which 1 means complete agreement and 5 means complete disagreement. The total score ranges from 49 to 245, with higher scores indicating a better quality of life. Wang, Jian-Guo, Jun-Tao[25] translated the SS-QOL questionnaire into Chinese version and investigated the reliability and validity in patients with stroke in China. All 12 domains showed excellent reliability in test-retest, inter-test (kappa coefficient ranged from 0.82–1.00) and internal consistent (Cronbach’s coefficient >0.76).[25]

2.2.2. Personal and environmental factors. Personal factors, including age, gender, education level, and marriage status (Fig. 2), were examined. Characteristics of disease including hemorrhage type, comorbidities, time before treatment and whether to receive surgical treatment were also collected. Environmental factors included social support, insurance, and financial status. Participants were categorized as having adequate social support if they lived with caregivers, such as family members and formal caregivers.

2.2.3. Impairments in body structure or function. The Hamilton Rating Scale for Anxiety (HAMA) and the Hamilton Rating Scale for Depression (HAMD) were used to measure the impairments in mental function of patients.[26,27] The HAMA consists of 14 items measured on a 5-point scale. A score of 0 represents no and 4 represents extremely. The total score was classified as no (<7), potential (7–13), assured (14–29) and severe anxiety (>29). The HAMD contains 24 items, of which 9 items are defined from 1 to 2, 1 item is defined from 0 to 2, and 14 items are defined from 0 to 4. The total score was classified as no (<8), potential (8–19), assured (20–35), and severe depression (>35).

Impairments in neurological function were assessed using the Scandinavian Stroke Scale (SSS).[28] The SSS contains 9 items measured with 1 item scoring 0 to 2, 1 item scoring 0 to 4, 5 items scoring 0 to 6, 1 item scoring 0 to 10 and 1 item scoring 0 to 12. The total score was classified as good (>50), intermediate (30–49) or poor (<29).

2.2.4. Activity limitations. Limitations in performing self-care activities and mobility were examined in relation to HRQOL (Fig. 2). Self-care activities referring to the performance of daily tasks such as dressing and washing were evaluated via the ability of daily living. The Barthel Index (BI) served to measure the activities of daily living of the patients.[29] The scale describes 10 tasks in daily life and is scored according to the time or assistance required by patients. Two items range from 0 to 5, 6 range from 0 to 10 and 2 range from 0 to 15. The total score ranges from 0 to 100, with a higher score representing greater independence.

2.3. Procedure

Participants were screened from the medical records and interviewed over the telephone or face-to-face to obtain informed consent. Then, social-demographic data and clinical characteristics were collected from the medical records. The HRQOL, body function or structure impairment and activity limitation were assessed using SS-QOL, HAMA, HAMD, SSS, and BI scales by a trained research assistant.

2.4. Data analysis

Descriptive statistics of the predictor variables based on the ICF framework (Fig. 2) were presented as the mean, standard deviation, median, interquartile range and percentages as appropriate. Path analysis was used to examine the hypothesized causal relationships between social-demographic factors, mental impairments, neurological function impairments, activity limitations and quality of life. As illustrated in Fig. 2, these relationships were assumed to be unidirectional. Structural equation modeling (SEM) was conducted to assess the fitness of the final model. The $\chi^2$ static was used to assess the magnitude of the discrepancy between the sample and fitted covariance matrices, where the $P > 0.05$ indicated that the model and data were consistent.[30] The root mean square error of approximation (RMSEA), the goodness-of-fit index (GFI), the adjusted goodness-of-fit index (AGFI), the standardized root mean square residual (SRMR), and the Akaike Information Criterion (AIC) was used to evaluate the optimum model, with RMSEA < 0.08, GFI > 0.90, AGFI > 0.90, and SRMR < 0.05 indicating good model fitness. The path analysis was performed using Amos 24 (SPSS Inc., Chicago, IL).

3. Results

3.1. Social-demographic and clinical characteristics

Table 1 shows the social-demographic and clinical characteristics of patients. The participants consisted predominantly of females (53.5%) with a mean age of 54.6 years old (SD = 11.1). Of all participants, 25.6% had a low to moderate education level, and only 44.6% had medical insurance. Subarachnoid hemorrhage was the main cause of hemorrhagic stroke in this sample (57.4%). Most of the participants underwent surgical treatment (69.3%). The participants had moderate activity limitations and neuro-
**Table 1**

| Characteristics          | n (%) | Mean (SD) |
|--------------------------|-------|-----------|
| Age                      | 35.0  | 29.0 (11.0) |
| Gender                   |       |           |
| Female/Age               | 40 (53.5) | 39 (11.9) |
| Male/Age                 | 42 (45.5) | 42 (12.7) |
| Education                |       |           |
| ≤ 6 years                | 24 (31.5) | 24 (11.1) |
| 6–12 years               | 56 (70.5) | 56 (12.7) |
| > 12 years               | 4 (5.0) | 4 (12.7) |
| Married (yes)            | 168 (83.6) | 168 (10.9) |
| Insurance (yes)          | 90 (44.6) | 90 (11.9) |
| Time before treatment    |       |           |
| [Median (IQR) hour]      | 5 (2–10) | 5 (2–10) |

**Hemorrhage type**
- Subarachnoid hemorrhage: 116 (74.4)
- Intracranial hemorrhage: 69 (46.4)
- Others: 13 (6.5)

**Comorbidities (yes)**: 25 (15.6)
- Depression: 26 (5.6)
- Diabetes: 30 (18.2)
- Heart disease: 20 (12.1)
- Cough: 20 (12.1)
- Asthma: 13 (8.5)
- Others: 5 (3.1)

**Surgical treatment (yes)**: 140 (69.3)
- Craniotomy: 136 (69.3)
- Burr hole: 4 (2.0)
- Others: 10 (5.3)

**Financial status**
- Poor: 9 (5.0)
- General: 15 (8.5)
- Good: 26 (14.5)

**BI**
- 54.4 (29.5)

**HAMD**
- 16.5 (11.2)

**HAMA**
- 15.9 (11.2)

**SSS**
- 46.8 (10.9)

**SS-QOL**
- 164.2 (51.4)

Bi = Barthel Index, HAMD = Hamilton Rating Scale for Anxiety, HAMA = Hamilton Rating Scale for Depression, SS-QOL = Stroke-specific Quality of Life, SSS = Scandinavian Stroke Scale.

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logical function impairments. The participants experienced assured anxiety and potentially borne depression.

### 3.2. Relationships between SS-QOL, personal and environmental factors

A path analysis model based on the ICF framework (Fig. 1) was used to examine the relationships between personal and environmental factors, mental function impairments, neurological impairments, and activity limitations with quality of life. This model explained 82% of all variance in SS-QOL and had a standardized coefficient of determination (R²) of 0.82. The model showed that SS-QOL was significantly associated with factors such as age, hemorrhage type, comorbidities, and financial status. The neurological function impairments were also indirectly related to SS-QOL through HAMA (β = 0.18, P < .001) and socially mediated through the activity limitation (β = 0.57, P < .001). Additionally, the depression had no contribution to SS-QOL through either direct or indirect pathway. Table 2 shows the standardized direct, indirect and total effects of personal and environmental factors, function impairments and activity limitation on SS-QOL.

### 3.3. Relationship between SS-QOL, function impairments and activity limitations

The relationships between SS-QOL function impairments and activity limitations are displayed in Fig. 3b. Activity limitation (β) directly contributed to SS-QOL most strongly (β = 0.57, P < .001). Anxiety (HAMA) and neurological function impairments (SSS) were also significant contributing factors. Anxiety contributed to SS-QOL directly via a relationship with SS-QOL (β = 0.11, P = .004) and indirectly through activity limitation (β = –0.37, P < .001). Similarly, the contribution of neurological function impairments was established via a direct relationship with SS-QOL (β = 0.31, P < .001) and indirectly through activity limitation (β = 0.57, P < .001). However, depression had no contribution to SS-QOL through either direct or indirect pathway. Table 2 shows the standardized direct, indirect and total effects of personal and environmental factors, function impairments and activity limitation on SS-QOL.

### 3.4. The model of SS-QOL in patients with hemorrhagic stroke

The statistics to assess the fit of the model are shown in Table 3. The χ² test was non-significant (P = .246) and the RMSEA was 0.032 (less than 0.08), indicating that the model and data were consistent. The discrepancy divided by degrees of freedom was 1.248 (less than 2), suggesting a good fit of this model. The GFI and AGFI were 0.973 and 0.941 (greater than 0.90). The SRMR was 0.048 (less than 0.05). All the statistics showed the good fitness of this model.

### 4. Discussion

The model examined in this study reasonably fitted with the data, indicating that it was effective to identify the key predictors of HRQOL in patients with hemorrhagic stroke using path analysis. Complex interrelationships between personal and environmental factors, body function or structure impairments and activity limitations were found and explained 82% of all variance in the HRQOL of patients with hemorrhagic stroke.

#### 4.1. Direct effects

The model coefficients showed that the activity limitation, mental and neurological impairments had direct effects on HRQOL. The activity limitation had the largest effect and explained most of the variance in HRQOL.

Similar to previous studies,[13,14,31] the activity limitation in this study significantly predicted the HRQOL of patients with hemorrhagic stroke. The stroke survivors always lived with functional impairments, such as motor disorder, sensory disorder or language disorder.[31–33] The decrease of function level decided the lower ability to cope with self-care activities of daily living, which played an important role in HRQOL. In fact, the quality of life was the individual’s self-perceived position within the social contexts.[34] The decrease of activities of daily living indicates the care needs from others, which may change the self-perceived position of patients, leading to a decrease in quality of life. This result indicated that professions should provide more interventions to improve the daily living activities of patients. Functional rehabilitations for patients were confirmed to be effective in improving the HRQOL.[12] Additionally, using supportive tools appropriately for the daily activities of stroke survivors may also be useful to obtain better quality of life.[35]

The neurological function impairments were also a strong predictor of HRQOL in this study. This result confirmed the significant relationship between neurological function and HRQOL in previous studies.[15,36] Cognitive impairment was the most common outcome caused by neurological function impairments. The relationship between HRQOL and cognitive function was well-known.[17] Carod-Artal, Trizotto, Coral, Moreira[36] found that better cognitive function predicted better communication and memory domains of HRQOL in patients...
with stroke. Cognitive function was also recognized as a predictor of the physical function domain of HRQOL. The motor disorder also had side effects on the HRQOL of patients. Typically, the upper limb and lower extremity motor function recovery helped patients perceive HRQOL. This result indicated that neurological function recovery was the key component for patients’ rehabilitation after discharge, including sensory, memory, motion, cognition, and communication exercise.

It was not surprising to find that anxiety was a determinant of HRQOL. In fact, anxiety was a common psychological problem for stroke survivors with a prevalence of 22% to 25% in the first 6 months after stroke. The health consequences, evolution, and fear of aggravation of disease, the change in financial status and care needs brought serious anxiety to patients. High anxiety easily decreased energy levels and led to the social isolation of patients, thereby influencing HRQOL. This finding revealed that mental health of patients with stroke was important. Detailed disease and cure information, available health care services after discharge and early interventions for anxiety symptoms may reduce anxiety. However, there was no evidence that depression had any impact on HRQOL in this study. This finding was inconsistent with previous studies, most of which suggested that depression or both anxiety and depression were related to HRQOL in patients with stroke. Although patients with stroke had a higher prevalence of depression (24% to 30%) than anxiety, depression showed less stability and persistence than anxiety, especially at the early stage after stroke. This may explain why depression did not predict the HRQOL in this study. More
**4.2. Indirect effects**

Although all personal and environmental factors had no direct effects on HRQOL, age, hemorrhage type, and financial status had indirect effects on HRQOL via neurological impairments. This result may imply that patients with older age, subarachnoid hemorrhage, and poor financial status had lower HRQOL through the level of neurological function. Additionally, comorbidities had effects on HRQOL through anxiety and neurological impairments, confirming the effects of comorbidities on HRQOL in previous studies.\(^{[41,36]}\)

Apart from the direct effect, anxiety had an indirect influence on HRQOL through activity limitations. Anxiety certainly influences the physical rehabilitation of patients by intensifying fear, nervousness and muscle tension.\(^{[43]}\) This effect may result in a decrease of ability in daily living. As noted in a previous study, patients with post-stroke anxiety suffered more daily life activity limitations than social function limitations.\(^{[43]}\) This may explain how anxiety influences HRQOL through activity limitation. Therefore, clinical professions should take into account the influence of anxiety on physical rehabilitation and HRQOL. Neurological impairments also had an indirect effect on HRQOL through activity limitation. The motor disorder, vision disorder, cognitive impairments and some other function impairments caused by neurological impairments made daily activities, such as bathing, washing and dressing much harder for patients. This result indicated that the rehabilitation of neurological function may improve HRQOL by increasing the activities of daily living of patients with stroke.

**5. Limitations**

There are several limitations in this study. First, the convenience sample with a Glasgow Coma Scale score of 9 or above reduces the representativeness of this study. However, it is necessary to exclude patients with Glasgow Coma Scale scores under 9 due to the anxiety and depression assessment. Second, the small sample size regarding the number of parameters that were examined in this study made it hard to draw definitive causal inferences. A larger sample size study is required to provide better insight into how patients with hemorrhagic stroke perceive HRQOL. Finally, the cross-sectional study cannot describe the change in HRQOL over time. Longitudinal studies are needed to examine the change of HRQOL over time and its determinants at different time points.

**6. Conclusion**

This study showed that anxiety, neurological impairments and activities of daily living of patients with hemorrhagic stroke directly contributed to HRQOL. The study also demonstrated a model based on the ICF framework including personal and environmental factors, body function or structure impairments and activity limitations. The determinants of HRQOL and the interrelationships between these factors may help clinical professions to develop specific interventions for some modifiable factors, such as anxiety, to improve the HRQOL of patients with hemorrhagic stroke.

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**Author contributions**

The manuscript has been read and approved by all authors. The authors are alone responsible for the content and writing of the paper. All authors contributed toward data analysis, drafting and revising the paper and agree to be accountable for all aspects of the work.

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