ORIGINAL REPORT

FATIGUE IS ASSOCIATED WITH REDUCED PARTICIPATION AND HEALTH-RELATED QUALITY OF LIFE FIVE YEARS AFTER PERIMESENCEPHALIC SUBARACHNOID HAEMORRHAGE: A MULTICENTRE CROSS-SECTIONAL STUDY

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Objective: To determine whether fatigue is associated with participation and health-related quality of life 5 years after perimesencephalic subarachnoid haemorrhage.

Design: Multicentre cross-sectional study.

Subjects: Forty-six patients with perimesencephalic subarachnoid haemorrhage.

Methods: Fatigue was assessed with the Fatigue Severity Scale, participation (frequency, restrictions, satisfaction) with the Utrecht Scale for Evaluation of Rehabilitation Participation, health-related quality of life with the Stroke-Specific Quality of Life Scale-12, symptoms of depression and anxiety with the Hospital Anxiety and Depression Scale, and coping with the Coping Inventory for Stressful Situations.

Results: A total of 46 patients were included (63% men, mean age 50.4 ± 9.4 years), with a mean time of 4.7 ± 1.6 years after perimesencephalic subarachnoid haemorrhage onset. Fatigued patients (33%) had worse participation (p < 0.01) and health-related quality of life (p < 0.001) than non-fatigued patients, and more often had hypertension, depression, anxiety and emotion-oriented coping (p < 0.05). Fatigue severity was inversely and independently (p < 0.005) associated with participation frequency (B = −3.62), satisfaction (B = −4.54), having restrictions (odds ratio = 2.48, 95% confidence interval 1.079–5.685), and health-related quality of life (B = −0.19), adjusted for depression, anxiety, and/or hypertension.

Conclusion: Five years after perimesencephalic subarachnoid haemorrhage, one-third of patients still reported fatigue, which was associated with worse participation and health-related quality of life. Future studies should examine whether these patients may benefit from rehabilitation aimed at fatigue.

Key words: fatigue; participation; health-related quality of life; mood; coping; perimesencephalic subarachnoid haemorrhage; stroke; subarachnoid haemorrhage.

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Lay abstract

A subarachnoid haemorrhage (SAH) is a subtype of stroke. Of all patients with SAH, approximately 10% are diagnosed with non-aneurysmal perimesencephalic subarachnoid haemorrhage (PM-SAH). PM-SAH is generally considered a benign form of SAH; however we have previously found that one-third of patients with PM-SAH are still fatigued 5 years after PM-SAH. Fatigue may be related to reduced participation and health-related quality of life, both of which are considered important rehabilitation outcomes. Therefore, this study examined whether fatigue is associated with participation and health-related quality of life after PM-SAH. The results showed that, 5 years after PM-SAH, fatigued patients had worse participation and quality of life than non-fatigued patients. In addition, more severe fatigue was associated with worse participation, regarding frequency, satisfaction and restrictions, and with worse health-related quality of life. Further studies are necessary to determine whether patients with PM-SAH may benefit from rehabilitation aimed at fatigue.
long-term follow-up after treatment, and hence they often have a shorter hospital stay than patients with aneurysmal SAH (A-SAH) (4, 5). Because of their good prognosis, patients with PM-SAH are generally not referred to rehabilitation programmes. However, studies focusing on long-term neuropsychological outcome have shown that up to 7.5 years after onset, 26–62% of patients with PM-SAH report symptoms such as headaches, irritability, depression, forgetfulness, deconditioning and fatigue (6–8).

Fatigue is one of the most common and debilitating sequelae of SAH (9). In a previous study, we have found that 5 years post-PM-SAH onset, 33% of patients were still fatigued and that more severe fatigue was related to depression, comorbidity and worse cognitive functioning (8). In patients with A-SAH, the prevalence of fatigue 4–5 years after ictus was 60% (10) to 65% (11), respectively, and, in healthy subjects, 18% of people were found to be fatigued based on the same criteria as used in our previous study (12). Although the prevalence of fatigue is lower in patients with PM-SAH than in patients with A-SAH, fatigue is substantially more common in patients with PM-SAH than in healthy subjects. This endorses the importance of studying the possible negative consequences fatigue may have in this patient group.

Chronic fatigue may affect participation (13) and health-related quality of life (HR-QoL) (14), which are both considered important rehabilitation outcomes. Studies on participation and HR-QoL after PM-SAH are scarce, and studies that have been conducted have either a small sample size and/or did not assess outcome with validated questionnaires. Therefore, the results of these studies are difficult to compare and inconclusive: some studies have shown reduced HR-QoL (15, 16) and difficulties returning to work (6, 17), while others have reported good HR-QoL (18) and no problems returning to work (19, 20).

In patients with PM-SAH, it is not known what the role of fatigue is in participation and HR-QoL outcome. However, in patients with A-SAH, fatigue has been found to be related to worse HR-QoL (14, 21) and reduced participation in terms of functional performance and autonomy and restrictions in participation (13). Just as in patients with A-SAH, participation and HR-QoL might also be reduced in those patients with PM-SAH who experience long-term fatigue. If so, this may indicate that patients with PM-SAH could benefit from rehabilitation aimed at managing and coping with fatigue, since this may enhance their HR-QoL and participation. The aim of the current study was therefore to determine whether fatigue is associated with participation and HR-QoL in the long-term after PM-SAH, taking into account patient characteristics, depression, anxiety and coping as potential confounders. It was hypothesized that fatigued patients with PM-SAH would have lower levels of participation and HR-QoL than non-fatigued patients, and that fatigue severity would be independently associated with HR-QoL and participation.

**METHODS**

**Participants and procedures**

Patients diagnosed with PM-SAH, were identified from a data registry of all consecutive SAH patients who were hospitalized between 2006 and 2012 at the neurology or neurosurgery departments of Erasmus MC University Medical Centre Rotterdam or University Medical Center Utrecht. After screening for eligibility, patients were invited to participate in a cross-sectional study via a letter from their neurologist, sent by post, comprising information about study protocol, rationale and burden of participating in the study. Patients who were interested in participating were asked to return a reply form using a pre-paid envelope. Patients were included if they survived PM-SAH and were at least 18 years of age. Exclusion criteria were: serious (neurological) comorbidity and insufficient understanding of the Dutch language. PM-SAH was defined by an accumulation of blood in the cisterns around the mesencephalon on computed tomography (CT) and the absence of an aneurysm on 4-vessel angiogram (2, 3). Prior to inclusion, all patients signed an informed consent form. The study was approved by the medical ethics committee of Erasmus MC and UMC Utrecht.

**Data collection**

After patients provided informed consent, they were visited at home by a research psychologist to collect data on multiple outcome measures using validated questionnaires (8). In addition, patient characteristics, including years of education, employment status (paid job vs retired/no paid job), hypertension at time of home visit (yes/no), and discharge destination (home/home with outpatient rehabilitation vs rehabilitation centre/nursery home) were collected in a structured interview. In order to minimize fatigue during testing, the home visit was designed in such a way that fatigue was assessed in the first part of the test battery. In addition, all tests were provided in a fixed order with regular breaks between tests. Duration of the home visit varied, with a maximum of 150 minutes. If the tests were not completed after 150 minutes, participants were asked to return any remaining questionnaires by post the next day. In this study, a secondary analysis was conducted.
on the data of the self-report questionnaires on fatigue, participation, quality of life, depression and anxiety, coping and comorbidity.

**Measurement instruments**

Fatigue was assessed with the Fatigue Severity Scale (FSS) (22), which consists of 9 statements on the impact of fatigue in daily life situations, which are rated on a 7-point Likert-scale, ranging from 1 (strongly disagree) to 7 (strongly agree). A cut-off score of 4 or higher was used to distinguish fatigued patients from non-fatigued patients. The FSS shows good internal consistency and test-retest reliability in patients with stroke (12, 22).

Participation was assessed with the Utrecht Scale for Evaluation of Rehabilitation-Participation (USER-P) (23). The USER-P assesses 3 aspects of participation, frequency of participation, restrictions in participation and satisfaction with participation. For all 3 aspects questions are related to work/education, household activities, sports/physical exercise, relationship with a partner and social and leisure activities. The frequency subscale is divided into 2 parts; the first part consists of 4 items about vocational activities in an ordinary week, scored on a scale from 0 (not at all) to 5 (≥36 h). The second part consists of 7 items about participation in social and leisure activities in the last 4 weeks, scored on a scale from 0 (not at all) to 5 (19 times or more). The restrictions subscale consists of 11 items to assess whether the person experienced participation restrictions due to his/her health condition. Items are scored on a scale from 0 (not possible at all) to 5 (very much). In case the experienced restrictions are not a result of the health condition, there is a “not applicable” answer option. Satisfaction with participation was assessed with 10 items, scored on a scale from 0 (not satisfied at all) to 4 (very satisfied), including a “not applicable” option for the items on work/education and relationship with a partner. Sum scores of the applicable questions are converted to a score ranging between 0 and 100 for all 3 subscales, where higher scores reflect better participation (higher frequency, less restrictions, higher satisfaction). In addition, the score on the USER-P Restrictions scale was dichotomized (USER-P-Restricted), where a score <100 indicates having restrictions and a score of 100 equals no restrictions. The USER-P has been found to be valid in patients who visited outpatient rehabilitation in the Netherlands, including stroke patients (23).

HR-QoL was assessed with the Stroke-Specific Quality of Life-12 (SS-QoL-12) scale and is expressed in a total, physical, and psychosocial score. The questionnaire contains 12 items, where 6 items relate to the physical scale and 6 items to the psychosocial scale. Items are rated on a scale from 1 (poor HR-QoL) to 5 (good HR-QoL). The physical scale encompasses domains of self-care, mobility, upper extremity function, language, vision, and work, and the psychosocial scale includes thinking, family roles, social roles, personality, mood, and energy domains. Mean scores are calculated for the total scale and the 2 subscales, with higher scores indicating better quality of life (24). The SS-QoL-12 has been validated in patients with stroke, including patients with SAH (24).

Symptoms of anxiety and depression in the past 4 weeks were assessed with the Hospital Anxiety and Depression Scale (HADS) (25), which consists of 7 questions related to anxiety and 7 questions related to depression. Patients rate all items on a 4-point scale ranging from 0 (no symptoms) to 3 (serious symptoms). The total sum score ranges between 0 and 21 for both the depression and anxiety scale. Higher scores reflect more depressive or anxiety symptoms. A cut-off score of 8 or higher indicates the presence of depression/anxiety (25). The Dutch version of the HADS has been validated in several (patient) groups (26).

Coping strategy was assessed with the Coping Inventory for Stressful Situations (CISS) (27). The CISS consists of 48 items, divided over 3 subscales each with 16 questions regarding a task-oriented, an emotion-oriented, or an avoidant coping strategy. Items are scored on a 5-point scale, ranging from 1 (not at all) to 5 (very much). Sum scores (range 16–80) are calculated and indicate to what extent people react in a certain way when they encounter a stressful situation. Higher scores on each subscale reflect using that coping strategy more often (27). The psychometric properties of the CISS have been found acceptable to good in a Dutch population of patients with acquired brain injury (including stroke patients) (28).

Comorbidity was assessed with the Cumulative Illness Rating Scale (CIRS), which measures impairments in 14 body systems on a scale from 0 (no impairments) to 4 (life-threatening impairments). Sum scores are calculated, ranging from 0 to 56, with higher scores indicating more severe comorbidity (29).

**Statistical analysis**

Statistical analyses were performed using IBM SPSS Statistics version 25 (SPSS Inc., Chicago, IL, USA). Descriptive statistics were used to summarize patient characteristics and outcomes regarding depression and anxiety of the total study population and the fatigued and non-fatigued subgroups. Dependent on the assumption of normally distributed data, either independent t-tests or Mann–Whitney U tests were carried.
out to examine differences in patient characteristics, and participation and quality of life outcomes between fatigued and non-fatigued patients. χ² tests were used for categorical variables.

Univariate and multivariate linear regression analyses were performed to examine the association of fatigue with participation (USER-P Frequency and USER-P Satisfaction) and HR-QoL (SS-QoL-12 total score), adjusting for patient characteristics, depression and anxiety, coping and comorbidity. In addition, univariate and multivariate logistic regression analyses were conducted to examine relationships of the aforementioned determinants with the dichotomized variable USER-P Restricted. Variables that were significant (p < 0.05) in univariate regression analyses, were included in the multivariate regression using backward selection. Assumptions of linearity, multicollinearity, and normally distributed, homoscedastic and independent residuals, were met. A significance level of p < 0.05 was used and a Bonferroni correction was applied to correct for multiplicity.

RESULTS

Study population
In total, 46 patients were included, with a mean time post-onset of PM-SAH of 4.7 years (SD 1.6). Mean age was 50.4 years (SD 9.4) and 63% of subjects were men. Overall, the mean depression and anxiety scores were below the cut-off score of 8, indicating no symptoms of depression or anxiety. Mean fatigue score of the total group was 3.4 (SD 1.3) and 33% had long-term fatigue (8). No significant differences were found between fatigued and non-fatigued patients in patient characteristics, except for hypertension. Of all 46 patients, 1 patient received inpatient rehabilitation and 45 were discharged to their home. None of the patients who were discharged home received outpatient rehabilitation. The mean scores on the depression and anxiety subscales of the HADS differed significantly (p < 0.001) between fatigued and non-fatigued patients. The proportion of patients with depressive (p < 0.05) or anxiety (p < 0.001) symptoms was also significantly larger in the fatigued group than in the non-fatigued group. Fatigued patients used an emotion-oriented coping strategy more often than non-fatigued patients (p < 0.05) and had more comorbidity (Table I).

Fatigued and non-fatigued patients

Fatigued patients scored significantly worse than non-fatigued patients on the frequency subscale (p = 0.001), and the restrictions and satisfaction subscales (p < 0.001) of the USER-P (Fig. 1). In addition, there were significantly (p < 0.001) more patients who had restrictions in participation in the fatigued group (n = 13, 86.7%) than in the non-fatigued group (n = 5, 16.1%). Also regarding HR-QoL, fatigued patients scored significantly worse than non-fatigued patients (p < 0.001), and both the physical (p < 0.001) and psychosocial scales (p < 0.001) (Fig. 1). Correlations between participation and HR-QoL outcomes were: USER-P Frequency and SS-QoL-12: r = 0.341, p = 0.20; USER-P Restrictions.

Table I. Patient characteristics and outcome 5 years after perimesencephalic subarachnoid haemorrhage (PM-SAH) onset

| Patient characteristics | Total group (n = 46) | Fatigued, FSS ≥ 4 (n = 15) | Non-fatigued, FSS < 4 (n = 31) |
|-------------------------|---------------------|---------------------------|-------------------------------|
| Age at onset, years, mean (SD) | 50.4 (9.4) | 50.5 (9.3) | 50.3 (9.6) |
| Sex, men, n (%) | 29 (63.0) | 10 (66.7) | 19 (61.3) |
| Partner, yes, n (%) | 39 (84.8) | 11 (73.3) | 28 (90.3) |
| Education, years, mean (SD) | 14.4 (3.9) | 13.2 (2.8) | 15.0 (4.3) |
| Employed before SAH, n (%) | 36 (78.3) | 11 (73.3) | 25 (80.6) |
| Hypertension, yes, n (%) | 12 (26.1) | 7 (46.7) | 5 (16.1)* |
| Discharge destination, n (%) | | | |
| Home | 45 (97.8) | 15 (100) | 30 (96.8) |
| Inpatient rehabilitation | 1 (2.2) | 0 (0) | 1 (3.2) |
| Outcome after 5 years | | | |
| HADS Depression, mean (SD) | 3.6 (3.6) | 6.4 (3.0) | 2.2 (3.0)** |
| HADS Anxiety, mean (SD) | 5.5 (4.0) | 8.9 (4.6) | 3.9 (2.5)** |
| HADS Depression ≥ 8, n (%) | 7 (15.2) | 5 (33.3) | 2 (6.5)* |
| HADS Anxiety ≥ 8, n (%) | 11 (23.9) | 9 (60.0) | 2 (6.5)** |
| Coping strategy (CISS) | | | |
| Task-oriented, mean (SD) | 55.9 (9.7) | 52.3 (8.8) | 57.5 (9.8) |
| Emotion-oriented, mean (SD) | 36.7 (11.9) | 42.2 (12.5) | 34.4 (11.1)* |
| Avoidant, mean (SD) | 36.3 (9.7) | 35.2 (9.6) | 36.7 (9.8) |
| Fatigue Severity Scale | 36.7 (9.8) | 39 (84.8) | 25 (80.6) |
| Comorbidity (CISS) | 4.0 (3.0) | 6.5 (3.5) | 2.8 (1.7)** |

*p < 0.05; **p < 0.001. HADS Depression and Anxiety range 0–21. CISS subscales range 16–80. HADS and CISS: higher score means worse condition. CISS higher score reflects use of that coping strategy more often.
FSS: Fatigue Severity Scale; HADS, Hospital Anxiety and Depression Scale; CISS: Coping Inventory for Stressful Situations; CIRS: Cumulative Illness Rating Scale Missing; CISS total=2, 2 in fatigued group.
and SS-QoL-12: $r = 0.697, p < 0.001$; USER-P Satisfaction and SS-QoL-12: $r = 0.681, p < 0.001$. Outcomes of the total group and the non-fatigued and fatigued group are shown in Table S1.

**Associations between fatigue, and participation and HR-QoL outcomes**

Table II shows the outcomes of the univariate and multivariate linear regression analyses. In the univariate models, years of education, fatigue, depression, anxiety, comorbidity and emotion-oriented coping were significantly associated with participation in both the USER-P Frequency and Satisfaction domains, and with HR-QoL. In the multivariate model, fatigue severity remained significantly ($p < 0.005$) associated with the frequency of participation, ($R^2 = 0.257$), and with participation satisfaction, adjusted for depression ($R^2 = 0.594$). In addition, more severe fatigue was significantly ($p < 0.001$) associated with worse HR-QoL, adjusted for depression and anxiety ($R^2 = 0.791$) (Tables II and III).

Univariate logistic regression analyses showed that hypertension, fatigue, depression, anxiety and comorbidity were significantly ($p < 0.05$) associated with having restrictions in participation. In multivariate analysis more severe fatigue remained significantly ($p < 0.005$) associated with having restrictions in participation 5 years post PM-SAH onset, adjusted for anxiety, hypertension and comorbidity (Table SII).

**DISCUSSION**

The results of this study suggest that fatigue plays a key role in participation and HR-QoL after PM-SAH. This is supported by our finding that 5 years after PM-SAH fatigued patients participated less frequently in

| Table II. Univariate and multivariate regression models for participation and health-related quality of life outcome 5 years after perimesencephalic subarachnoid haemorrhage (PM-SAH) |
|---------------------------------------------------------------|
| **User-P Frequency**                                          | **User-P Satisfaction** | **SS-QoL-12 Total** |
| **Variables**                                                 | Univariate | Multivariate | Univariate | Multivariate | Univariate | Multivariate |
| Age at onset, per year                                        | B          | p-value      | B          | p-value      | B          | p-value      | B          | p-value      | B          | p-value      |
| Sex, men vs women                                            | -0.17      | 0.235        | -0.17      | 0.492        | 0.01       | 0.338        |
| Partner, yes vs no                                           | -1.22      | 0.662        | -7.23      | 0.119        | -0.04      | 0.805        |
| Education, per year                                          | 0.85       | 0.011        | 1.40       | 0.013        | 0.04       | 0.040        |
| Pre-SAH employment status, employed vs unemployed            | -0.85      | 0.795        | -1.98      | 0.719        | 0.01       | 0.949        |
| Hypertension, yes vs no                                      | -3.35      | 0.271        | -6.00      | 0.242        | -0.13      | 0.487        |
| FSS Fatigue                                                  | -3.70      | 0.000        | -3.62      | 0.000        | -3.01      | 0.000        | -3.01      | 0.000        | -0.11      | 0.000        | -0.03      | 0.075        |
| HADS Depression                                              | -1.15      | 0.001        | NS         | -3.01      | 0.000        | -2.07      | 0.000        | -3.01      | 0.000        | -0.11      | 0.000        | -0.03      | 0.075        |
| HADS Anxiety                                                 | -0.87      | 0.007        | NS         | -2.40      | 0.000        | NS         | -0.11      | 0.000        | -0.05      | 0.001        |
| CISS Task-Oriented                                           | 0.18       | 0.217        | 0.14       | 0.564        | 0.01       | 0.174        |
| CISS Emotion-Oriented                                        | -0.32      | 0.049        | NS         | -0.60      | 0.001        | NS         | -0.02      | 0.000        | NS         | -0.01      | 0.475        |
| CISS Avoidant                                                | -0.17      | 0.237        | 0.01       | 0.985        | -0.01      | 0.000        | -0.078     | 0.005        | NS         | -0.01      | 0.475        |

**Fig. 1.** (A) Participation and (B) health-related quality of life of fatigued and non-fatigued patients 5 years after perimesencephalic subarachnoid haemorrhage (PM-SAH). *p = 0.001; **p < 0.001. USER-P: Utrecht Scale for Evaluation of Rehabilitation Participation; SS-QoL-12: Stroke Specific Quality of Life-12; FSS: Fatigue Severity Scale; Fatigued (FSS ≥ 4) n = 15; non-fatigued (FSS < 4) n = 31. USER-P and SS-QoL-12 higher score means better condition.
daily life activities, had more participation restrictions and were less satisfied with their participation than non-fatigued patients. In addition, fatigue severity was associated with both participation and HR-QoL outcomes, independent of depression, anxiety, hypertension or comorbidity.

Overall, 5 years post-PM-SAH patients have a rather good participation regarding frequency, restrictions and satisfaction. However, strikingly, both frequency of participation and perceived restrictions in participation in the subgroup of fatigued patients in the current study were comparable to patients 1 year post-stroke (75% ischaemic) (30) and to patients up to 4 years post-stroke (92% ischaemic) (31), respectively. In addition, satisfaction with participation in fatigued patients with PM-SAH was only slightly higher than in patients 1 year post-stroke (30). Thus, although, in general, patients with PM-SAH seem to have a benign course regarding long-term participation outcome, the current results suggest that a subgroup of patients with PM-SAH who experience fatigue have a comparable level of problems in participation as patients post-stroke.

Regarding HR-QoL, patients with PM-SAH overall reported a rather good HR-QoL as assessed with the SS-QoL-12. This is in agreement with studies in patients with PM-SAH, who used the Short-Form-36 (SF-36) (15, 16, 20) and the Sickness Impact Profile (SIP) (18) to assess HR-QoL. In the current study, fatigued patients had worse HR-QoL than non-fatigued patients, and the psychosocial domain was most affected. This is in agreement with a previous study in patients with SAH, which found that HR-QoL, also assessed with the SS-QoL-12, was lowest in the psychosocial domain and primarily regarding the items fatigue and memory (32). Apparently, patients with PM-SAH, and especially those who are fatigued, mainly experience a lower quality of life in domains regarding thinking, memory and social roles and less in domains regarding self-care, mobility and work.

This study found that the severity of fatigue was inversely associated with the level of participation in the 3 domains of frequency, restrictions and satisfaction and with HR-QoL, 5 years after PM-SA. This agrees with studies in patients with A-SAH, that showed that fatigue is related to reduced HR-QoL (21) and reduced participation in terms of functional performance and autonomy and restrictions in participation (13). Other psychological variables related to participation and HR-QoL in the current study included depression, anxiety and emotion-oriented coping. These factors are known to be associated with participation, HR-QoL and fatigue after A-SAH and stroke (13, 14, 21, 33–36). Interestingly, the associations between fatigue and satisfaction with participation, between fatigue and having restrictions in participation, and between fatigue and HR-QoL, remained significant, even after adjusting for depression, and/or anxiety, respectively. Emotion-oriented coping dropped out of the multivariate models, probably because this coping style is also associated with depression and anxiety, which had a stronger association with participation and HR-QoL (28, 37).

Patients with PM-SAH are generally not referred to inpatient or outpatient rehabilitation programmes, which is also reflected in the current study sample, where only 1 patient received rehabilitation. The current results indicate that it would be useful to screen patients with PM-SAH for fatigue in outpatient rehabilitation centres or at the general practitioner, using, for example, the FSS as screening tool, with the cut-off score of 4. In addition, it seems important to be aware of additional psychological characteristics of fatigued patients, such as symptoms of depression or anxiety, which may also be related to an emotion-oriented coping strategy (28, 37). For this subgroup, especially, it should be considered to refer them to rehabilitation programmes aimed at reducing fatigue and coping with fatigue, since this may improve not only the frequency

### Table III. Model selection and performance

| Model | USER-P | | USER-P | | SS-QoL-12 |
|-------|--------|--------|--------|--------|--------|
|       | Frequency | Change | p-value | Satisfaction | Change | p-value | Total | Change | p-value |
| 1     | Education, FSS, HADS-D, HADS-A, CISS-E, CIRS | 0.352 | 0.010 | Education, FSS, HADS-D, HADS-A, CISS-E, CIRS | 0.624 | 0.000 | Education, FSS, HADS-D, HADS-A, CISS-E, CIRS | 0.796 | 0.000 |
| 2     | Education, FSS, HADS-D, CISS-E, CIRS | –0.001 | 0.832 | Education, FSS, HADS-D, HADS-A, CISS-E, CIRS | –0.001 | 0.821 | Education, FSS, HADS-D, HADS-A, CISS-E, CIRS | 0.000 | 0.842 |
| 3     | Education, FSS, HADS-D, CISS-E, CIRS | –0.004 | 0.739 | Education, FSS, HADS-D, HADS-A | –0.004 | 0.512 | Education, FSS, HADS-D, HADS-A | –0.000 | 0.791 |
| 4     | Education, FSS, CISS-E | –0.014 | 0.242 | FSS, HADS-D, HADS-A | –0.014 | 0.235 | FSS, HADS-D, HADS-A | –0.005 | 0.338 |
| 5     | Education, FSS | –0.033 | 0.168 | FSS, HADS-D | –0.011 | 0.295 | |
| 6     | FSS | –0.044 | 0.118 | |
|       | R total | 0.257 | | R total | 0.594 | | R total | 0.791 |

USER-P: Utrecht Scale for Evaluation of Rehabilitation-Participation; SS-QoL-12: Stroke Specific Quality of Life-12; FSS: Fatigue Severity Scale; HADS: Hospital Anxiety and Depression Scale; HADS-D: Depression; HADS-A: Anxiety; CISS-E: Coping Inventory for Stressful Situations Emotion-oriented coping; CISS: Cumulative Illness Rating Scale.
of participation, but also the experienced restrictions and satisfaction with participation, and HR-QoL in the long-term.

One of the strengths of the current study is that different aspects of participation and HR-QoL were assessed with validated questionnaires in a relatively large group of patients with PM-SAH, particularly considering the rather low incidence of PM-SAH (1). In addition, the current study not only considered participation and HR-QoL of the total group, but also of the fatigued group and non-fatigued group separately. This revealed that the FSS and the cut-off score of 4 can be a useful and appropriate tool to distinguish fatigued from non-fatigued patients in clinical practice. Finally, depression, anxiety, coping, and patient characteristics were taken into account, thereby enabling analysis of the independent role of fatigue in participation and quality of life outcome.

One of the limitations of the study is that it is not known whether fatigue is causally related to participation and HR-QoL, since the study had a cross-sectional design. It would be of interest to examine a possible causal relationship in future longitudinal cohort studies. In addition, it was necessary to preselect possible predictors for the multivariate analysis, since the number of predictors to take into account in the multivariate analysis was limited due to the small sample size of 46 patients. Finally, other personality factors, such as self-efficacy and neuroticism, are known to be associated with fatigue, quality of life and participation after SAH or stroke (14, 35, 38, 39). These variables were not taken into account in the current study, but this would be of interest in future studies.

To summarize, 5 years after non-aneurysmal PM-SAH, fatigued patients had worse participation and HR-QoL than non-fatigued patients, and fatigue severity was related to multiple aspects of participation and HR-QoL. Future studies should examine whether patients with PM-SAH may benefit from rehabilitation aimed at fatigue, in terms of enhanced participation and HR-QoL.

The authors have no conflicts of interest to declare.

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