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Does upper limb strength play a prominent role in health-related quality of life in stroke patients discharged from inpatient rehabilitation?

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ABSTRACT

Background: Impairments in arm function are a common problem in stroke survivors and have a large impact on health-related quality of life (HRQoL). Little is known about the longitudinal relationship between recovery of upper limb strength and changes in HRQoL.

Objectives: This study aimed to determine to what extent changes in HRQoL are related to changes in upper limb strength after discharge from inpatient rehabilitation.

Methods: 250 patients from an RCT were assessed at discharge from inpatient rehabilitation (baseline) and at 12 weeks post-discharge (follow-up). The Stroke Impact Scale was used to measure HRQoL, and the Motricity Index Arm was used to measure upper limb strength. Hierarchical regression analysis was performed to determine the predictive value of upper limb strength on HRQoL, relative to demographic and clinical characteristics. Regression analysis was used to determine the relation between upper limb strength improvement and HRQoL improvement.

Results: Upper limb strength at baseline was a major predictor of HRQoL at follow-up, after accounting for demographic and clinical characteristics (p < .05). Improvement in HRQoL was positively related to improvement in upper limb strength (F(1, 240) = 18.351, p < .0005).

Conclusions: These findings highlight the importance of upper limb strength in HRQoL, as HRQoL is associated with improvement in upper limb strength recovery. Better monitoring of recovery and treatment of upper limb strength during the outpatient rehabilitation period and beyond, i.e. outside the typical time-window of recovery in the first 3 months post-stroke, might contribute to higher quality of life for stroke survivors.

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KEYWORDS

Upper limb function; selfreported functional status; quality of life; stroke; outcome measures; rehabilitation

Introduction

Stroke is a major health problem across the world causing complex disability.¹ The impact of this common and serious condition on an individual's life is considerable: physical, psychological, and social consequences can be experienced.^{2–4} Upper limb paresis, a muscle weakness in the affected limb to one side of the body, is one of the most frequently occurring conditions (up to 85% of stroke survivors).^{5,6} Improvement in upper extremity motor function occurs mainly in the first few months after stroke.⁷ At 6 months post-stroke, estimates pointed out that some dexterity in the paretic arm is found in 38% of the stroke patients who show no dexterity in the first week post-stroke.^{7–9}

Arm function plays a critical role in the performance of daily life activities. Most everyday activities require the use of both hands, and because of this, performance of bimanual activities receives considerable attention in the rehabilitation setting.¹⁰ Improved arm and hand function positively contribute to societal participation and (health-related) quality of life.^{1,10,11} Health-related quality of life (HRQoL) can be defined as an individual's (or group's) perceived physical and mental health over time.¹² There is a growing body of literature that recognizes that different factors influence HRQoL after stroke.^{1,4,13,14} A cross-sectional study has shown that the extent of upper limb improvement positively influences a patient's perception of what

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activities can be performed, which in turn enhances HRQoL.¹ Incomplete motor recovery of the upper and lower extremities has been found to be the strongest predictor of a lower HRQoL in an observational study.⁴

Whilst some research has been carried out on the association between arm function and HRQoL^{1,4,13,14}, there has been no longitudinal investigation of improvement in HRQoL in relation to improvement in arm recovery. Obtaining insights into this relationship will be useful for understanding problems faced by patients and for planning and optimization of rehabilitation treatment after stroke.

The first aim of the present study was to identify the relation between upper limb strength and HRQoL at discharge from inpatient rehabilitation and at follow-up (12 weeks later). Second, we aimed to determine whether upper limb strength at discharge from inpatient rehabilitation predicts HRQoL at follow-up, when corrected for patient and stroke characteristics. Third, we aimed to determine whether a change in upper limb strength is related to a change in HRQoL over time. We hypothesized that an improvement in upper limb strength is positively related to an improvement in HRQoL.

Methods

Data for this study were collected during the FITstroke trial.¹⁵ The FIT-Stroke trial is a stratified, multicentre single-blinded randomized controlled trial to investigate the effectiveness of taskoriented circuit class training. Stroke patients were recruited in nine outpatient rehabilitation centers in the Netherlands, between June 2008 and December 2010. All included patients completed an inpatient rehabilitation period (average of 3 months) and were included at the start of their outpatient rehabilitation period. The inpatirehabilitation consisted ent period of a multidisciplinary approach to reach complex (physical and cognitive) rehabilitation goals. The patients were treated by nurses, physical therapists, occupational therapists, speech therapists, social workers, and psychologists. Outcomes from assessments at baseline (start outpatient rehabilitation period) and 12 weeks after randomization were

used in this study. Full details of the FIT-Stroke trial have been reported elsewhere.¹⁵

The FIT-Stroke protocol has been approved by the Medical Ethics Committee of the University Medical Center Utrecht and all the participating rehabilitation centers. The trial is registered in the Dutch Trial Register (NTR1534). The content of this manuscript conforms to STROBE guidelines.¹⁶

Participants

For inclusion, patients were eligible if they 1) had a verified stroke according to the WHO definition; 2) were discharged from a rehabilitation center; 3) needed to continue physiotherapy during outpatient care to improve walking competency or physical condition, or both; and 4) were able to give informed consent and motivated to participate in a 12-week intensive program of physiotherapy. Patients affected by cognitive deficits (Mini Mental Status Examination, <24 points), who were not able to communicate (Utrechts Communicatie Onderzoek (UCO), <4 points) or lived more than 30 km from the rehabilitation center were excluded. All participants provided written informed consent. The study size was based on a power analysis.¹⁵

Outcome measures

Health-related quality of life: stroke impact scale (SIS), version 3

The SIS 3.0 is a multidimensional, self-reported stroke-specific 59-item instrument that assesses HRQoL in eight domains related to activities and participation (including strength, memory, emotion, communication, activities of daily living (ADL), mobility, hand function, and participation). The SIS has been shown to have good psychometric properties in a group of stroke survivors.¹⁷ All 59 items are rated using a 5-point Likert scale and scored from "unable to complete the item" to "no difficulty experienced completing the item." Each domain of the SIS has a range of 0-100 with higher scores indicating better quality of life. An extra question in the recovery domain assesses how much the patient feels that he/she has recovered from his/her stroke.¹⁷

Independent variables

Motricity index (MI), arm

The upper extremity subscale of the Motricity Index (MI arm) was used to assess muscle strength of the paretic arm. The MI arm gives a rapid and reliable measure of the upper limb strength and uses a 6-point ordinal scale, with higher scores indicating better upper limb strength. The MI has been shown to provide good validity and reliability as a tool in stroke research.¹⁸ Pinch grip, elbow flexion, and shoulder abduction were assessed. On each dimension, scores ranged from 0 (no activity) to 33 (maximal muscle force).¹⁸

Motricity index (MI), leg

The lower extremity subscale of the Motricity Index (MI leg) was used to assess muscle strength and voluntary movement of the paretic leg. Ankle dorsiflexion, knee extension, and hip flexion were assessed. The scoring system is similar to the MI arm described above. The MI has been shown to provide good validity and reliability as a tool in stroke research.¹⁸

Hospital anxiety and depression scale (HADS)

The HADS is a self-reported measure for assessing anxiety and depression in patients, but it does not provide a specific diagnosis.¹⁹ It is a 14-item scale with 7 items for anxiety and 7 items for depression. The items use a four-point rating scale and patients can score between 0 and 3 points per item. The HADS has been proven to be a reliable, valid, and practical psychological screening tool, with lower scores indicating lower risk of anxiety and/ or depression disorders.¹⁹

Fatigue severity scale (FSS)

The FSS is a self-administered questionnaire with nine questions that examine the perceived severity of fatigue symptoms in different situations in the past week.²⁰ The patient indicates to what extent fatigue determines functioning. Scores for each item range from 1 (strong disagreement) to 7 (strong agreement). The FSS is a valid and reliable scale to measure fatigue in stroke.²⁰

Demographic and clinical variables

Age, gender, stroke type, side of hemiplegia and dominant side affected were assessed as potential predicting variables of HRQoL.

Data analysis

We statistically tested for differences in change scores on HRQoL, arm- and leg function between the circuit class training group and the usual care group 12 weeks after randomization to determine if both groups could be included in the analysis. SIS total score was the dependent variable in the regression analysis. Age, gender, clinical variables (stroke type, side of hemiplegia and dominant side affected) and clinical scales (MI arm and leg, HADS, FSS) at baseline and at follow-up were considered as independent variables. We tested whether the assumptions for a linear regression analysis were met. Change-frombaseline scores of the SIS and MI-arm were computed to describe the data.

To determine the relationship between HRQoL and upper limb strength, cross-sectionally at baseline and follow-up, univariate regression analysis was performed. Variables demonstrating *p*-values <0.20 were included in the hierarchical regression analysis.

To determine how much variance in HRQoL at follow-up was explained by upper limb strength and other demographic and clinical variables at baseline, hierarchical regression analysis was performed. HRQoL at baseline post-stroke was entered as first block into the analysis (to control for the effect of the dependent variable) and upper limb strength as second block. Clinical variables that could be included in the hierarchical regression analysis were combined as the third block and demographic and stroke characteristics were added in the fourth block. Potential predictor variables were examined for collinearity by inspection of the correlation coefficients (no multicollinearity when coefficients <0.7) and Tolerance/VIF values (Tolerance needed to be >0.1 and VIF values <10).

Univariate regression analysis was also performed with change-from-baseline scores for HRQoL, as dependent variable, and change-frombaseline scores in upper limb strength. Effect sizes were classified as .02, .15 and .35 as small, medium, and large effect sizes.²¹ Significance levels were set at p = .05. Statistical analyzes were performed with SPSS, version 25.0.

Results

In total, 250 participants were included in this study (flowchart; Supplementary Figure 1). Of the 250 included patients, one patient in the circuit training group and seven in the usual care group were excluded from the analysis. Reasons were withdrawal from participation (n = 3), death from cancer (n = 2), and recurrent stroke (n = 2), while one patient missed the 12-week assessment visit because of change of address.¹⁵ The change scores on HRQoL, arm- and leg function from circuit class training group and control group did not differ significantly from each other 12 weeks after randomization. Table 1 lists the clinical and demographic characteristics of the patients. The baseline assessment was done 102 days (SD 64) post-stroke and the follow up assessment 12 weeks after baseline. The average age in the population was 57 years (SD 10), 65% (n = 162) was male, the majority (n = 203) had an ischemic stroke, and almost half of the patients (n = 116) had a right hemisphere lesion. From this table, we can see that there are significant improvements in mean scores over time for most clinical variables (p < .05), except the emotion subscale (SIS), FSS, and HADS.

For HRQoL (SIS), the mean score at baseline was 70.30 (SD 11.54) and 3 months later the mean was 75.61 (SD 13.78). The highest score at baseline was 94. One patient reached the maximum score of 100 at the follow-up assessment. Change-from-baseline scores show that 57 patients (24%) had a negative change score (decline), and the remaining patients had a neutral (change score of 0; 3%) or positive change score (improvement; 73%). The lowest and highest change-from-baseline scores were -23 and +40, respectively.

Table 1. Baseline characteristics (n=250)

Characteristics	Baseline	Follow-up*	P value
Age (years)	57 (10)		
Gender, n (% male)	162 (65)		
Stroke type, n (%)			
Ischemic	203 (81)		
Haemorrhagic	47 (19)		
Site of stroke, n (%)			
Right hemisphere	116 (46)		
Left hemisphere	91 (36)		
Brainstem	20 (8)		
Cerebellum	4 (2)		
Time since onset, days	102 (64)		
Stroke Impact Scale ^a			
Strength (0–100)	51.80 (20.27)	59.61 (22.82)	<.01
Memory (0–100)	81.83 (17.48)	87.08 (15.92)	<.01
Emotion (0–100)	82.56 (13.57)	81.89 (14.51)	.49
Communication (0–100)	84.89 (18.94)	86.51 (18.01)	.04
Activities (0–100)	70.13 (15.44)	77.43 (16.57)	<.01
Mobility (0–100)	79.32 (14.11)	85.56 (12.90)	<.01
Affected hand (0–100)	45.10 (35.59)	55.68 (37.69)	<.01
Participation (0–100)	66.75 (20.84)	71.94 (19.59)	<.01
Recovery (0–100)	56.53 (16.53)	63.56 (17.34)	<.01
Physical functioning (0–100)	60.74 (16.16)	69.57 (19.36)	<.01
Total (0–100)	70.30 (11.54)	75.61 (13.78)	<.01
Fatigue Severity Scale (1–7) ^b	3.98 (1.69)	4.03 (1.67)	.43
Hospital Anxiety and Depression Scale ^b			
Depression (0–21)	4.40 (3.23)	4.28 (4.00)	.53
Anxiety (0–21)	3.63 (3.27)	3.66 (3.55)	.64
Nottingham Extended ADL ^a			
Mobility (0–18)	10.90 (4.21)	13.62 (4.14)	<.01
Kitchen (0–15)	10.06 (3.80)	12.45 (3.53)	<.01
Domestic (0–15)	4.53 (3.93)	7.95 (4.08)	<.01
Leisure (0–18)	7.31 (2.87)	10.39 (3.60)	<.01
Motricity Index ^a		,	
Arm (0–100)	60.37 (26.41)	65.35 (26.27)	<.01
Leg (0–100)	68.13 (20.16)	72.92 (20.41)	<.01

Values are displayed as mean (SD) unless otherwise indicated; *Follow-up was 12 weeks after the baseline assessment, n = 242; ^aHigher mean scores reflect better function; ^bLower mean scores reflect better function.

Mean upper limb strength (MI-arm) was 60.37 (SD 26.41). Eleven patients (4%) had a baseline score of 0 in comparison to 21 patients (8%) with the maximum score of 99 at baseline. Six months post-stroke, mean MI-arm was 65.35 (SD 26.27), and only seven patients (3%) had a score of 0 while the number of patients reaching the maximum score increased to 40 (17%). Change-from-baseline scores showed that 39 patients (16%) had a negative change score, and the remaining patients had a neutral (no change) (28%) or positive change score (56%). The lowest and highest change-from-baseline scores were -23 and +52 points, respectively.

Upper limb strength and health-related quality of life cross-sectionally in time

Linear regression analysis showed that HRQoL was statistically significantly related to upper limb strength at baseline, F(1, 248) = 165.023, p < .0005 and upper limb strength accounted for 40% of the explained variability in HRQoL (Table 2). At follow-up HRQoL was also significantly related to upper limb strength, F(1, 240) = 225.191, p < .0005 and upper limb strength accounted for 48% of the explained variability in HRQoL (Table 2). Table 2 also shows that age, sex, side of hemiplegia, MI-leg, HADS, and FSS in relation to HRQoL at baseline can be included as independent variables in the regression analysis (all p < .2). At follow-up, age and sex

(baseline), MI-leg, HADS, and FSS in relation to HRQoL emerged as independent variables (all p < .2).

Upper limb strength, stroke- and clinical variables, and health-related quality of life over time

HRQoL at baseline (Model 1) was significantly related to HRQoL at follow-up ($R^2 = .644$, F(1,(229) = 414.546, p < .0005) (Table 3). The addition of (improvement in) upper limb strength to the prediction of HRQoL at follow-up (Model 2) led to a statistically significant increase in R^2 of .078, F(2,227) = 31.983, *p* < .0005. The addition of MI-leg, FSS, HADS (Model 3) led to small, non-significant changes in \mathbb{R}^2 (.004) (Table 3). The addition of age, sex, and hemiplegia (full model) led to a small increase in R^2 of .010, which was statistically significant (p < .05, Table 3 for full details). Clinical variables, like leg function (MI-leg), did not show a significant correlation with HRQoL (p > .05). Sex was the only stroke-related characteristic with a significant relation with HRQoL (p < .05).

Changes in upper limb strength and changes in health-related quality of life over time

Change scores in upper limb strength between baseline and follow-up accounted for 7.1% of the variation

	Univariate regression analysis Stroke Impact Scale									
			Baseline				Follow-up			
Candidate predictors	В	95% CI	Beta	р	R ²	В	95% CI	Beta	р	R ²
Demographic										
Age	.136	(003,.276)	.122	.055*	.015	.113	(057,.282)	.084	.191*	.007
Sex	-2.059	(-5.064,.947)	085	.179*	.007	-4.943	(-8.561, -1.326)	171	.008*	.029
Stroke type	.512	(-3.174, 4.199)	.017	.785	.000	.068	(-4.464, 4.600)	.002	.976	.000
Side of hemiplegia ^a	-2.815	(-5.738,.108)	122	.059*	.015	-2.344	(-5.945, 1.257)	084	.201	.007
Dominant side affected ^b	1.183	(-1.764, 4.131)	.051	.430	.003	.241	(-3.383, 3.864)	.009	.895	.000
Clinical										
MI-arm	.276	(.234,.319)	.632	.000*	.400	.365	(.317,.413)	.696	.000*	.484
MI-leg	.278	(.215,.341)	.486	.000*	.236	.364	(.292,.436)	.539	.000*	.291
HADS	-1.048	(-1.247,849)	550	.000*	.303	-1.244	(-1.476, -1.013)	564	.000*	.318
FSS	-1.599	(-2.433,766)	233	.000*	.054	-2.390	(-3.376, -1.404)	295	.000*	.087

Table 2. Univariate regression analyzes SIS and potential predictors, baseline, and follow-up.

MI-arm/leg Motricity Index arm/been; *HADS* Hamilton Anxiety and Depression Scale; *FSS* Fatigue Severity Scale; **p* < 0.2

^aDummy coding: 0 = left, 1 = right.

^bDummy coding: 0 = dominant side affected, 1 = dominant side unaffected; *p*-values are used to determine inclusion in hierarchical regression analysis.

Table 3. Hierarchical regression analysis.

	Stroke Impact Scale, at follow-up								
	В	SE B	β	р	R ²	R ² change	F	р	
Baseline									
Model 1					.644		414.546	.000*	
SIS total	.965	.047	.803	.000*					
Model 2					.722	.078	31.983	.000*	
SIS total	.745	.054	.620	.000*					
MI-arm	.159	.024	.302	.000*					
MI-arm (baseline-follow-up)	.290	.049	.211	.000*					
Model 3					.726	.004	1.040	.376	
SIS total	.699	.068	.582	.000*					
MI-arm	.145	.030	.274	.000*					
MI-arm (baseline-follow-up)	.283	.050	.206	.000*					
MI-leg	.042	.033	.060	.206					
FSS	.231	.331	.028	.484					
HADS	131	.107	057	.223					
Model 4					.736	.010	2.807	.040	
SIS total	.675	.070	.561	.000*					
MI-arm	.152	.030	.288	.000*					
MI-arm (baseline – follow-up)	.276	.049	.201	.000*					
MI-leg	.041	.033	.060	.209					
FSS	.166	.338	.020	.624					
HADS	128	.107	056	.230					
Age	051	.049	038	.297					
Sex	-2.615	1.042	090	.013*					
Hemiplegia ^a	-1.130	1.013	041	.266					

MI-arm/leg Motricity Index arm/been; *HADS* Hamilton Anxiety and Depression Scale; *FSS* Fatigue Severity Scale. ^aDummy coding: 0 = left, 1 = right; *p < .05.

 R^2 = .071; adjusted R^2 = .067 (small to medium effect size, according to Cohen, 1988); F(1, 240) = 18.351, p < .0005.

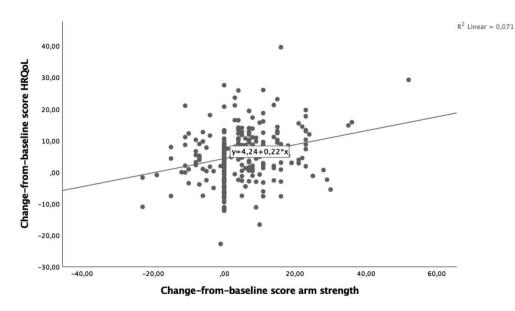


Figure 1. Scatterplot and linear regression fit for change-from-baseline scores between baseline and follow-up.

in change scores in HRQoL between baseline and follow-up, a small to medium effect size (Figure 1).²¹ Change scores in upper limb strength between baseline and follow-up significantly predicted change scores in HRQoL between baseline and follow-up (F (1, 240) = 18.351, p < .0005).

Discussion

This is the first study that assesses the degree and relative impact of the relationship, both cross-sectionally and longitudinally, between upper limb strength and HRQoL. Upper limb strength was significantly related to HRQoL at baseline and follow-up and is an important significant predictor of HRQoL, even after correcting for clinical and stroke characteristics. Furthermore, an improvement in upper limb strength was positively related to an improvement of HRQoL, which suggests that recovery of upper limb strength is also important during the outpatient rehabilitation period and beyond.

Our finding that upper limb strength is independently related to HRQoL is in line with findings from other studies.^{1,4,13,14,22,23} In contrast to earlier studies, however, our study is able to demonstrate the relation between upper limb strength and HRQoL in the sub-acute outpatient rehabilitation phase. In addition, patients were enrolled at a specific moment during stroke rehabilitation, namely upon discharge from inpatient rehabilitation. We considered it useful to explore this sub-acute phase, since it is characterized by minimal rehabilitation support and is beyond the sensitive time-window of recovery.^{24,25} Most earlier studies assessed patients within a broad time frame after stroke (e.g. 2-68 months after stroke place onset), and took in the chronic phase.4,13,14,22,23

Our study highlights the unique contribution of upper limb strength to HRQoL and its importance compared to other predictors. In previous studies arm function was included as one of the many potential predictors of HRQoL and can, therefore, have been easily overlooked, as these studies did not focus solely on arm function.^{1,4,13,14,22} Some previous studies did show that specific domains of HRQoL, i.e. ADL and participation were related to arm function^{13,14}, and that autonomy in daily life activities was strictly related to recovery of the affected arm.²⁶ The current study also demonstrates that upper limb strength contributes more to HRQoL than other predictors, like leg function, which is consistent with the findings of a comparable study⁴, indicating that upper limb strength is the strongest predictor of HRQoL. It should be noted that this has only been demonstrated in mildly to moderately impaired stroke patients, because the inclusion criteria were limited to this group of patients.⁴

In addition to the importance of upper limb strength, there are several factors that predict HRQoL in stroke patients. Our study demonstrated that HRQoL was also cross-sectionally predicted by age, sex, side of hemiplegia, anxiety and depression, leg function, and fatigue. In accordance with most studies, symptoms of anxiety and depression and post-stroke fatigue could affect a patient's motivation to participate in rehabilitation programs and influence (the rate) of recovery.^{1,13,27} Older age is associated with poorer HRQoL in stroke patients as in the general population. The association between (female) gender and poorer HRQoL might be explained by a higher prevalence of anxiety post-stroke.¹ Dominant side of hemiplegia has been repeatedly shown to be associated with HRQoL. Most patients are right-handed with a left hemisphere stroke. Since the left hemisphere controls speech and language function may affect HRQoL through an altered communication ability.^{1,28}

In contrast to the current study, previous studies have not assessed the effect of changes in upper limb strength over time on HRQoL, as HRQoL assessed at a single timewas point.^{1,4,23,29} The current study shows that improvements in upper limb strength are positively related to improvements in HRQoL during outpatient rehabilitation. This suggests that there should also be a focus on upper limb strength recovery in the outpatient rehabilitation phase, in order to possibly expect improvement in HRQoL. The improvement in upper limb strength found during outpatient rehabilitation raises the possibility that there is potential to train outside the sensitive time-window (i.e. 0-3 months poststroke).^{30,31} A recent study that investigated an intensive upper limb neurorehabilitation programme in chronic stroke patients, found large clinical improvements in measures of impairment and activity.³² However, future studies are recommended to confirm the results and unanswered questions about potential for arm recovery beyond the sensitive time-window. Currently, patients receive little to no rehabilitation support when discharged to the community.^{24,25}

The present study had some limitations. Firstly, included patients were indicated for inpatient rehabilitation when discharged from the hospital and had only mild to moderate stroke, which limits the generalizability of the trial. In addition, the included patients were specifically selected for inclusion in a task-oriented circuit class training because of primary problems in walking competency, what might result in a selection of patients with relatively mild upper limb impairment. Nonetheless, only a few patients reached a maximum score on the upper limb clinical measures at baseline. Secondly, we chose to limit this study to upper limb strength, while sensory function may play a prominent role in a patient's perception of arm function.³³ Future studies should determine the role of sensory function in the relation between upper limb strength and HRQoL.

Our study provides evidence that upper limb strength is an important independent predictor of HRQoL in the sub-acute phase after stroke. Upper limb strength appears to be the most important predictor of HRQoL, besides other predictors as leg function and anxiety, and an improvement of upper limb strength contributes positively to HRQoL. Current arm function recovery therapies, e.g. constraint-induced movement therapy,³⁴ should be optimized and evaluated since improvement in upper limb strength positively influences HRQoL. Future studies on the current topic, and outside the sensitive timewindow of recovery, are therefore recommended.

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