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What is the longitudinal profile of impairments and can we predict difficulty caring for the profoundly-affected arm in the first year post-stroke?

Allison, Rhoda; Kilbride, Cherry; Chynoweth, Jade; Creanor, Siobhan; Frampton, Ian; Marsden, Jonathon

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1 **What is the longitudinal profile of impairments and can we predict difficulty caring for**
2 **the profoundly-affected arm in the first year post-stroke?**

3 **Abstract**

4 Objective: To establish the longitudinal profile of impairments of body functions and activity
5 limitations of the arm, and evaluate potential predictors of difficulty caring for the
6 profoundly-affected arm post-stroke.

7 Design: Prospective cohort study.

8 Setting: Three UK stroke services.

9 Participants: People unlikely to regain functional use of the arm (N=155) were recruited at 2-
10 4 weeks post-stroke, and followed up at 3, 6 and 12 months. Potential predictors at baseline
11 were hypertonicity, pain, motor control, mood, sensation/perception, age and stroke severity.

12 Interventions: NA

13 Main Outcome Measures: Difficulty caring for the arm (LASIS), pain, hypertonicity, range of
14 movement, arm function and skin integrity. Multi-variable linear regression identified the
15 best fitting model for predicting LASIS at 12 months.

16 Results: One hundred and ten participants (71%) were reviewed at one year. There was a
17 large variation in the profile of arm functions and activity limitations. Inability or severe
18 difficulty caring for the arm affected 29% of participants. Hypertonicity developed in 77%,
19 with severe hypertonicity present in 25%. Pain was reported by 65%, 94% developed
20 shoulder contracture and 6% had macerated skin. Difficulty caring for the arm increased with
21 age, greater level of hypertonicity and stroke classification; collectively these factors
22 accounted for 33% of the variance in LASIS.

23 Conclusions: At one year post-stroke, there was a high incidence of impairments of body
24 functions and activity limitations in people with a profoundly-affected arm. Individual

25 profiles were very variable and although some pre-disposing factors have been identified, it

26 remains difficult to predict who is at greatest risk.

27 **Key words:** stroke, upper limb, spasticity, pain, contracture

28 **List of abbreviations:**

29 HAEM – Haemorrhage stroke

30 LACS lacunar stroke on Oxford classification

31 LASIS Leeds Arm Spasticity Impact Scale

32 MAL-14 Motor Activity log

33 MMAS Modified modified Ashworth Scale

34 PACS – Partial anterior circulation stroke on Oxford classification

35 POCS – Posterior circulation ischaemic stroke on Oxford classification

36 Q1, Q3 First quartile and third quartile of the inter quartile range

37 TACS total anterior circulation stroke on Oxford classification

38 Three quarters of people with stroke will experience arm weakness, and 62% of these will not
39 recover dexterity at six months¹. For the purposes of this research, the term 'profoundly
40 affected arm' is used to describe the situation where a stroke survivor has no movement in the
41 affected arm or when movement is not functionally useful². While current physical therapies
42 in stroke rehabilitation are based predominantly on exercise and task-specific training^{3,4},
43 additional therapy and practice of tasks does not improve active function in those with the
44 most significant weakness⁵. Hence for those most unlikely to regain active function, a focus
45 on managing activity limitations and avoiding secondary complications may be more
46 appropriate. This approach involves maintaining the ability to care for the arm including tasks
47 such as hand-washing and nail-cutting (i.e. passive function activities⁶ which may be
48 conducted by the person themselves or their carer).

49 Previous research shows that hypertonicity is present as early as one week post-stroke⁷ and
50 affects up to 47% of survivors⁸. Pain can also occur within one week⁹ with an incidence up to
51 49%¹⁰. Contracture is apparent by two weeks and affects 50%¹¹. Previously reported
52 predictors of hypertonicity include reduced motor control^{7,8}, and increased stroke severity^{8,12}.

53 The most common predictors of pain are reduced sensation^{13,14}, and weakness¹³. The
54 significance of depression is not clear, with some studies identifying a positive link with
55 pain¹⁵, and others discounting this⁸. Contracture is most frequently predicted by
56 weakness^{16,17}. However, this previous research is limited as all of these studies have been
57 conducted on general populations of stroke survivors and not targeted at those with the most
58 significant weakness. Furthermore, none of these previous studies has evaluated the profile or
59 potential predictors of difficulty caring for the arm after stroke in a systematic way².

60 Despite the high proportion of people with a profoundly-affected arm post-stroke (62%¹),
61 there is currently no targeted research on (1) the longitudinal profile of activity limitation in
62 caring for the arm, (2) the proportion of people who develop associated impairments of body

63 functions and (3) the relationship between initial clinical findings and subsequent difficulty
64 caring for the arm. The aim of this study was to establish the longitudinal profile of
65 impairment of body functions and activity limitation in people with a profoundly-affected
66 arm, and evaluate potential predictors of difficulty caring for the arm, in the first year post-
67 stroke.

68 **Methods**

69 **Participants**

70 In a prospective, longitudinal study, all adult patients with first or subsequent stroke admitted
71 to three stroke units in the UK over 30 months from September 2011, and still under the care
72 of the stroke team at 2 weeks post-stroke, were screened for inclusion. Criteria included
73 stroke within the past 2-4 weeks and a Fugl-Meyer upper extremity score of equal to or less
74 than 11 points at 2 weeks, or 15 points at 3 weeks, or 19 points at 4 weeks post-stroke. These
75 scores are strongly associated with high probability of not regaining function in the arm¹.
76 Patients who were unable to use their arm before the stroke were excluded. Potential
77 participants were assessed for their ability to consent using the Mental Capacity Act¹⁸. Those
78 with capacity were asked for their consent to participate. If the potential participant was
79 judged by the researcher not to have capacity to make this decision, a consultee was
80 approached if available. A consultee is someone who knows the person well but is not acting
81 in a professional capacity, who can consider the persons beliefs and provide assent on their
82 behalf if this is in line with their interests¹⁹.

83 Participants' baseline data were collected at the point of consent and at 3, 6, and 12 months
84 later in the setting of their choice. These time scales allow comparison with previous
85 studies^{7,8,9,10}. Throughout the study, all participants received usual care under the UK NHS.

86 **Baseline predictor variables and demographic variables**

87 Five potential predictors of difficulty caring for the arm and related impairments were
88 identified²: motor control, mood, sensation/perception, hypertonicity and pain. As the
89 primary outcome related to passive care activities, hand dominance was not considered as a
90 predictor. To maximise inclusivity wherever possible the measures used were suitable for
91 people with aphasia or cognitive impairment. This included using pictographic resources,
92 observational tools and measures with evidence of validity when completed by proxy. The
93 predictor measures are summarised in Table 1^{20,21,22,23,24,25,26,27}. Scores for hypertonicity with
94 the Modified Modified Ashworth Scale (MMAS)²² were applied to the five arm muscles
95 identified as commonly affected (i.e. shoulder adductors and internal rotators, and elbow,
96 wrist and finger flexors)⁸. The single worst score of any muscle group ("*worst hypertonicity*")
97 and the summed score of hypertonicity in all five groups ("*total hypertonicity*") were
98 considered (independently) as predictors. Summary scores of this type have been developed
99 and validated²⁸.

100 In addition to these pre-specified predictors, demographic data were also collected including
101 age, sex and type of stroke using the Oxfordshire Community Stroke Project Classification²⁹.

102 **Outcome measures (3, 6 and 12 months post-stroke)**

103 The primary outcome measure was a scale of difficulty caring for the arm: the Leeds Arm
104 Spasticity Impact Scale²⁵ (LASIS). This is an item bank of 12 tasks of caring for the arm
105 including aspects of washing, nail-cutting and dressing. The participant rates each relevant
106 task with degree of difficulty using a scale from 0 to 4, and scores are then averaged. Test-
107 retest reliability has been established with a minimally detected change of 0.5³⁰.

108 Secondary outcomes included passive range of movement, pain, hypertonicity, active
109 function, and skin integrity. The measures are summarised in Table 1. A protocol for
110 conducting the predictor and outcome measure assessments was developed and demonstrated
111 a good degree of inter-rater reliability, with Kappa scores of 0.82 for MMAS scores, 1.0 for

112 pain, 0.8 for LASIS and 93% agreement for measuring range of movement to within 15
113 degrees.

114 **Statistical Analysis**

115 All statistical analyses were performed using the statistical programming language R³¹.

116 Summary statistics were produced. Where the data was normally distributed, means and
117 standard deviations were used. Otherwise median and inter-quartile ranges were given.

118 Individual profile plots were constructed to visualise each participant's LASIS average across
119 follow-up points. The linear association between each continuous predictor and LASIS
120 average at 12 months was summarised using Pearson's correlation coefficient, whilst
121 descriptive statistics for LASIS average at 12 months are presented for each level of each
122 categorical predictor.

123 Multi-variable linear regression was used to identify models of predictors for LASIS average
124 at 12 months post-stroke. For brevity, this paper reports only the overall best fitting model.

125 **Sample size**

126 Sample sizes for multi-variable linear regression are based on the minimum R^2 value of
127 interest and the number of independent predictors. Whilst there were five potential predictors
128 of interest, three are categorical: pain (three categories), sensation/perception (three
129 categories) and hypertonicity (five categories), with two continuous predictors (Fugl-Meyer
130 and mood scores). After recoding categorical predictors as indicator variables, as required
131 for modelling, statistically, it may be considered that there are 10 possible explanatory
132 variables/predictors. Assuming a significance level of 10%, a sample size of 120 participants
133 was required to detect a medium effect size of 0.15 (which corresponds to R^2 value of around
134 13%) with 90% power. Based on previous studies^{32,33}, it was estimated that there would be a
135 potential drop off of 10% per measurement session. Therefore, the recruitment target was set

136 at 165 participants, with the aim of following-up at least 120 participants at the 12 months
137 post-stroke time point.

138 **Ethics**

139 The study was approved by the NRES South West Ethics Committee (Reference:
140 11/SW/0149).

141 **Results**

142 Figure 1 illustrates the process of recruitment and follow-up, including reasons for
143 participants lost to follow-up: 833 people were screened for inclusion of which 216 (26%)
144 fulfilled the inclusion criteria, and 155 gave consent or consultee assent to participate (72%
145 of those eligible). At one year 110 participants (71%) were reviewed. Of the remaining 45
146 participants, 6 declined reassessment, 33 had died and 6 were unavailable.

147 Participant demographic data at baseline and the predictor measures are summarised in Table
148 2. The average age of participants was 74.7 years, with a higher proportion of women than
149 men, and almost half of the participants had a total anterior circulation stroke (TACS). At
150 baseline, 82.6% had already developed some hypertonicity, with 17.4% exhibiting pain and
151 31.8% demonstrating impairment of sensation/perception. Outcome measures at each follow-
152 up are summarised in Table 3 and briefly summarised below.

153 **Longitudinal profiles of difficulty caring for the arm**

154 LASIS outcomes were collected from 104 participants at all time points. The mean LASIS at
155 3 and 6 months were similar (1.7 and 1.6 respectively) and by 12 months had increased to
156 2.0. However, there was a large variation in the profiles of each participant's scores, as
157 shown in the individual profile plots in Figure 2, with some showing increasing difficulty
158 over time, some decreasing difficulty and some broader variation. At the 12 month time-point
159 over half (59%) of participants reported no or little difficulty with care tasks but 12%

160 reported moderate difficulty and 29% indicated they either had a great deal of difficulty or
161 were unable to perform tasks such as washing or dressing.

162 **Longitudinal profiles of related impairments of body functions and activity limitation**

163 *Active function*

164 As anticipated, the majority of participants had not recovered active use of the arm at 12
165 months, with 73% scoring between 0 and 1 (inclusive) on the Motor activity log (MAL-14)
166 and median values remaining at 0 across time points. However, fifteen participants (14%)
167 regained some use of the arm (scoring two or more on MAL-14). The baseline characteristics
168 of those who regained some use are shown in Table 4.

169 *Hypertonicity*

170 Individual profiles of hypertonicity were very variable over the three time-points, although
171 median hypertonicity total score was 4.0 at all time points (see Table 3). Some participants
172 showed trends for increasing hypertonicity over time, some decreasing and some with no
173 discernible pattern. At one year 77% of survivors had developed some hypertonicity in at
174 least one muscle group (MMAS score at least 1), with severe hypertonicity in at least one
175 muscle group present in 25% of participants (MMAS score at least 3). The muscle groups
176 most commonly affected by severe hypertonicity were elbow and wrist flexors (affecting
177 14% of participants each), shoulder internal rotators (13%), finger flexors (10%) and shoulder
178 adductors (6%).

179 *Pain*

180 Profiles of pain were also very variable within the group of participants, although a larger
181 proportion reported pain at follow-up compared to baseline, when the vast majority (83%)
182 were pain free. At 12 months, pain in some part of the arm was reported by 65% of
183 participants.

184 *Range of movement*

185 Individual profiles of range of movement were variable over time at all the joints assessed,
186 with some participants having increasing and some decreasing range between 3 and 12
187 months. However, over the three time points, the mean range of movement, particularly at the
188 shoulder and wrist, was less than would be expected in healthy older adults³⁴. Range of
189 movement in the fingers was less reduced. Table 3 includes range for the index finger
190 proximal interphalangeal joint as an example. Other studies have defined contracture as the
191 loss of at least 30% of the available range of movement¹⁷. Using these criteria, 94% of
192 participants had developed shoulder contracture, 9% elbow contracture, 54% wrist
193 contracture and 7% finger contracture at 1 year.

194 *Skin integrity*

195 Seven participants (6%) developed macerated skin in the hand or elbow-crease at 12 months.
196 None of the participants had broken skin at any point.

197 **Predicting difficulty caring for the arm**

198 Table 5 summarises the bivariate relationships between LASIS average at 12 months and
199 each of the predictors. There was evidence of a positive relationship between the LASIS
200 average and age, hypertonicity total score and mood, although only the linear association
201 between LASIS average and age was statistically significant. We used hypertonicity total
202 scores in the best fitting model because they explained a greater percentage of variance for
203 the LASIS average than worst hypertonicity.

204 The overall best fitting linear model was derived from the five pre-specified predictors and
205 the four additional baseline variables. After the removal of three outliers, the final model was
206 fitted to data from 106 participants and included age ($p<0.001$), hypertonicity total ($p=0.002$)
207 and stroke classification (participants who have suffered from lacunar stroke (LACS)²⁹
208 compared to (a) participants who have suffered from total anterior circulation stroke
209 (TACS)²⁹ ($p=0.004$) and (b) participants who have suffered from a haemorrhage

210 (p=0.010))(see Table 6). Collectively, these three variables explained approximately one third
211 (adjusted $R^2=33\%$) of the variance in the LASIS average at 12 months. From the linear
212 regression coefficients from this final best fitting model:

- 213 • A one year increase in age at baseline increases the LASIS average at 12 months by an
214 average of 0.050 units (standard error (SE) 0.008);
- 215 • A one unit increase in hypertonicity total at baseline increases the LASIS average at 12
216 months by an average of 0.109 units (SE 0.035);
- 217 • The mean LASIS average for the group of participants who had suffered from LACS was
218 0.935 units (SE 0.314) lower than participants who had suffered from TACS and 0.962 units
219 (SE 0.367) lower than the group of participants who had suffered from a haemorrhage.

220 **Discussion**

221 This is the first longitudinal study, to our knowledge, of people with a profoundly-affected
222 arm after stroke. Whilst the sample included a high proportion of people with more severe
223 classifications of stroke this was not surprising given the target population. Many studies
224 restrict recruitment and do not involve people with severe communication or cognitive
225 limitations but we have demonstrated it was possible to include them, by supporting them
226 with enhanced communication resources or using proxies.

227 Given that participants were those with severe arm weakness at 2-4 weeks post-stroke,
228 observable patterns between impairments and activity limitation were thought to be a
229 possibility. However, this was not the case and longitudinal profiles of these factors were
230 highly individual.

231 The incidence of impairments in the arm was high when compared to studies that have
232 included general populations of stroke survivors. For example, 77% of our participants who
233 had severe weakness at baseline presented with hypertonicity at one year compared to 49% of
234 those who initially presented with milder weakness at baseline⁸. In addition, 65% of our

235 participants reported pain in the arm compared to 49% of a general population of stroke
236 survivors reporting pain in any part of the body⁹. Incidence of contracture of the shoulder and
237 wrist were also higher than that recorded in general populations of stroke survivors¹⁷,
238 although this was not the case for the elbow. It is unclear why so many of our participants
239 developed loss of range of the shoulder and wrist while the elbow and fingers remained less
240 severely affected. The shoulder is typically held in adduction and internal rotation at rest so
241 may be more vulnerable, while gravity may assist with extension of the elbow. The wrist is a
242 complex joint and contracture of the finger flexor muscle-tendon units may impact on range
243 of movement at the wrist in addition to the fingers. Differences in muscle architecture
244 surrounding connective tissue may also contribute to the variation in contracture between
245 muscles. Recent work in animal studies suggests that there is a direct relationship between
246 muscle atrophy and fibrosis. Cytokine myostatin, for example, is not only central to the
247 pathways that mediate muscle atrophy but can also activate fibroblasts and stimulate
248 fibrosis^{35,36}. Thus, differences in the weakness of individual muscles may impact on their
249 relative degree of contracture development and increase in passive stiffness^{35,37}.

250 The incidence of difficulty caring for the arm was high, as 29% of participants were either
251 unable to care for their arm or described significant difficulty. A number of predictors of
252 difficulty caring for the arm that can be assessed early after stroke were evaluated. The best
253 linear predictive model based on these included age, hypertonicity and stroke classification,
254 although these factors explained only 33% of the total variability in the LASIS average at one
255 year post-stroke. Our previous review did not identify other impairments that are likely to
256 influence longer-term outcome in caring for the arm². Previous studies have not considered
257 the use of biomarkers as predictors of outcome in this targeted group and it is possible they
258 may add to the predictive value.

259 There are a number of clinical implications of this work. Whilst recognising that people with
260 profoundly-affected arm gain little from active exercise to improve function⁴, given the high
261 incidence of pain, hypertonicity and contracture they may benefit from an educational
262 intervention to reduce the impact of these impairments, and from longer term monitoring.
263 With regard to the important risk factors identified, age and stroke classification cannot be
264 influenced in treatment after stroke but it is possible that early manifestations of hypertonicity
265 can be altered and research could explore if targeting hypertonicity early after stroke can
266 reduce the risk of difficulty caring for the arm longer term, particularly in those with other
267 risk factors.

268 **Study limitations**

269 This study has a number of limitations. Whilst every attempt was made to include measures
270 that had been validated in people with aphasia and cognitive impairment, this was not always
271 possible. The Fugl-Meyer score, in particular, has not been validated in this group. Equally
272 the anticipated sample size of 120 participants at 1 year was not achieved so adequate
273 statistical power may be lacking. Hand dominance was not considered as a predictor variable
274 and this may have an influence in self-care. Finally, no attempt was made to assess the
275 amount or content of rehabilitation that participants received so it is possible that any such
276 interventions may have impacted on outcomes. Therefore conclusions should be drawn with
277 caution.

278 **Conclusions**

279 At one year post-stroke, there was a high incidence of difficulty caring for the arm (measured
280 with LASIS) and of pain, hypertonicity and contracture. Notably, individual profiles were
281 very variable and although some pre-disposing factors have been identified, it remains
282 difficult to predict who is at greatest risk.

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378 apoptosis. *J Cell Sci*. 2012;125:3957-65.

379 Figure legends:

380 Figure 1: Flow diagram detailing recruitment and progression of participants

381 Figure 2 Individual participants LASIS scores at each time point. Each box contains 5
382 participants in their order of recruitment- where data is missing the participant was lost to
383 follow up.. (N=127 at 3 months, N= 117 at 6 months, N= 111 at 12 months).

384

385 **Table 1: Battery of predictor and outcome measures**

Predictors	Name	Scoring
Motor control	Fugl-Meyer Upper limb score ¹⁴	0-66, higher score indicates better control
Pain	Yes/no response to pain at rest and on passive movement ¹⁵	0,1 or 2, higher score indicates more pain
Hypertonicity	Modified Modified Ashworth scale ¹⁶	0-4, higher score indicates higher tone
Perception/ sensation	Find the thumb test ¹⁷	0,1 or 2, higher score indicates worse perception
Mood	Stroke Aphasic Depression Questionnaire-10 ¹⁸	0-30, higher score indicates lower mood
Outcomes		
Difficulty caring for arm	Leeds Arm Spasticity Impact Scale ¹⁹	0-4, higher score indicates more difficulty
Pain	As above	
Hypertonicity	As above	
Passive range of movement	Goniometry of shoulder flexion, abduction, external rotation; elbow flexion, extension; wrist extension, index, little finger and thumb extension at each joint ²⁰	Range measured in degrees of movement
Skin integrity	Axilla, elbow and hand, classified as dry/ intact; macerated or broken.	0,1 or 2, higher score indicates worse skin condition
Active arm	Motor activity log-14 ²¹	0-70, higher score indicates better

function		use
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388 **Table 2: Descriptive statistics of participant characteristics and potential predictors at**
 389 **baseline (n=155)**

Age	
Mean (SD)	74.7 (12.8)
Range	[38.0, 96.0]
Sex	
Female, n (%)	89 (57%)
Male, n (%)	66 (43%)
Stroke classification	
Not reported, n (%)	1 (0.6%)
Haemorrhage, n (%)	25 (16.2%)
Total anterior circulation stroke, n (%)	73 (47.4%)
Partial anterior circulation stroke, n (%)	30 (19.5%)
Lacunar stroke, n (%)	23 (14.9%)
Posterior circulation stroke, n (%)	3 (1.9%)
Fugl-Meyer upper limb score	
Median [Q1, Q3]	2.0 [2.0, 6.0]
Range	[0.0, 16.0]
Hypertonicity (worse MMAS score)	
0, n (%)	27 (17.4%)
1, n (%)	52 (33.5%)
2, n (%)	53 (34.2%)
3, n (%)	23 (14.8%)
4, n (%)	0 (0%)
Hypertonicity (total score)	

Median [Q1, Q3]	3.0 [1.0, 5.0]
Range	[0.0, 12.0]
Pain	
No pain at rest or movement, n (%)	128 (82.6%)
Pain on movement only, n (%)	21 (13.5%)
Pain at rest & on movement, n (%)	6 (3.9%)
Mood (SADQ-10)	
Mean (SD)	8.2 (4.5)
Range	[0.0, 23.0]
Sensation/Perception (Find the Thumb test)	
Not reported, n (%)	1 (0.6%)
Able to find affected thumb, n (%)	105 (68.2%)
Able to find affected arm only, n (%)	19 (12.3%)
Unable to find affected arm, n (%)	30 (19.5%)

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392 **Table 3: Descriptive statistics of outcome measures at each follow-up time point**

	3 months n=120	6 months n=113	12 months n=110
LASIS average			
Mean (SD)	1.7 (1.0)	1.6 (1.1)	2.0 (1.3)
Range	[0.08, 4.00]	[0.00, 4.00]	[0.00, 4.00]
Hypertonicity (worse score)			
Not reported, n (%)	2 (1.7%)	4 (3.5%)	1 (0.9%)
0, n (%)	16 (13.6%)	19 (17.4%)	24 (22%)
1, n (%)	31 (26.3%)	33 (30.3%)	29 (26.4%)
2, n (%)	52 (44.1%)	32 (29.4%)	29 (26.4%)
3, n (%)	19 (16.1%)	22 (20.2%)	24 (22%)
4, n (%)	0 (0%)	3 (2.8%)	3 (2.7%)
Hypertonicity (total score)			
Median [Q1, Q3]	4.0 [2.0, 8.0]	4.0 [1.0, 9.0]	4.0 [1.0, 7.8]
Range	[0.0, 15.0]	[0.0, 16.0]	[0.0, 15.0]
Pain			
No pain at rest or movement, n (%)	32 (26.7%)	31 (27.4%)	38 (34.5%)
Pain on movement only, n (%)	60 (50%)	56 (49.6%)	56 (50.9%)
Pain at rest & on movement, n (%)	28 (23.3%)	26 (23.0%)	16 (14.5%)
Passive range shoulder abduction			
Mean (SD)	76.8° (24.4°)	74.3° (22.6°)	79.9° (28.8°)
Range	[25.0°, 170.0°]	[10.0°, 160.0°]	[20.0°, 180.0°]
Passive range shoulder external rotation			
	22.8° (24.8°)	24.8° (22.7°)	25.0° (27.4°)

Mean (SD)	[-80°, 65.0°]	[-70.0°, 75.0°]	[-60.0°, 90.0°]
Range			
Passive range elbow extension			
Mean (SD)	165.0° (20.0°)	165.0° (20.8°)	164.8° (21.0°)
Range	[100.0°, 180.0°]	[90.0°, 180.0°]	[100.0°, 180.0°]
Passive range wrist extension			
Mean (SD)	38.6° (21.6°)	37.7 (26.7°)	43.7° (26.6°)
Range	[-60.0°, 90.0°]	[-50.0°, 80.0°]	[-60.0°, 80.0°]
Passive range index PIP extension			
Mean (SD)	175.3° (11.5°)	172.1° (17.3°)	174.2° (16.3°)
Range	[100.0°, 180.0°]	[90.0°, 180.0°]	[100.0°, 180.0°]
Skin integrity			
Not reported, n (%)	0	2 (1.8%)	0
Dry intact, n (%)	118 (98.4%)	105 (92.9%)	103 (93.6%)
Macerated, n (%)	2 (1.6%)	6 (5.3%)	7 (6.4%)
Broken, n (%)	0	0	0
Active function (MAL-14)			
Median [Q1, Q3]	0.0 [0.0, 0.8]	0.1 [0.0, 1.0]	0 [0.0, 1.2]
Range	[0, 4.21]	[0.0, 5.00]	[0.0, 4.38]

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Table 4: Descriptive statistics at baseline by functionality at 12 months (n=109)

	Participants with MAL less than two at 12 months (n=94)	Participants with MAL two or greater at 12 months (n=15)
Age		
Mean (SD)	72.7 (12.9)	68.1 (13.7)
Range	[38.0, 95.0]	[40.0, 96.0]
Gender		
Female, n (%)	49 (52%)	9 (60%)
Male, n (%)	45 (48%)	6 (40%)
Stroke classification		
Not reported, n (%)	0 (0%)	0 (0%)
Haemorrhage, n (%)	19 (20.2%)	1 (6.7%)
Total anterior circulation stroke, n (%)	45 (47.9%)	6 (40.0%)
Partial anterior circulation stroke, n (%)	16 (17.0%)	4 (26.7%)
Lacunar stroke, n (%)	12 (12.8%)	4 (26.7%)
Posterior circulation stroke, n (%)	2 (2.1%)	0 (0%)
Fugl-Meyer upper limb scores		
Median [Q1, Q3]	2.0 [2.0, 5.0]	5.0 [4.0, 10.5]
Range	[0.0, 15.0]	[2.0, 15.0]
Hypertonicity (worse MMAS score)		
0, n (%)	16 (17.0%)	3 (20.0%)
1, n (%)	26 (27.7%)	6 (40.0%)
2, n (%)	38 (40.4%)	5 (33.3%)
3, n (%)	14 (14.9%)	1 (6.7%)
4, n (%)	0 (0%)	0 (0%)
Hypertonicity (total score)		
Median [Q1, Q3]	3.0 [1.0, 5.8]	2.0 [1.0, 3.0]
Range	[0.0, 12.0]	[0.0, 7.0]

Pain		
No pain at rest or movement, n (%)	77 (81.9%)	13 (86.7%)
Pain on movement only, n (%)	12 (12.8%)	1 (6.7%)
Pain at rest & on movement, n (%)	5 (5.3%)	1 (6.7%)
Mood (SADQ-10)		
Mean (SD)	8.3 (4.8)	6.1 (4.2)
Range	[0.0, 23.0]	[1.0, 16.0]
Sensation/Perception (Find the Thumb test)		
Not reported, n (%)	1 (1.1%)	0 (0%)
Able to find affected thumb, n (%)	68 (72.3%)	11 (73.3%)
Able to find affected arm only, n (%)	10 (10.6%)	2 (13.3%)
Unable to find affected arm, n (%)	15 (16.0%)	2 (13.3%)

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399 **Table 5: Summary statistics of bivariate relationships between predictors and LASIS**
 400 **average at 12 months**

	Mean (SD) [range]	Correlation Coefficient (95% CI)
Age	-	0.39 [0.22, 0.54]
Sex		-
Female	2.2 (1.4) [0.00, 4.00]	
Male	1.7 (1.3) [0.00, 4.00]	
Stroke classification		-
Lacunar stroke (LACS)	1.4 (0.9) [0.00, 3.82]	
Partial anterior circulation stroke (PACS)	1.5 (1.3) [0.09, 4.00]	
Posterior circulation stroke (POCS)	0.9 (0.7) [0.42, 1.44]	
Total anterior circulation stroke (TACS)	2.3 (1.4) [0.00, 4.00]	
Haemorrhage	2.2 (1.3) [0.45, 4.00]	
Fugl-Meyer upper limb score	-	-0.18 [-0.36, 0.01]
Hypertonicity (worse MMAS score)		-
0	1.7 (1.3) [0.27, 4.00]	
1	1.9 (1.4) [0.00, 4.00]	
2	1.9 (1.3) [0.00, 4.00]	
3	2.8 (1.3) [0.55, 4.00]	
4	NA	
Hypertonicity (total score)	-	0.19 [0.00, 0.37]
Pain		-
No pain at rest or movement	1.9 (1.4) [0.00, 4.00]	
Pain on movement only	2.6 (1.3) [0.10, 4.00]	

Pain at rest & on movement	1.7 (0.6) [0.80, 2.27]	
Mood (SADQ-10)	-	0.19 [0.00, 0.36]
Sensation/Perception (Find the Thumb)		-
Able to find affected thumb	1.8 (1.3) [0.00, 4.00]	
Able to find affected arm only	2.3 (1.1) [0.25, 4.00]	
Unable to find affected arm	2.3 (1.5) [0.36, 4.00]	

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403 **Table 6: Regression statistics for the overall best fitting model for LASIS average at 12**
404 **months**

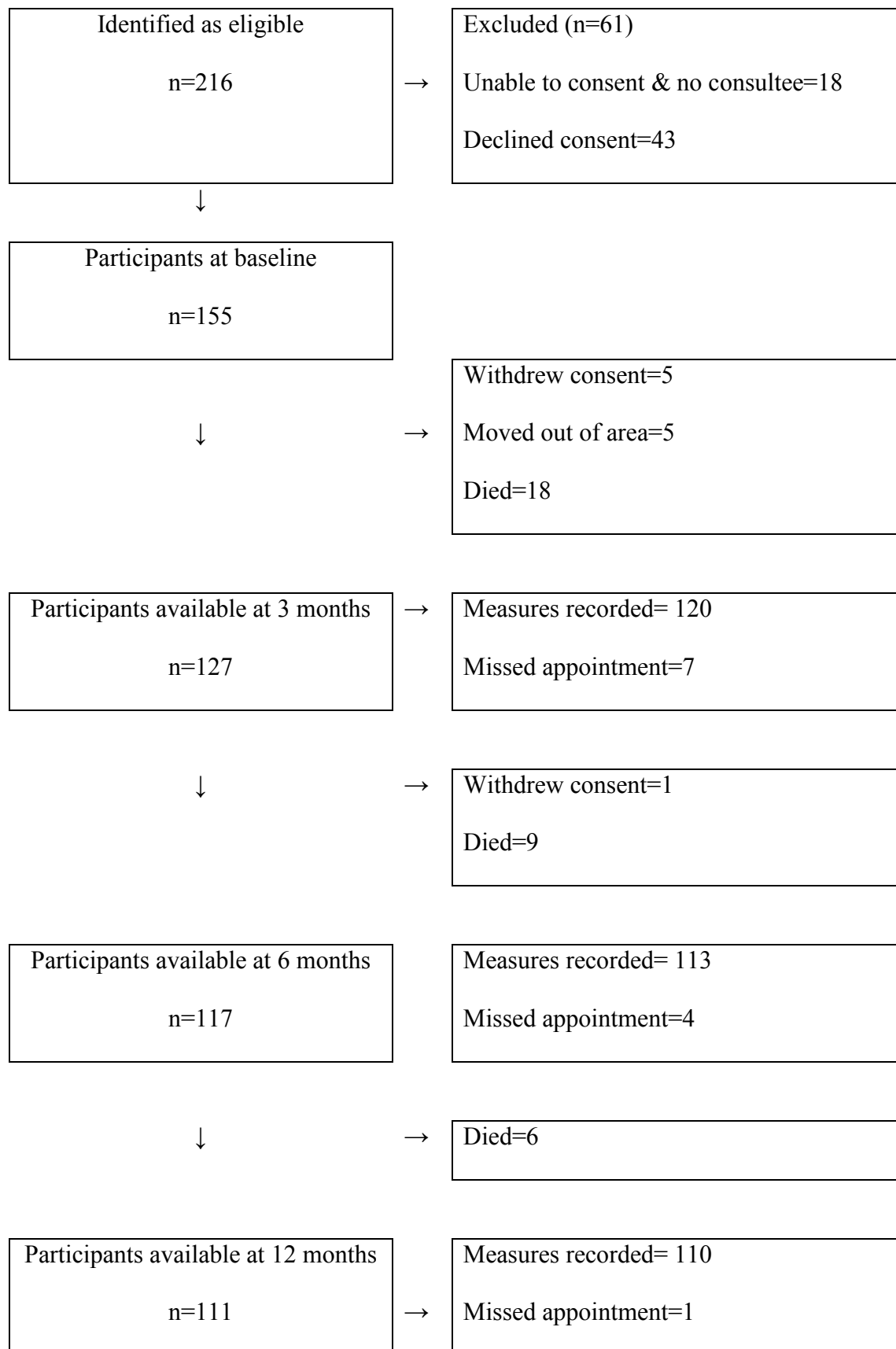
	Coefficient	95% confidence interval	p-value
Intercept	-2.658	[-4.028, -1.288]	<0.001
Age	0.050	[0.034, 0.066]	<0.001
Hypertonicity total	0.109	[0.040, 0.178]	0.002
stroke class POCS	-0.200	[-1.809, 1.409]	0.808
stroke class PACS	0.121	[-0.608, 0.850]	0.744
stroke class TACS	0.935	[0.320, 1.550]	0.004
stroke class HAEM	0.962	[0.243, 1.681]	0.010
Residual standard error: 1.091 on 99 DF Multiple R-squared: 0.37 Adjusted R-squared: 0.33 F-statistic: 9.6 on 6 and 99 DF, p-value: <0.001			

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406 LACS (Lacunar stroke) is the baseline level for stroke classification.

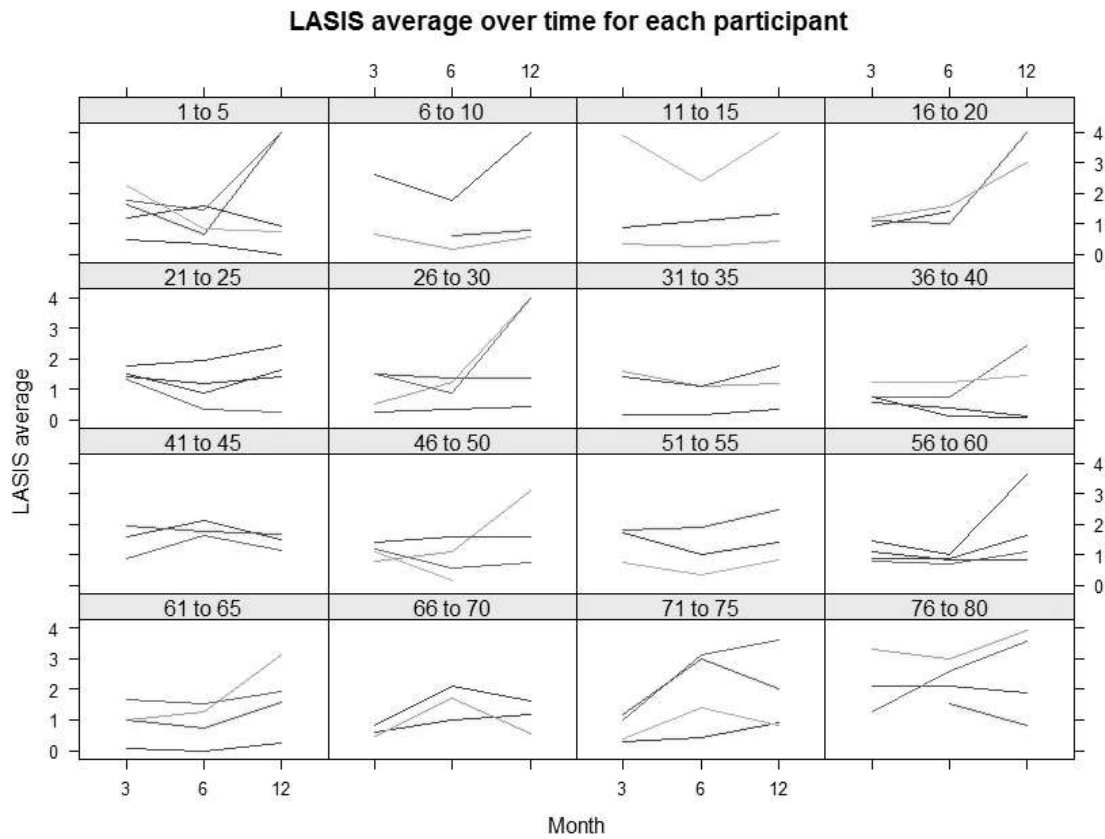
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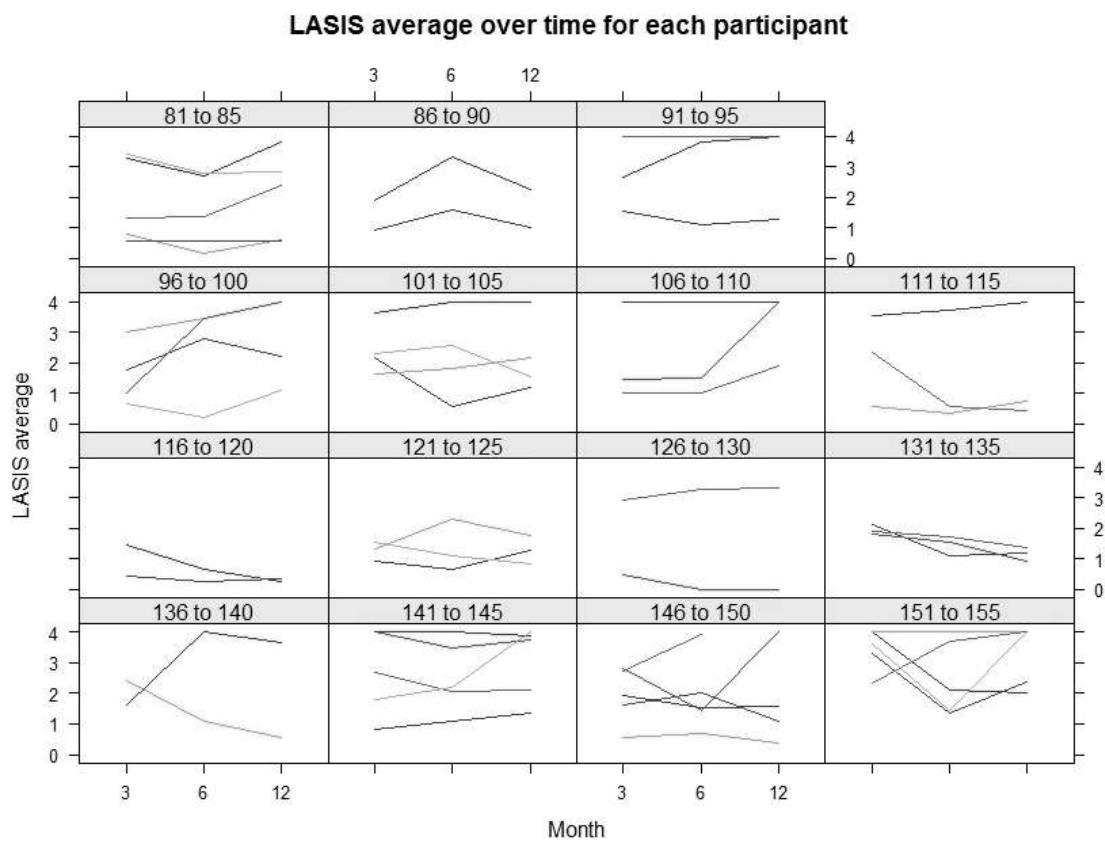


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412 Figure 2



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