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Rehabilitation of the upper arm early after stroke: video games versus conventional rehabilitation. A randomized controlled trial

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1 **Rehabilitation of the upper arm early after stroke: video games versus conventional**
2 **rehabilitation. A randomized controlled trial**

3

4

5 **Abstract**

6 **Background.** Few rehabilitation methods have proven their efficacy in increasing sensori-
7 motor recovery and/or function of the upper limb (UL) after stroke. Video games (VGs) are
8 promising tools in this indication.

9 **Objective.** To compare UL rehabilitation by using VGs and conventional rehabilitation (CR)
10 in patients with sub-acute stroke.

11 **Design.** Single-blind, multicentric trial, with central randomization and stratification by
12 center.

13 **Setting.** Physical and rehabilitation medicine departments of 2 university hospitals.

14 **Participants.** Adults within 3 months after a first vascular cerebral accident, with UL Fugl
15 Meyer Score (UL-FMS) <30/66 and without major cognitive impairment.

16 **Intervention.** A 45-min additional session of conventional occupational therapy (OT) or a
17 VG-based OT session as add-on therapy to usual rehabilitation programs, 5 days/week for 6
18 weeks.

19 **Main Outcome Measures.** Primary outcome: UL-FMS. Secondary outcome: Box and Block
20 Test (BBT), Wolf Motor Function test (WMFT), Motor Activity Log (MAL), Barthel Index
21 and quality of life (SF-36).

22 **Results.** We included 51 patients (20 women) at a mean (SD) of 27.2 (19.4) days post-stroke
23 (mean age 58 years [range 24-83]), 26 in the CR group and 25 in the VG group (23 in each
24 group at 6-month follow-up). The mean duration of the additional rehabilitation session was
25 similar in both groups: 29.3 (4.3) vs 28.0 (4.4) min in CR and VG groups. Shoulder pain

26 occurred in 4 patients in the VG group versus 7 in the CR group. At day 45, gain in UL-FMS
27 did not significantly differ between the groups (CR mean 17.8 [14.6] vs VG 24.1 [14.8];
28 $p=0.10$), whereas gain in BBT was doubled in the VG group (CR 7.4 [12.2] vs VG 15.7
29 [16.3]; $p=0.02$). At 6-month follow-up, the study was inconclusive about between-group
30 differences in UL-FMS, BBT and other criteria. Post-hoc analysis showed that gains in UL-
31 FMS or BBT were significantly higher in the VG than CR group for patients included within
32 30 days post-stroke.

33 **Conclusion.** In general, we cannot conclude that video gaming and conventional OT led to
34 different long-term sensorimotor recovery of the UL after sub-acute stroke. However, when
35 applied within the first month after stroke, video gaming was more efficient than conventional
36 rehabilitation on both sensorimotor recovery and gross grasping function.

37

38 Trial registration. [ClinicalTrials.gov \(NCT01554449\)](https://clinicaltrials.gov/ct2/show/study/NCT01554449)

39

40 **Key Words.** stroke; upper limb; video games; rehabilitation

41

42 **Introduction**

43 More than 75% of stroke patients have remaining upper-limb (UL) impairment at the chronic
44 stage (1), which renders the rehabilitation of the UL after stroke challenging. We suspect that
45 UL rehabilitation must be intensive, repetitive and task-oriented (2). Despite an increasing
46 interest during the past 2 decades, few rehabilitation methods have proven their efficacy in
47 increasing sensori-motor recovery and/or function of the UL after stroke (3).

48 Video games (VGs) are promising tools in this indication. However, rehabilitative
49 VGs include different kind of devices, so comparison between studies is difficult. The first
50 studies were mainly based on the use of “commercially available VGs”, whereas more

51 recently, “specifically designed VGs” for the rehabilitation of stroke patients have been
52 developed and tested. Subsequently, non-immersive games and immersive ones (virtual
53 reality) have been tested, despite different potential efficacy (4). The technical devices
54 supporting the games widely varied in the literature, from historical dedicated consoles such
55 as Nintendo (Wii[®]), Sony (Playstation[®]) or Microsoft (Xbox[®]), to personal computers,
56 numeric tablets or customized stations. The playing interface also widely varied among
57 surveys: mouse or joystick, graphic tablet, optic motion capture sensors such as Kinect[®],
58 mechanical support such as Armeo[®], or motorized interface such as InMotion Robot or
59 Reaplan[®] bridging the gap with rehabilitation robotics.

60 The rationale of UL rehabilitation with VGs first relies on the opportunity to increase
61 repetition, duration, and intensity of exercises (5). VGs offer goal-oriented tasks and
62 repetitive exercises in an enriched and interactive environment, highly suspected to increase
63 recovery after neurological lesion. As compared with “commercially available VGs”,
64 “specifically-designed VGs” enable therapists to adapt gameplay and game design (6)(7) as
65 well as modulate the sensory feedback provided to the patient to enhance learning, through
66 the manipulation of visual, auditory, haptic/sensitive feedback (8) (9). Whatever their
67 technical specificities, VGs are known to increase motivation and engagement of patients in
68 their rehabilitation program, thereby encouraging a higher number of repetitions and longer
69 periods of training (10). Specific effects of VG rehabilitation on neuroplasticity have been
70 poorly investigated and are still debated (11).

71 When we started the study in 2010, the literature was promising although inconclusive
72 about the efficacy of VGs when applied to sensorimotor rehabilitation of the UL after stroke.
73 The promising results from initial studies (12) (3) (4) (12) unfortunately relied on a large
74 amount of open-labelled and uncontrolled trials, reporting results of short trials on a small
75 sample of patients, mostly in the chronic phase after stroke. Finally, if VGs were suspected to

76 have some minor effects on body structure/function and activity, all authors agreed on the
77 lack of a demonstrated effect on participation outcomes or quality of life.

78 From these studies, we decided to conduct a randomised controlled trial (RCT) comparing
79 VG-based occupational therapy (OT) and the same amount of conventional OT based on 4
80 hypotheses:

81 - Main hypothesis: VG-based OT is more efficient for UL impairment and function
82 rehabilitation in patients with sub-acute stroke because VG more easily provides high-
83 intensity and repetitive task-specific practice.

84 - Secondary hypotheses:

85 ○ VG-based OT is more efficient than conventional OT in terms of general
86 function and quality of life.

87 ○ VG-based OT is safe and feasible.

88 ○ VG-based OT is more efficient than conventional OT when applied early after
89 stroke because the increased activity due to VGs aligns with the critical
90 window for neurological repair.

91 **Methods**

92 This study was a multicentric single-blind RCT. It was approved by the regional ethics
93 committee (IDRCB: 2010-A00596-33) and was registered in ClinicalTrials.gov
94 (NCT01554449). All patients gave their written informed consent before enrollment and
95 procedures conformed with the Declaration of Helsinki.

96 ***Participants***

97 Patients were recruited from physical and rehabilitation medicine departments of 2 university
98 hospitals between January 2011 and June 2016. Inclusion criteria were age > 18 years, first
99 ischemic or hemorrhagic stroke, time since stroke < 3 months, and UL Fugl Meyer Score
100 (UL-FMS) <30/66. Non-inclusion criteria were visual troubles limiting computer use,

101 cognitive impairment with Mini Mental Scale score < 22 (13), hemispatial neglect with
102 Catherine Bergego Scale score > 14/20 (14), aphasia with Boston Aphasia Quotient < 4/5
103 (15), and patient's or legal representative's refusal to participate in the study. A total of 51
104 patients with stroke were included in this trial, plus 12 healthy controls from a functional MRI
105 analysis that was previously published (16).

106

107 ***Study design***

108 The study was a single-blind RCT. Patients were randomly assigned at a 1:1 ratio to receive
109 30/45 min of additional conventional OT rehabilitation (CR) or non-immersive video gaming
110 rehabilitation (VG) under OT supervision. The randomization sequence was centralized
111 (online using Ennov Clinical software) and computed in permuted blocks by the statistician
112 who used SAS (SAS Institute, Cary, NC) with stratification by center.

113

114 ***Rehabilitation program***

115 All patients underwent the same CR protocol, whose content was harmonized between the 2
116 participating centers (see Table 2). This protocol included 2 sessions/day of 30 min each of
117 physiotherapy focusing on passive and active mobilization of the limbs, balance exercises,
118 muscular strengthening, and walking. It also included a 30- to 60-min OT session to improve
119 the patient's autonomy in activities of daily living, including prehension and grasping
120 exercises. Patients with swallowing and/or language disorders received up to 2 additional
121 sessions of speech therapy. The total duration of this "baseline" rehabilitation program ranged
122 from 90 to 180 min/day depending on patient's needs.

123 All patients received an additional session of rehabilitation under the close supervision of
124 an occupational therapist, 5 days/week, for 6 consecutive weeks. The duration of this

125 additional session was adapted depending on the patient's weakness and tolerance: at least 15
126 min/day (usually at the onset of the rehabilitation period), at most 45 min/day.

127 - In the CR group, patients received conventional OT focusing on intensive, repetitive,
128 and task-oriented movements of the affected UL. When patients presented grasping
129 capacities, various prehension exercises were added. Therapists were not allowed to
130 use VGs, electromechanical or robotic devices in this group.

131 - In the VG group, patients received OT based on video games played with the affected
132 UL. During the first half of the session, patients played VGs that were configurable by
133 the therapist in terms of speed, difficulty, distance and size of the targets, design of the
134 scenes, number of distracters, and visualization of the scoring of the performance. One
135 game automatically adapted target distance depending on the patients' reaching zone
136 (7) and was used at least 10 min/day. During the second half of the session, patients
137 played online games that were previously selected by a group of therapists with the
138 following criteria: slow and easy-to-play, as few as possible distracters, requiring
139 repetitive movements of the mouse, and low cognitive demand. These 30 games were
140 grouped on a dedicated server that was the same at both experimental sites. The
141 objective was to provide a large number of games to vary the content of the sessions
142 and to adapt the games to each patient's motor and cognitive impairment.

143 ***Outcomes***

144 Patients were assessed before the beginning of the rehabilitation protocol (V0) and at the end
145 of the program, between days 45 and 60 (V1). They had a follow-up assessment at 6 months
146 (V2) after inclusion. At each visit, the same trained physician conducted a detailed clinical
147 examination and the tests described below. This physician was blinded to the randomization
148 group of the patient and was not involved in administering study interventions. Adverse
149 events were collected throughout the study.

150 The UL Fugl Meyer Score (UL-FMS, maximal score 66) was used as a primary
151 endpoint to assess UL sensorimotor impairment (17). In the sub-acute stroke stage, a
152 difference of 9/10 points on the UL-FMS is considered a Minimal Clinically Significant
153 Difference (18).

154 The Box and Block Test (BBT) was used to assess overall gross grasping capacities
155 (19). The BBT consists in grasping, moving and releasing a maximum number of 2.5-cm
156 wooden cubes from one box to another in 1 min. Healthy persons move from 65 to 75
157 blocks/min, depending on the age. The Minimal Clinically Significant Difference of the BBT
158 in patients with sub-acute stroke is unknown (20), but its Minimally Detectable Change is
159 known to be 5.5 blocks/min (19).

160 We used the Wolf Motor Function Test (WMFT) to measure motor function of the UL
161 (20) and the Motor Activity Log (MAL) to estimate perceived upper-extremity participation
162 restrictions (21). Quality of life was assessed by the Medical Outcomes Study Short Form 36
163 (SF-36) (22) and independence for activities in daily living by the Barthel Index on a 100-
164 point score (23).

165

166 *Statistics*

167 Sample size estimation

168 According to the team's ongoing work at the time of the protocol development, we expected a
169 difference of gain in UL-FMS total score of 10 points \pm 10 between the 2 groups. With a 2-
170 sided α error of 0.05 and a β error of 0.10, the sample size was estimated at 22 patients per
171 group. Considering a dropout rate of about 8%, we needed to include a total of 50 patients.

172

173 Statistical analysis

174 Categorical variables are expressed as number (%) and quantitative variables as mean (SD) or
175 median [Q1-Q3]. Shapiro-Wilk's test was used to test the normality of continuous variables.
176 The primary outcome was the gain in total UL-FMS at V1 from baseline (V1-V0). Secondary
177 outcomes were gains at V1 or V2 (V2-V0) in distal or proximal UL-FMS, BBT score of the
178 affected hand, WMFT, MAL, Barthel Index and SF-36. In the main analysis, differences in
179 gains at V1 and V2 between the 2 groups were analyzed by Student *t* test or Wilcoxon test,
180 according to the normality of data distribution. Analyses were computed on an intention-to-
181 treat basis.

182 Linear mixed models for repeated measures were performed to analyze the association
183 between intervention group and evolution in UL-FMS and BBT score of the affected hand
184 over the 6-month follow-up. Because the post-stroke delay was strongly associated with the
185 variation of scores and cerebral plasticity is highest during the first month after stroke (24),
186 models were performed in the 2 sub-groups classified by the post-stroke delay (time to first
187 protocol session ≤ 30 and > 30 days). Time, Group (video game vs control) and Group x Time
188 interactions (to test the differences in changes over time between groups) were modeled as
189 fixed effects with the test scores as dependent variables. Participants were considered a
190 random effect.

191 $P < 0.05$ was considered statistically significant. The statistical analysis was performed
192 at the epidemiology and clinical research unit of Montpellier University Hospital with SAS
193 Enterprise Guide v7.13 (SAS Institute, Cary, NC).

194

195 **Results**

196 Recruitment of patients lasted from October 2012 to February 2016 after reaching the number
197 of subjects to treat, and the study follow-up ended in August 2016. The total study sample
198 was 51, with 25 patients in the VG group and 26 controls (Fig. 1). All participants completed

199 the program and the first assessment (week 6, primary outcome). At the second assessment
200 (month 6), 46 patients were evaluated, 23 in each group.

201 Mean age of the sample was 58 years; 61% were male. Baseline characteristics were
202 comparable between groups (Table 1). Upon inclusion in the study, the median [Q1–Q3] time
203 since stroke was 21 [12–36] days, with 61% of the sample included before 30 days. The
204 number and duration of therapy sessions were similar in both groups (Table 2).

205 The primary outcome, gain in total UL-FMS at week 6, did not significantly differ
206 between the 2 groups (Fig. 2): 17.8 (14.6) versus 24.1 (14.8) for control and VG groups
207 ($p=0.10$). Gains at week 6 were significantly higher in the VG than control group for distal
208 UL-FMS ($p=0.02$) and BBT ($p=0.02$) but not proximal UL-FMS, MAL items, WMFT, SF-36,
209 or Barthel Index (Table 3, left columns). At 6 months, BBT showed higher gains in the VG
210 than control group but not significantly ($p=0.055$); other results were not significantly
211 different between the 2 groups (Table 3, right columns).

212 In post-hoc multivariate analyses, we found a significant increase in total UL-FMS
213 over time ($p<0.0001$), comparable in both groups, and strongly explained by post-stroke delay
214 ($p=0.009$). As a consequence, we ran separate analyses as a function of time since stroke with
215 a threshold at 30 days. If VG-based OT rehabilitation started within 30 days after stroke, VG
216 therapy was more efficient for total UL-FMS (Time x Intervention interaction, $p=0.02$), but
217 not if rehabilitation started after 30 days ($p=0.85$). Similar results were found for distal UL-
218 FMS and BBT scores (Fig. 3).

219 Eight participants exhibited adverse events related to the study: 5 in the control group
220 and 3 in the VG group. These events included shoulder pain (6), complex regional pain
221 syndrome (1), and worsening of spasticity (1).

222

223 **Discussion**

224 Our study shows that non-immersive VG-based rehabilitation of the UL is not superior to a
225 same amount of conventional OT in patients with subacute stroke, when considering UL
226 sensori-motor impairment (UL-FMS), UL function (BBT and WMFT), activities of daily
227 living (Barthel Index) and quality of life (SF-36). These results are consistent with the recent
228 literature in the field.

229 Some early studies conducted in the 2000s and reviewed by Saposnik and Levin (25)
230 suggested a weak positive effect of VG-based rehabilitation as compared with conventional
231 rehabilitation programs. Despite encouraging recent reviews partially confirming these
232 positive results (26)(27), the evidence of efficiency is very low, and no conclusions can be
233 drawn from most of these trials because of methodological weakness or very low samples.

234 Over the last 5 years, an increasing number of negative RCTs have been published that
235 also temper the initial enthusiasm. These studies failed to demonstrate the positive effect of
236 VG on impairment and function, especially when they compared the same dose of
237 rehabilitation with conventional methods. Choi et al. (28) published a small but convincing
238 negative RCT of 20 subacute stroke patients, comparing Nintendo Wii® rehabilitation and the
239 same duration of conventional rehabilitation. Prange et al. (29) compared the effect of weight-
240 supported arm training combined with computerized exercises with dose-matched
241 conventional training of the UL in 70 subacute patients. Both failed to demonstrate any
242 difference between the 2 methods on arm function and activity. Adie et al. (30) included 209
243 subacute stroke outpatients in a multicentric RCT. The participants received daily UL
244 rehabilitation with the Nintendo Wii® system or conventional arm exercises. VG
245 rehabilitation was not superior to arm exercises on UL function; also, Wii®, despite being well
246 tolerated, was more expensive than arm exercises at home. Saposnik et al. (31) also drew the
247 same conclusion from the Evrest Study comparing 2 groups of 141 patients with subacute

248 stroke receiving a 2-week program of intensive rehabilitation with commercial non-immersive
249 Nintendo Wii® games compared to the same duration of recreational OT. Those results were
250 synthesized by Laver et al. (32) in a recent Cochrane review including 72 trials and 2470
251 patients studying the effect of virtual reality (VR) and/or VG-based rehabilitation. Among the
252 72 trials, the authors selected 22 (1038 patients) comparing VR/VG rehabilitation to the same
253 dose of conventional rehabilitation of the UL. They concluded that “the use of VR and
254 interactive video gaming is not more beneficial than conventional approaches in improving
255 upper limb function”.

256 The main results of our study confirm this conclusion, but it is also possible that the
257 effect of VG was undermined in our study. Our protocol ensured that no lack of chances
258 occurred for patients, so the selective impact of VG was more difficult to reveal because of
259 the multiplicity of common interventions (i.e., when comparing physical therapy + OT + OT
260 vs physical therapy + OT + VG) and also because the additional OT session in the control
261 group included high repetitive and intensive training of the UL. To evaluate whether VR/VG-
262 based rehabilitation has a positive effect, while being more efficient on therapist’s
263 investment/costs, it should be studied against “no rehabilitation” or “self-rehabilitation” and
264 not against conventional OT or physical therapy. Kwon et al. (33) studied the effect of VR
265 when added to conventional rehabilitation in a group of 26 patients with acute stroke versus
266 conventional rehabilitation alone and showed a significant effect on motor function (UL-FMS
267 and Function Motor test) in the experimental group. Sin and Lee (34), in a randomized
268 controlled study, compared 2 groups of 40 patients total with chronic stroke, one receiving an
269 additional period of VR rehabilitation of the UL with the Xbox Kinect system®. At the end of
270 the intervention and at follow-up, the 2 groups significantly differed in range of motion, UL-
271 FMS and BBT scores. On the basis of 10 trials (210 participants), Laver et al. (32) concluded
272 that “VR may be beneficial in improving upper limb function and activities of daily function

273 when used as an adjunct to usual care”, which suggests that VR/VG as a rehabilitation tool
274 mainly act by increasing the overall therapy time. From this hypothesis, video games could be
275 particularly interesting after discharge from the rehabilitation centre, as a “tele-rehabilitation
276 home-based device”, when human resources supporting rehabilitation in the chronic phase are
277 scarce.

278 One could also suspect that the selection of patients with severe stroke, with UL-FMS
279 < 30/66, influenced the results of our study. However, when considering less impaired
280 patients only (UL-FMS > 20), we did not find any difference between the 2 groups.
281 Furthermore, our sample was similar to comparable studies.

282 Finally, the small sample size, as well as the discrepancy between the non-significant
283 results on the main judgment criterion (UL-FMS) and the significant positive functional
284 results (BBT), are questioned. These 2 points may suggest an underpowered study. Indeed,
285 the large variability in primary endpoint (UL-FMS) has been underestimated and may have
286 hidden positive results. The choice of the UL FMS as a primary endpoint is also questionable
287 because its metrological properties are controversial, and a more functional criterion could
288 have been more interesting. Not assessing patients’ and therapists’ opinion on the games
289 prevents us from understanding “external causes” of this negative result.

290 Nevertheless, a potentially important result of this trial is that early VG-based
291 rehabilitation of the UL could be more efficient for UL motor recovery and function than
292 conventional rehabilitation, when started within the first 30 days after stroke (Fig. 3b). Of
293 course, some caution is required because this conclusion relies on a post-hoc analysis of the
294 main results. Still, because the VG and control groups had similar characteristics at inclusion
295 (Table 1) and different characteristics after early treatment, the VG therapy likely favorably
296 acts on cerebral plasticity and recovery but only when applied early (Fig. 3).

297 The question of the usefulness of very early (during the first week) or early (during the
298 first month) rehabilitation after stroke has aroused the interest of many authors recently.
299 Because cerebral plasticity is known to be crucial during the first month after stroke (24),
300 some authors suggest that early training, at the right dose and with the correct methods, will
301 aid recovery when applied during this period (35). However, despite an increasing interest, we
302 have limited evidence to guide practice and draw recommendations. We know that very acute
303 rehabilitation may be deleterious immediately (<24 hr) after stroke (36)(37). Some studies
304 support the idea that very early rehabilitation is not useful. For example, the AMOBES Study
305 (38) of an additional 45 min of early intensive physical therapy initiated during the first 72 hr
306 after stroke included 104 patients and reported negative results on motor function as
307 compared with conventional rehabilitation. However, some other studies demonstrated that
308 early mobility training and mobilization appear well tolerated and concluded few reasons to
309 delay initiating some rehabilitation within the first month (24).

310 So far, the question of a specific effect of VG on motor recovery after stroke remains
311 unsolved. In our study, patients received 2 sessions of UL rehabilitation each day,
312 corresponding to a daily dose of 60 to 120 min of exercises. This dose of UL rehabilitation is
313 very high and exceeds the dose usually used in this population in other studies. Indeed, in
314 2016, Serrada et al. (39) performed a systematic review of 94 trials including 3236
315 participants and concluded that only 7.9 min/day was devoted to UL therapy within the 4
316 weeks of post-stroke rehabilitation. Because our 2 groups of patients were dose-matched, this
317 finding precludes that the increase in motor function and grasping capabilities is linked to the
318 duration and intensity of rehabilitation. We conclude that our study strongly suggests that VG
319 is responsible for this increase of recovery in the early-treated group, which has not been
320 previously studied (32).

321 Because the answer to each rehabilitation protocol and thus the recovery potential
322 could have differed across groups due to differences in the location of brain lesions, the
323 variability of lesions should be considered. As shown in Table 1, most of our patients had
324 subcortical or total lesions, with no difference between the 2 groups. Except for the fact that
325 damage to the cortico-spinal tract is highly suspected to be associated with reduced UL motor
326 recovery (40), the current literature on the link between lesion characteristics in structural
327 imaging and recovery potential is inconclusive (41). As suggested by some papers in the field,
328 functional MRI as well as diffusion tensor imaging to “structural imaging” should be added in
329 future studies.

330 Because some characteristics of the VG used in our trial might have influenced the
331 outcome, this work could be continued with an early inclusion and with improved VG. First,
332 even if our patients played a large part of each session with “specifically-designed VGs” that
333 are highly suspected to be more efficient than commercially available ones, they played only
334 10 min with the most interesting game offering automatic adjustments to their performance.
335 Such “adaptive VGs” have been poorly investigated in the medical literature (42)(7) and their
336 specific interest is seriously suspected, even if not formally demonstrated yet (43).

337 Second, our VGs were not immersive ones, which could have impeded their
338 efficiency. Indeed, and even if still debated, the immersive properties of VGs are suspected to
339 provide more beneficial results on motor recovery (43).

340 Third, our games did not include manipulation of visual or auditory feedback provided
341 to patients to favour after-effect and learning, for example by increasing errors in trajectories
342 formation. In the same way, based on mirror neurons theories (44), providing patients with a
343 corrected visual feedback through an avatar could also increase the therapeutic effects of the
344 games.

345 Fourth, we found a significantly greater improvement in distal UL-FMS with a
346 functional translation in the BBT at 6 weeks in the VG group (Table 3). This result is
347 somewhat surprising because all the games in the VG group mainly involved shoulder-elbow
348 movements. However, half of the games were played with a mouse-like device, so distal
349 extensors and pinch muscles probably played an important role in controlling the movement
350 in the game space. Thus, patients in the VG group may have learned to use the distal joints
351 while playing VGs with a mouse-like playing interface, so that VG therapy reduced under-use
352 of the distal joints.

353

354 **Conclusion**

355 Considering all patients, our study is inconclusive about the hypothesized superiority of VG-
356 based rehabilitation of the UL as compared with the same duration of conventional OT in
357 patients with subacute stroke. This finding suggests that in general, VG-based methods should
358 not replace conventional methods but can be considered additional therapy, thus increasing
359 the amount of rehabilitation.

360 Considering only patients for whom rehabilitation started within 30 days post-stroke,
361 our study concludes that VG-based UL rehabilitation is more effective than conventional
362 rehabilitation. Although the latter result needs to be confirmed by further studies, this finding
363 suggests that early after stroke, VG therapy may be effective and could be systematically
364 considered a complement to conventional methods.

365

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371

372 **Conflict of Interest.** None declared.

373

374 **Legends**

375

376 **Figure 1.** Flow of participants in the study.

377

378 **Figure 2.** Neurological and functional changes in the video game and control groups.

379

380 **Figure 3.** Neurological and functional changes by intervention delay.

381

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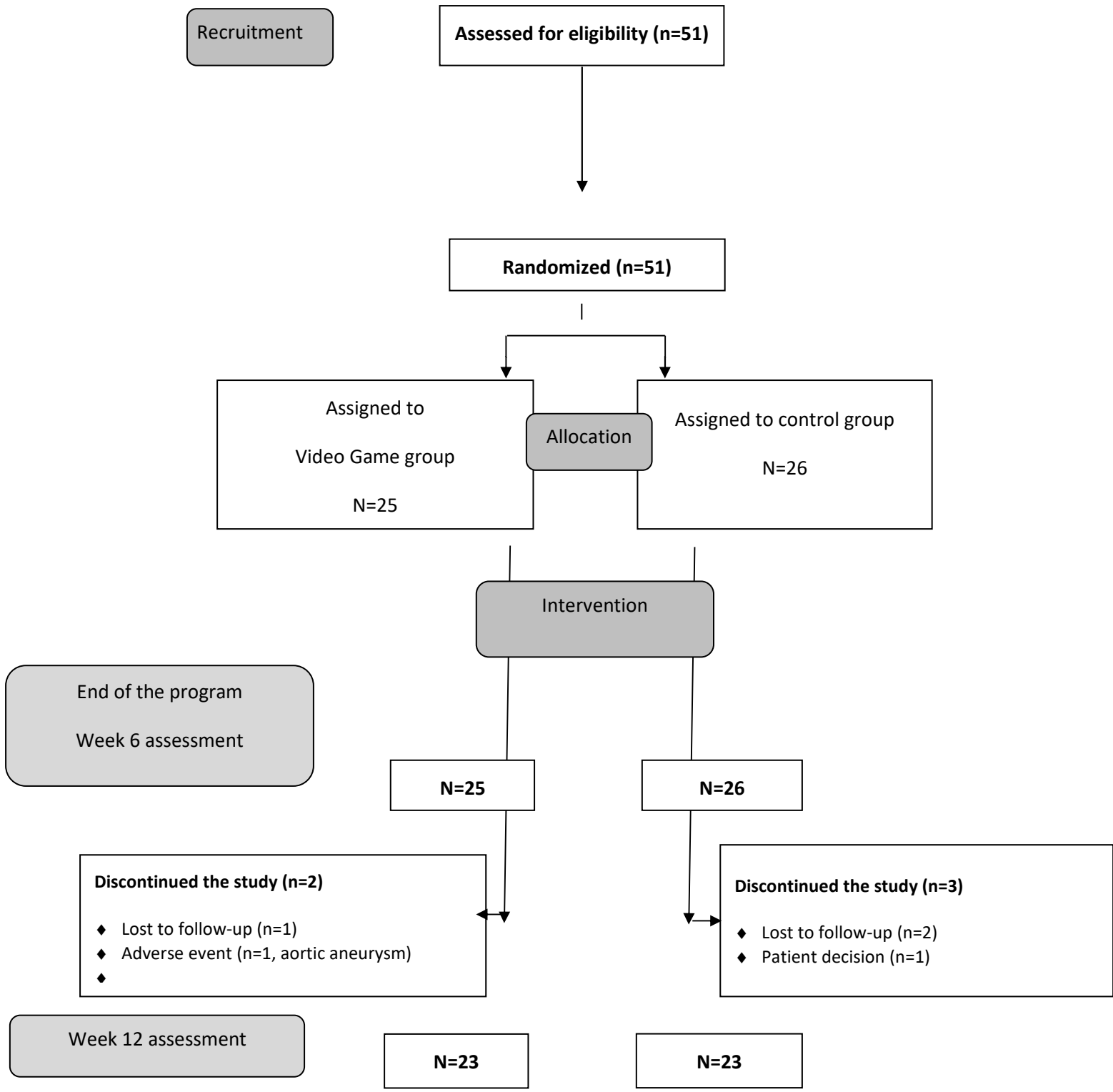
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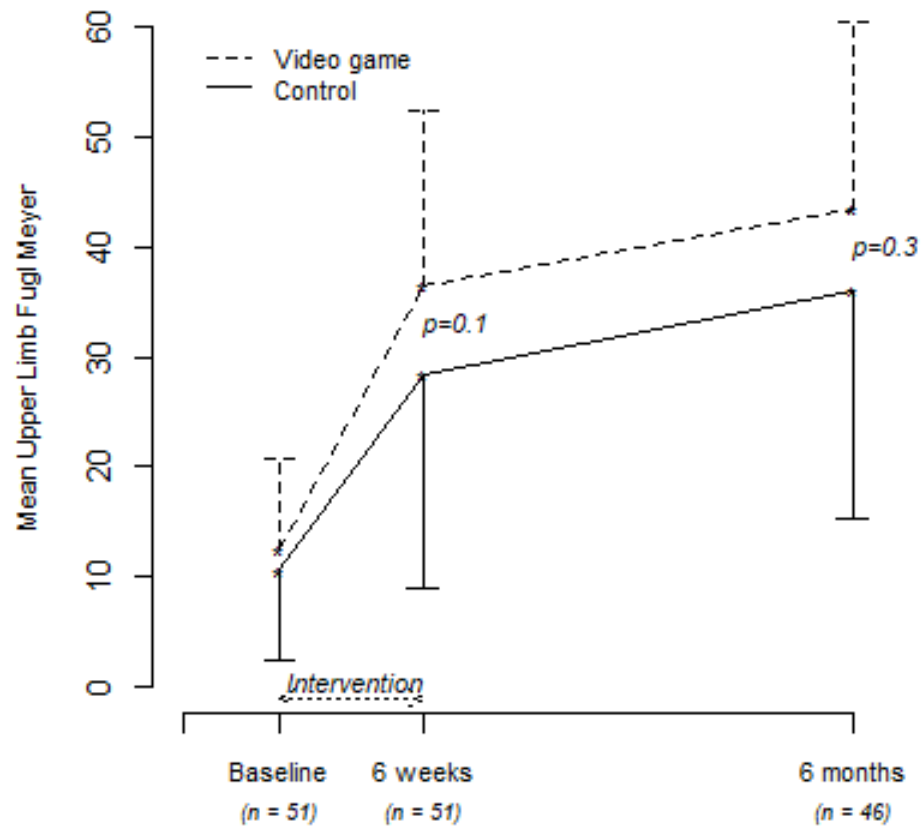


Fig2a

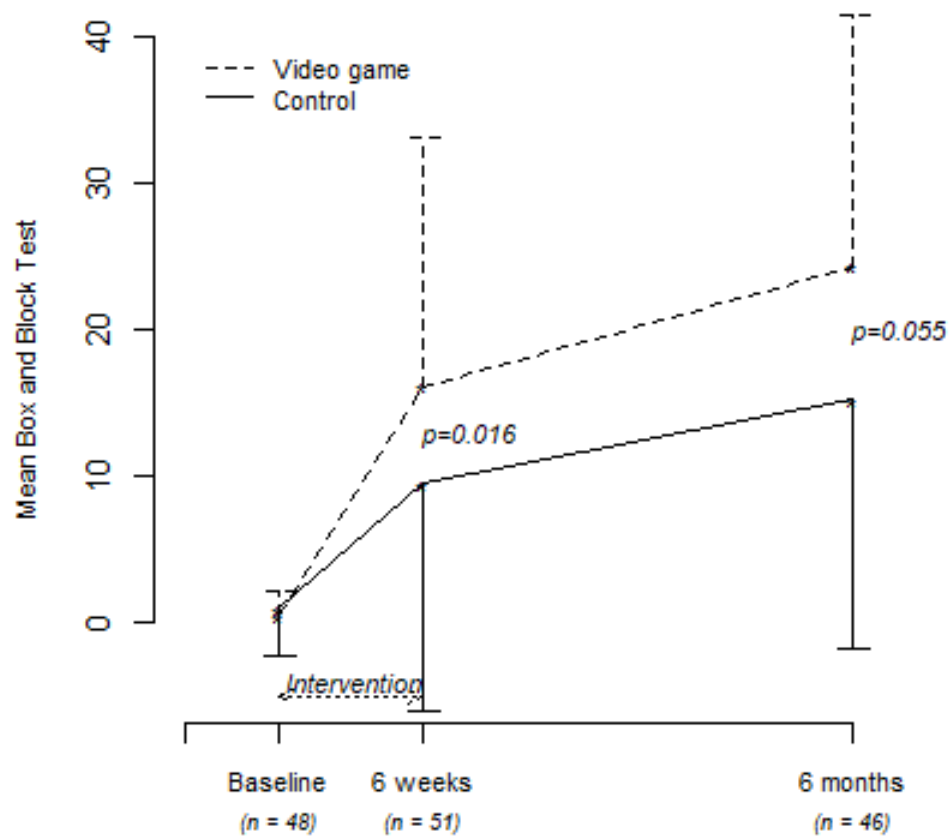


Fig2b

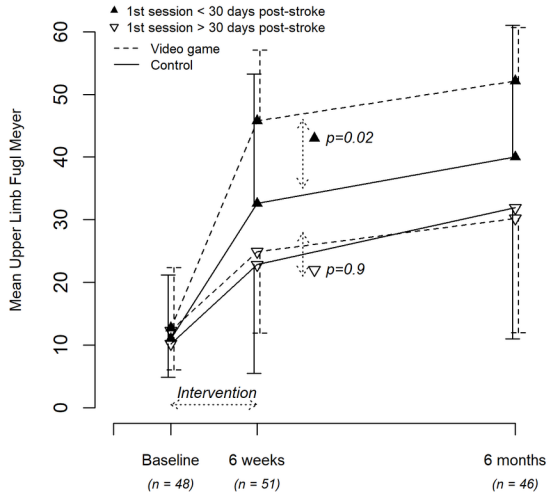


Fig3a

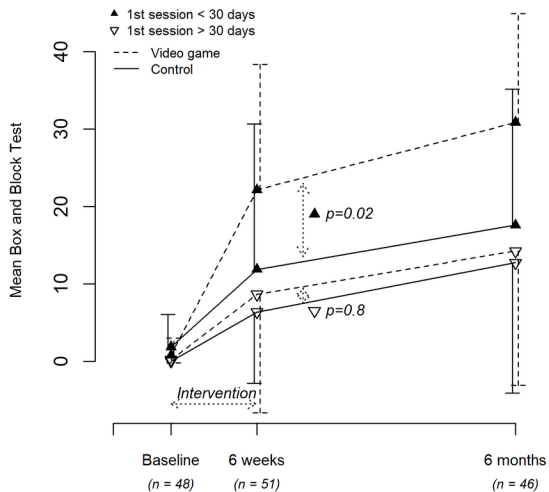


Fig3b

Table 1. Patient characteristics at inclusion for the control and video games groups.

	Control group (n=26)	Video games group (n=25)
Sex		
◦ Female	11 (42%)	9 (36%)
• Male	15 (58%)	16 (64%)
Stroke side		
◦ Right	12 (46%)	13 (52%)
• Left	14 (54%)	12 (48%)
Stroke type		
◦ Ischemic	16 (65%)	22 (88%)
• Haemorrhagic	10 (35%)	3 (12%)
Stroke lesion		
◦ Cortical	1 (5%)	0 (0%)
• Sub Cortical	6 (29%)	9 (47%)
• both	4 (19%)	7 (37%)
Age (yr)	55.8 (14.0)	60.8 (14.1)
	56.47 [15.16]	64.22 [20.77]
NIHSS	11.6 (6.1)	9.5 (4.5)
	12 [8]	9 [4]
Time since stroke (d)	27.5 (19.4)	27.0 (19.9)
	20.5 [23]	22 [26]
UL Fugl-Meyer Proximal	7.9 (5.2)	9.9 (5.1)
	6 [5]	8 [7]
UL Fugl-Meyer Distal	2.8 (4.7)	2.6 (4)
	0.5 [4]	1 [5]
UL Fugl-Meyer Total	10.7 (8.3)	12.5 (8.2)
	6.5 [12]	9 [12]
BBT affected arm	1.0 (3.3)	0.5 (1.7)
	0 [0]	0 [0]
WMFT time	81.0 (41.8)	88.6 (41.3)
	82.83 [67.13]	112.80 [52.00]
WMFT functional ability	0.7 (0.8)	0.58 (0.8)
	0.47 [1.20]	0.20 [0.93]
MAL amount of use	0.1 (0.1)	0.2 (0.6)
	0.00 [0.07]	0.00 [0.00]
MAL quality of movement	0.04 (0.1)	0.2 (0.5)
	0.00 [0.00]	0.00 [0.07]
Barthel Index	46.2 (25.2)	48.0 (23.9)
	40 [30]	45 [40]
SF36 Physical Composite Score	30.6 (5.7)	31.7 (5.5)
	30.27 [9.40]	31.42 [10.16]
SF36 Mental Composite Score	39.7 (14.3)	43.9 (11.1)
	34.37 [20.29]	41.98 [17.34]

Data are mean (SD) and median [Q1–Q3].

NIHSS, National Institute of Health Stroke Score (worse score = 42); UL, upper limb; BBT, Box and Block Test; WMFT, Wolf Motor Function Test; MAL, Motor Activity Log; SF36, Medical Outcomes Study Short Form 36.

Note that stroke lesion data were missing for some patients (n=11).

Table 2. Rehabilitation sessions for the control and video games groups during the 6-week intervention.

	Control group (n=26)	Video games group (n=25)
Standard care sessions (provided to both groups)		
Physiotherapy: total number	33.6 (8.4)	32.6 (9)
Physiotherapy: duration (min)	55.5 (7.8)	55.4 (8.3)
Occupational therapy: number	32.8 (8.5)	31.5 (9.4)
Occupational therapy: duration (min)	28.9 (4.2)	29.5 (4.1)
Speech therapy: number	12.3 (11)	8.8 (10)
Speech therapy: duration (min)	31.8 (5.5)	32 (6.1)
Additional sessions (different in the 2 groups)		
Number of sessions	32.7 (8.7)	32.7 (8.9)
Duration (min)	29.3 (4.3)	28.5 (4.4)
Nature of intervention	Occupational therapy	Video games

Data are mean (SD).

Table 3. Gains in upper limb motricity, function, independence, quality of life at 6 and 12 weeks in the control and video games group.

Assessments	Difference: week 6 from baseline				Difference: month 6 from baseline			
	Control (CR)	Video games (VG)	Number evaluated (CR/VG)	p	Control (C)	Video games (VG)	Number evaluated (CR/VG)	p
UL Fugl-Meyer Proximal	11.7 (10.2)	13.4 (9.5)	26/25	0.5	15.7 (10.2)	18.2 (9.4)	23/23	0.4
UL Fugl-Meyer Distal	6.2 (6.0)	10.7 (6.4)	26/25	0.02	9.9 (7.5)	13 (7.9)	23/23	0.3
UL Fugl-Meyer Total	17.8 (14.6)	24.1 (14.8)	26/25	0.1	25.6 (16.3)	31.2 (15.8)	23/23	0.3
BBT affected arm	7.4 (12.2)	15.7 (16.3)	23/25	0.02	14.4 (15.5)	23.8 (16.7)	21/23	0.055
WMFT time	-30.7 (39.7)	-47.3 (42.1)	25/25	0.16	-43.1 (37.1)	-56.8 (45.4)	22/23	0.39
WMFT Proximal functional ability	1.3 (1.2)	2.1 (1.4)	25/25	0.05	1.9 (1.3)	2.6 (1.6)	22/23	0.13
Nine-Hole Peg (sec)	-3.6 (5.5)	-13.4 (56.1)	23/25	0.7	-4.1 (5.5)	-14.3 (58.3)	21/23	0.5
MAL amount of use	0.8 (1.3)	1 (1.6)	25/25	0.5	1.3 (1.5)	2.2 (2)	22/23	0.2
MAL quality of movement	0.7 (1.1)	1 (1.5)	25/25	0.5	1.2 (1.4)	2 (1.8)	22/23	0.2
Barthel Index	28.6 (18.2)	33.4 (25.4)	25/25	0.4	42.7 (21.4)	40.9 (25.7)	22/23	0.8
SF36 Physical Composite Score	2.4 (8.5)	2.8 (6.5)	25/23	0.9	7.6 (8.8)	7 (9.7)	21/22	0.8
SF36 Mental Composite Score	3.6 (11.4)	2.9 (12.2)	25/23	0.8	1 (11.2)	2.8 (15.3)	21/22	0.7

Data are mean (SD).

Primary efficacy criterion = gain in UL Fugl-Meyer Total.

p-values correspond to Student *t* or Wilcoxon tests comparing gains (6 weeks and 6 months from baseline) between the 2 groups.

UL, upper limb; BBT, Box and Block Test (mean number of blocks); WMFT, Wolf motor function test; MAL, motor activity log; SF36, Medical Outcomes Study Short Form 36