

Rhythmic Sensory Stimulation on Gait in Patients With Chronic Stroke

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Research

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1 **Rhythmic Sensory Stimulation on Gait in Patients with Chronic Stroke**

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9

10 **Abstract**

11 **Background:** Patients with stroke suffer from impaired locomotion which exhibit unstable walking
12 with increased gait variability. Rhythmic sensory stimulation is one approach for improving the gait
13 of persons with stroke, Parkinson's disease, or the elderly. However, the effects of this approach on
14 the gait of patients with chronic stroke are unclear. This study was conducted to identify the effects
15 of rhythmic sensory stimulation on the gait of patients with chronic stroke.

16 **Methods:** Twenty 20 older adults with stroke (mean age \pm SD, 72.10 \pm 7.15 years; female/male,
17 8/12) and twenty age- and gender-matched healthy controls (mean age \pm SD, 72.65 \pm 6.93 years;
18 female/male, 8/12) walked 60 m under four conditions: (1) normal walking with no stimulation, (2)
19 walking with rhythmic auditory stimulation (RAS), (3) walking with rhythmic somatosensory
20 stimulation (RSS), and (4) walking with rhythmic combined stimulation (RCS: RAS + RSS). RAS
21 was applied through an earphone in the ear of each participant, while RSS was applied through a
22 haptic device on the wrist of the participant. RCS was applied simultaneously via an earphone and
23 haptic device. The gait performance (i.e., mean gait speed, stride length, gait cycle, cadence, stance
24 ratio, swing ratio, and double support ratio) and gait variability (i.e., coefficient of variation (CV)

25 value of stride length, gait cycle, stance ratio, swing ratio, and double support ratio) were evaluated.

26 **Results:** Gait performance in the stroke group was significantly improved in walking with RAS,
27 RSS, and RCS compared to normal walking with no stimulation ($P < 0.008$). Gait variability was
28 significantly decreased in the RAS, RSS, and RCS conditions compared to that during normal
29 walking ($P < 0.008$). The gait performance and variability in the healthy control group were not
30 significantly different under the RAS, RSS, or RCS conditions compared to those under normal
31 walking ($P > 0.008$).

32 **Conclusions:** Rhythmic sensory stimulation is effective in improving the gait of patients with
33 chronic stroke, regardless of the type of rhythmic stimuli, compared to healthy controls.

34 **Trial registration:** This study was approved by the Bioethics Committee (IRB-2019-04-003-001),
35 and all participants provided written informed consent.

36

37 **Keywords:** Stroke, hemiplegia, gait, stimulation

38

39

40 **Background**

41 Stroke is a neurological disorder caused by ischemic and hemorrhagic damage to cerebrovascular
42 vessels [1-4]. The most common symptom of stroke is a movement disorder defined as loss or
43 limitation of muscle control and motor function [5-7]. Patients with stroke suffer from impaired
44 locomotion in their daily lives due to movement disorders [8] and exhibit unstable walking, with
45 increased gait variability [7, 9]. Compared with healthy controls, patients with stroke have decreased
46 gait performance, generally expressed as the mean of spatiotemporal parameters [10-12], and
47 increased gait variability, expressed using the standard deviations (SDs) or coefficients of variation
48 (CVs) of the gait spatiotemporal variables [13]. Gait performance in individuals with post-stroke

49 hemiparesis is characterized by reduced walking speed, cadence, stride length, hip joint angle at peak
50 extension, knee joint angle at toe-off or during swinging [14, 15] and increased foot lateral
51 displacement during swinging [15, 16]. Changes in walking variability, known as physiological
52 signals that reflect alterations in walking characteristics due to aging and disease [17, 18], are
53 pronounced in patients with stroke [13]. For example, Balasubramanian et al. [13] reported that the
54 SDs of the spatiotemporal gait parameters, including the step length, swing, and stride time, for
55 patients with stroke were greater than those of healthy individuals in the same age group. Kao et al.
56 [19] also reported significant increases in the SDs of step length, step width, and margin of stability
57 compared to those of healthy controls while patients with stroke walked at four different speeds
58 (60%, 80%, 100% of their preferred speeds and as fast as possible). These results indicate that
59 increased walking variability is closely related to walking in patients with stroke. Reducing such
60 variability should be considered to improve walking in patients with stroke [9] because unstable
61 walking with increased gait variability in these patients can lead to falls [13, 19-23].

62 Rhythmic sensory stimulation, which can be used to improve the gait of persons with stroke,
63 Parkinson's disease, or the elderly, includes sensory feedback utilizing auditory, somatosensory, and
64 external visual stimuli that provide spatial and temporal information to promote locomotion [24, 25].
65 Rhythmic auditory stimulation (RAS), which stimulates hearing with fixed rhythms; rhythmic
66 somatosensory stimulation (RSS), which provokes the somatosensory system with rhythmic
67 vibrations; and rhythmic visual stimuli, which stimulates vision with constant patterns, affect the
68 motor system of the human body [26-28]. This rhythmic sensory stimulation enhances the walking
69 ability of patients with motion impairments and reduces walking variability. For example, RAS has
70 been shown to increase gait speed [29] and stride length [30] but to reduce the CVs of stance time
71 and double support time in patients with stroke [9]. RSS has been shown to increase the stride length
72 of patients with Parkinson's disease [31].

73 Considering these results, applying rhythmic sensory stimulation to patients with stroke
74 could potentially reduce gait variability and improve gait performance [9, 32, 33]. Nevertheless, the
75 effects of RAS, RSS, or rhythmic combined stimulation (RCS: RAS + RSS) on the gait of patients
76 with stroke are unclear. In addition, a previous study examined the ability of RAS to improve gait in
77 patients with stroke; however, the experiment included only one subject, which limits the
78 generalizability of the results [9]. Moreover, the effects of RSS and RCS on the gait of persons with
79 stroke remain unknown. Similar to previous studies [28, 34], which used a mixture of auditory and
80 visual stimulation, the main concern of this study was how the combined effects of auditory and
81 somatosensory stimuli affect the walking of patients with stroke. Therefore, this study aimed to
82 determine whether RSS, RAS, or RCS could improve the walking ability of persons with stroke. We
83 predicted that individuals with stroke will have worse walking performance and greater gait
84 variability than healthy controls and that RCS will induce greater improvement in gait in individuals
85 with stroke than RAS or RSS alone.

86

87 **Methods**

88 This study was approved by the Bioethics Committee (IRB-2019-04-003-001), and all participants
89 provided written informed consent.

90

91 ***Participants***

92 A total of 40 persons, including 20 participants with stroke and 20 gender- and age-matched healthy
93 controls, were enrolled (Fig 1). The patients with stroke were limited to those who were able to walk
94 independently and participate voluntarily in daily life, whereas the control group included healthy
95 community-dwelling participants of similar age, height, weight, and gender. Table 1 lists the specific
96 characteristics of the participants.

97 [Insert Table 1 near here]

98

99 Two 7D (3D accelerometer, 3D gyroscope, and 1D barometer) inertial measurement units
100 (IMUs; Physilog 5, Gaitup, Inc., Lausanne, Switzerland) that were validated by previous studies [35-
101 38], particularly for assessing the mobility of patients with stroke [39], were used to obtain the gait
102 measurements. Data were extracted from two 7-axis IMU sensors attached to the feet of the
103 participants and transferred to a spreadsheet file in conjunction with analysis software. The initial
104 three footprints of the collected data were manually excluded to minimize experimental variations.
105 The mean walking performance of the calculated data was determined, and the variability was
106 determined as the CV value, which was calculated as $([\text{standard deviation}/\text{mean}] \times 100)$ [18, 40].

107 Rhythmic sensory stimulation was presented as a metronome beat by linking the pulse
108 wearable haptic device (Soundbrenner, Inc., Berlin, Germany) and earphones to the tablet PC
109 application software provided by the manufacturer [41]. The wearable haptic device was portable
110 and could be worn on the wrist, whereas the earphone was a small device that could be worn on the
111 ear.

112

113 *Study design*

114 The Mini-Mental State Exam [42] and Rivermead Mobility Index (RMI) were used to assess the
115 cognitive and motor impairment levels of the participants, respectively [43]. In addition, Up and Go
116 tests were conducted to assess the quickness and dynamic equilibrium of the participants [44].
117 Thereafter, the participants rested for 5 min and then participated in gait experiments using rhythmic
118 sensory stimulation.

119 The participants received sufficient explanation from the investigators before participating
120 and performed preliminary exercises for 10 min to familiarize themselves with the experimental

121 protocol. Two 7-axis IMU sensors were attached to both feet in all patients, who were asked to return
122 from a 30 m “return point” for a total of 60 m of walking. The participants were tested under four
123 conditions (Fig 2, 3): (1) no stimulation, (2) RAS, (3) RSS, and (4) RCS. Under the first condition,
124 the participants walked normally. The investigator then matched the metronome beat with the
125 cadence collected from the normal walking of the participant and asked the participant to step in
126 time with the metronome beat [31]. Under the second condition, each participant wore earphones
127 and walked to the RAS in time with the metronome beat. Under the third condition, each participant
128 wore a wearable haptic device on the wrist without paralysis and walked to the return point following
129 vibration of the metronome. Under the fourth condition, each participant wore earphones and the
130 wearable haptic device and walked to the return point in time with the metronome beat and vibration.
131 The last three conditions according to these rhythmic sensory stimuli were performed in random
132 order. The experiment took approximately 50 min, and a 5 min rest period was provided for each
133 experimental condition. Depending on the condition of the participant, the rest time was adjusted to
134 5–15 min if needed.

135

136 *Data analysis*

137 The Shapiro-Wilks normality test was performed to confirm that the data in this study were normally
138 distributed. Gait analysis based on rhythmic sensory stimulation was performed by a two-way
139 repeated measures analysis of variance with one between factor and one within factor, and the mean
140 difference was verified at the 0.05 significance level. The independent variables were the group
141 (stroke or healthy control) and type of rhythmic sensory stimulation (no stimulation, RAS, RSS, or
142 RCS), and the dependent variables were the gait performance and variability. When there were
143 interaction effects, three tasks (RAS, RSS, and RCS) were post-tested using the multiple
144 comparisons, with Bonferroni correction. The significance level was corrected to $0.05/6 = 0.0083$

145 by the Bonferroni correction. All data analyses and statistical processes were performed using SPSS
146 ver. 20 (IBM Inc., USA) software. The effect size of the two-way repeated measures analysis of
147 variance was calculated as partial eta-squared (η_p^2). Here, η_p^2 values of 0.01, 0.06, and 0.14 represent
148 small, medium, and large effect sizes, respectively. In the post-hoc test for interactions, the effect
149 size was determined as Cohen's d, where values of 0.2, 0.5, and 0.8 correspond to small, medium,
150 and large effect sizes, respectively. The sample size of the two-way repeated measures in ANOVA
151 was calculated using G*power 3.1.9.4 software. As a result of setting the effect size (f) to 0.25,
152 significance level (α) to 0.05 and power ($1 - \beta$) to 95% in the number of samples, the optimal sample
153 size was determined to be 38 subjects.

154

155 **Results**

156 Table 2 presents the changes in gait performance and variability of the patients with stroke and
157 healthy controls owing to rhythmic sensory stimulation. The gait speed, stride length, cadence, and
158 swing ratio of the gait performance of the group of patients with stroke significantly decreased
159 compared to those of the healthy control group ($P < 0.05$). In addition, the gait cycle, stance ratio,
160 and double support ratio of the gait performance of the group of patients with stroke significantly
161 increased compared to those of the healthy control group ($P < 0.05$). There are significant differences
162 in gait speed, stride length, gait cycle, cadence, and double support ratio according to the rhythmic
163 sensory stimulation ($P < 0.05$), and there are significant differences in gait speed, stride length, and
164 double support ratio according to the interaction ($P < 0.05$).

165

166 [Insert Table 2 near here]

167

168 According to the post-hoc test, the stroke group had significantly faster gait speed under the

169 RAS ($P = 0.001$, $d = 0.29$), RSS ($P = 0.001$, $d = 0.30$), and RCS ($P = 0.001$, $d = 0.31$) conditions
170 compared to that for normal walking (Fig 4a), as well as significantly increased stride length under
171 the RAS ($P = 0.001$, $d = 0.30$), RSS ($P = 0.001$, $d = 0.27$), and RCS ($P = 0.001$, $d = 0.27$) conditions
172 compared to that for normal walking (Fig 4b). Moreover, the stroke group showed significantly
173 reduced double support ratios under the RAS ($P = 0.001$, $d = 0.28$) and RCS ($P = 0.001$, $d = 0.22$)
174 conditions compared to that for normal walking (Fig 4c).

175 The CVs of stride length, gait cycle, stance ratio, and swing ratio for the patients with stroke
176 were significantly higher than those of the healthy controls ($P < 0.05$). There are significant
177 differences in the CVs of stride length, gait cycle, stance ratio, and swing ratio according to the
178 rhythmic sensory stimulation ($P < 0.05$), as well as significant differences in the CVs of stride length,
179 gait cycle, stance ratio, swing ratio, and double support ratio according to the interaction ($P < 0.05$).

180 According to the post-hoc test, the stroke group had significantly reduced stride length CVs
181 under the RAS ($P = 0.001$, $d = 0.77$), RSS ($P = 0.001$, $d = 0.69$), and RCS ($P = 0.001$, $d = 0.73$)
182 conditions compared to that for normal walking (Fig 5a), as well as significantly reduced gait cycle
183 CVs under the RAS ($P = 0.001$, $d = 0.75$), RSS ($P = 0.001$, $d = 0.63$), and RCS ($P = 0.001$, $d = 0.62$)
184 conditions compared to that for normal walking (Fig 5b). In addition, the stroke group had
185 significantly reduced stance ratio CVs under the RAS ($P = 0.001$, $d = 0.78$), RSS ($P = 0.001$, $d =$
186 0.72), and RCS ($P = 0.001$, $d = 0.79$) conditions compared to that for normal walking (Fig 5c), as
187 well as significantly reduced swing ratio CVs for the RAS ($P = 0.001$, $d = 0.67$), RSS ($P = 0.001$, $d =$
188 0.55), and RCS ($P = 0.002$, $d = 0.65$) conditions compared to that for normal walking (Fig 5d).
189 Moreover, the stroke group had a significantly reduced double support ratio CV under the RCS ($P =$
190 0.002 , $d = 0.66$) condition compared to that for normal walking (Fig 5e).

191 However, the gait performance and variability in the healthy control group did not differ
192 significantly under the RAS, RSS, and RCS conditions from those for normal walking ($P > 0.008$).

193 In addition, there were no significant differences between the RAS, RSS, and RCS conditions in the
194 gait variability of the stroke and healthy control groups ($P > 0.008$).

195

196

197 **Discussion**

198 This study aimed to investigate the effects of rhythmic sensory stimulation on the gait performance
199 of patients with stroke. Gait data were collected through 7-axis accelerometers under four
200 experimental conditions for patients with stroke and healthy controls. The patients with stroke
201 showed decreased gait performance and increased gait variability while walking compared to the
202 healthy controls. In addition, the gait performance of the group of patients with stroke under the
203 RAS, RSS, and RCS conditions was improved; moreover, the gait variability decreased. The walking
204 performance of the healthy control group did not change under the same conditions. The main results
205 of this study are as follows.

206 First, RAS improved the gait performance and decreased the gait variability of patients with
207 stroke. Thus, the mean gait speed (8.6%) and stride length (7.6%), which are gait performance
208 parameters, increased with RAS during walking compared to those of normal walking without
209 stimulation. The double support ratio (6.6%) decreased. In addition, when RAS was applied during
210 walking, the gait variability parameters, i.e., the CVs of stride length (25.3%), gait cycle (28.1%),
211 stance ratio (21.7%), and swing ratio (25.9%), decreased. Similarly, previous studies have reported
212 that RAS improves gait performance and reduces variability in patients with stroke [7, 9]. For
213 example, Thaut et al. [7] found that RAS increases the mean gait speed and stride length in relation
214 to the walking performance of patients with stroke. Wright et al. [9] found that RAS reduces the CVs
215 of stance time and double support time in gait variability in patients with stroke. This improvement
216 in walking ability results from regular modulation of the central pattern generator of RAS [45, 46].

217 The regular pattern of rhythmic locomotion of the human body is controlled by the central nervous
218 system [47]. During walking, RAS is thought to help stabilize gait by inducing a certain movement
219 pattern by periodically stimulating the central nervous system [45]. In particular, the three rhythmic
220 sensory stimuli in this study showed significant effects in the five gait variability parameters and
221 three gait performance parameters. These results demonstrate that rhythmic sensory stimulation is
222 more effective at reducing gait variability than at improving overall gait performance in patients with
223 stroke.

224 Second, RSS improved the gait performance and decreased the gait variability of the patients
225 with stroke. The mean gait speed (8.6%) and stride length (7.6%), which are gait performance
226 parameters, increased with RSS during walking compared to those of normal walking without
227 stimulation in patients with stroke. In addition, when RSS was provided during walking, the gait
228 variability parameters, i.e., the CVs of stride length (23.7%), gait cycle (24.5%), stance ratio (22.0%),
229 and swing ratio (24.5%), decreased. Similarly, previous studies of patients with Parkinson's disease
230 and older adults who experienced falls reported that RSS improved gait performance and reduced
231 gait variability in both groups [26]. For example, van Wegen et al. [31] found that RSS increases the
232 mean step length in walking performance in patients with Parkinson's disease. Galica et al. [26]
233 argued that RSS reduces the SDs of gait variability, gait cycle, stance time, and swing time of older
234 persons with fall histories. Previous studies also indicated that a reduction in proprioception
235 increases gait variability and leads to unstable gait [48, 49]. RSS has been shown to induce a constant
236 motor pattern by periodically promoting somatosensory input in patients with stroke [45]. Similarly,
237 in this study, RSS helped induce a stable gait in patients with stroke. This is the first study to verify
238 the positive effect of RSS on the gait of patients with stroke.

239 Third, in this study, RCS improved the gait performance and decreased the gait variability of
240 patients with stroke. The gait performance parameters, mean gait speed (8.6%) and stride length

241 (7.6%), increased, and the double support ratio (5.3%) decreased with RCS compared to those of the
242 natural gait without stimulation. However, when RCS was provided during walking, the gait
243 variability parameters, i.e., stride length (25.1%), gait cycle (24.5%), stance ratio (22.7%), swing
244 ratio (26.0%), and double support ratio (17.3%), decreased. These results were somewhat different
245 from those of previous studies on the effects of mixed stimulation in patients with Parkinson's
246 disease. For example, Arias and Cudeiro [28] found that mixed stimulation of auditory and visual
247 information did not improve the mean speed in the walking performance of patients with Parkinson's
248 disease. Suteerawattananon et al. [34] reported that mixed stimulation of auditory and visual
249 information did not increase the mean stride length related to gait performance in patients with
250 Parkinson's disease. It is not possible to distinguish clearly whether these conflicting results are due
251 to the different subjects using the rhythmic sensory stimuli or the different combinations of sensory
252 stimuli, because the subjects of the previous studies by Arias and Cudeiro [28] and
253 Suteerawattananon et al. [34] were patients with Parkinson's disease, and the stimuli were different.
254 Therefore, follow-up studies on the effects of various forms of rhythmic sensory stimulation on gait
255 are required.

256

257 **Study Limitations**

258 First, the severity of patients with stroke was evaluated as RMI in this study; however, only one
259 stroke group was considered. Stroke severity could affect the sensitivity of the sensory receptors that
260 utilize the sensory signals in the gait process. Second, the walking speed of the patients with stroke
261 was not sufficiently considered. Understanding how rhythmic sensory stimulation affects patient gait
262 at various gait speeds can provide adequate information for designing an optimal program that will
263 further improve the gait abilities of patients with stroke during rehabilitation. For reference, this
264 study matched the cadence and metronome beats of the rhythmic sensory stimulation in the natural

265 gait of the patients with stroke; however, the gait change of these patients may differ at different gait
266 speeds.

267

268 **Conclusions**

269 Treatment methods using rhythmic sensory stimulation improved gait performance and decreased
270 gait variability of patients with stroke, regardless of the type of rhythmic stimuli. These benefits
271 were achieved despite the increases in walking speed or stride length of the patients with stroke.
272 Rhythmic sensory stimulation was more effective at stabilizing the walking of these patients
273 compared to that of the healthy control group. This study revealed that the use of rhythmic sensory
274 stimulation could improve the gait of patients with stroke with a relatively high risk of falling.

275

276 **List of abbreviations**

277 Coefficient of variation (CV), inertial measurement units (IMU), rhythmic auditory stimulation
278 (RAS), rhythmic combined stimulation (RCS), Rivermead Mobility Index (RMI), rhythmic
279 somatosensory stimulation (RSS).

280

281 **Declarations**

282 *Ethics approval and consent to participate*

283 This study was approved by the Bioethics Committee (IRB-2019-04-003-001). All participants
284 provided written informed consent.

285

286 *Consent for publication*

287 Not applicable.

288

289 ***Availability of data and materials***

290 The datasets used during the current study are available from the corresponding author on reasonable
291 request.

292

293 ***Competing interests***

294 The authors declare that they have no competing interests.

295

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298

299 ***Authors' contributions***

300 YL and SS designed the study. YL performed the data collection. YL and SS analyzed the data
301 interpreted and wrote the manuscript for publication. SS read and approved the final manuscript.

302

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305

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432

433

434 **Figure Captions**

435 Fig 1 Experimental design flow diagram showing experimental design.

436

437 Fig 2 a) Experimental scene of a patient with stroke during a 60-m walk. b) Rhythmic auditory
438 stimulation (RAS) condition. c) Rhythmic somatosensory stimulation (RSS) condition. d) Raw data
439 extracted from 7D IMU sensor.

440

441 Fig 3 Examples of gait cycle from a patient with stroke in all walking conditions. a) No stimulation:
442 normal walking; b) RAS: rhythmic auditory stimulation; c) RSS: rhythmic somatosensory
443 stimulation; and d) RCS: rhythmic combined stimulation (RAS + RSS). The horizontal lines are the
444 average gait cycle of patients with stroke. CV: Coefficient of variation.

445

446 Fig 4 Effects of rhythmic sensory stimulation on gait performance compared to normal walking. a)
447 speed; b) stride length; and c) double support. Normal: no stimulation; RAS: rhythmic auditory
448 stimulation; RSS: rhythmic somatosensory stimulation; RCS: rhythmic combined stimulation (RAS
449 + RSS). * indicates significant difference compared to normal condition ($P < 0.008$).

450

451 Fig 5 Effects of rhythmic sensory stimulation on gait variability compared to normal walking.
452 Coefficients of variation (CV) of a) stride length; b) gait cycle; c) stance; d) swing; and e) double
453 support. Normal: no stimulation; RAS: rhythmic auditory stimulation; RSS: rhythmic somatosensory
454 stimulation; RCS: rhythmic combined stimulation (RAS + RSS). * indicates significant difference
455 compared to normal condition ($P < 0.008$).

Figures

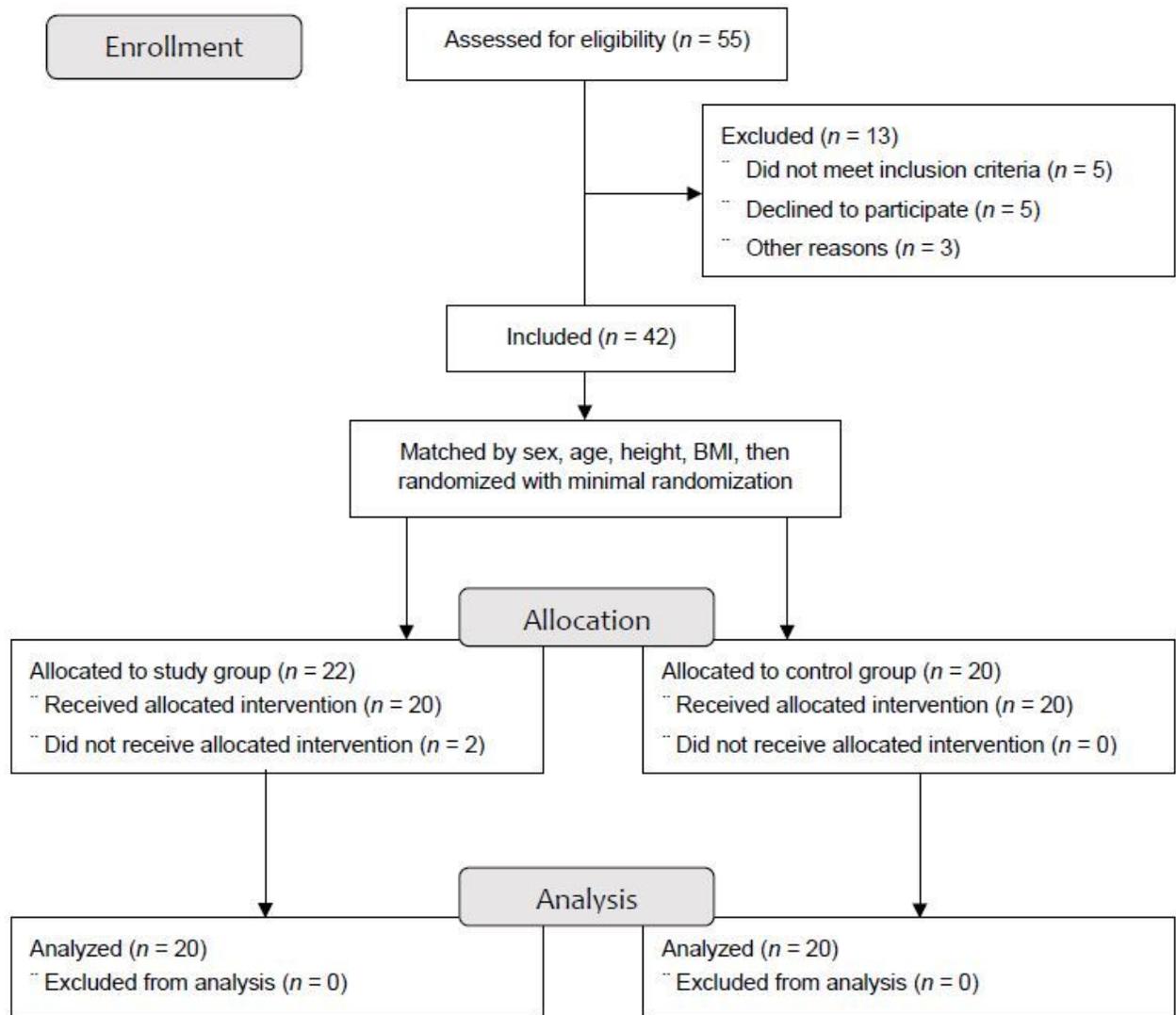


Figure 1

Experimental design flow diagram showing experimental design.

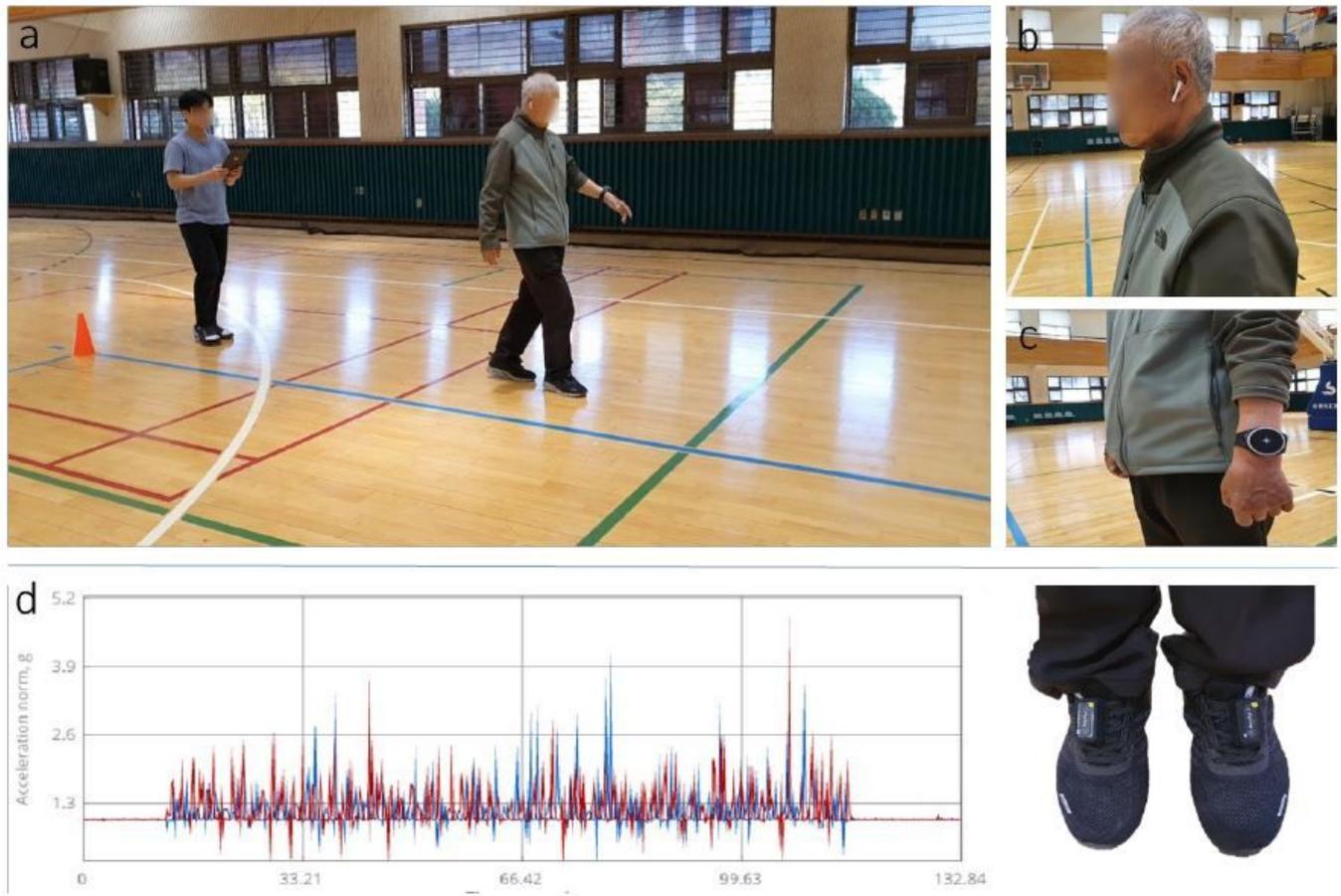


Figure 2

a) Experimental scene of a patient with stroke during a 60-m walk. b) Rhythmic auditory stimulation (RAS) condition. c) Rhythmic somatosensory stimulation (RSS) condition. d) Raw data extracted from 7D IMU sensor.

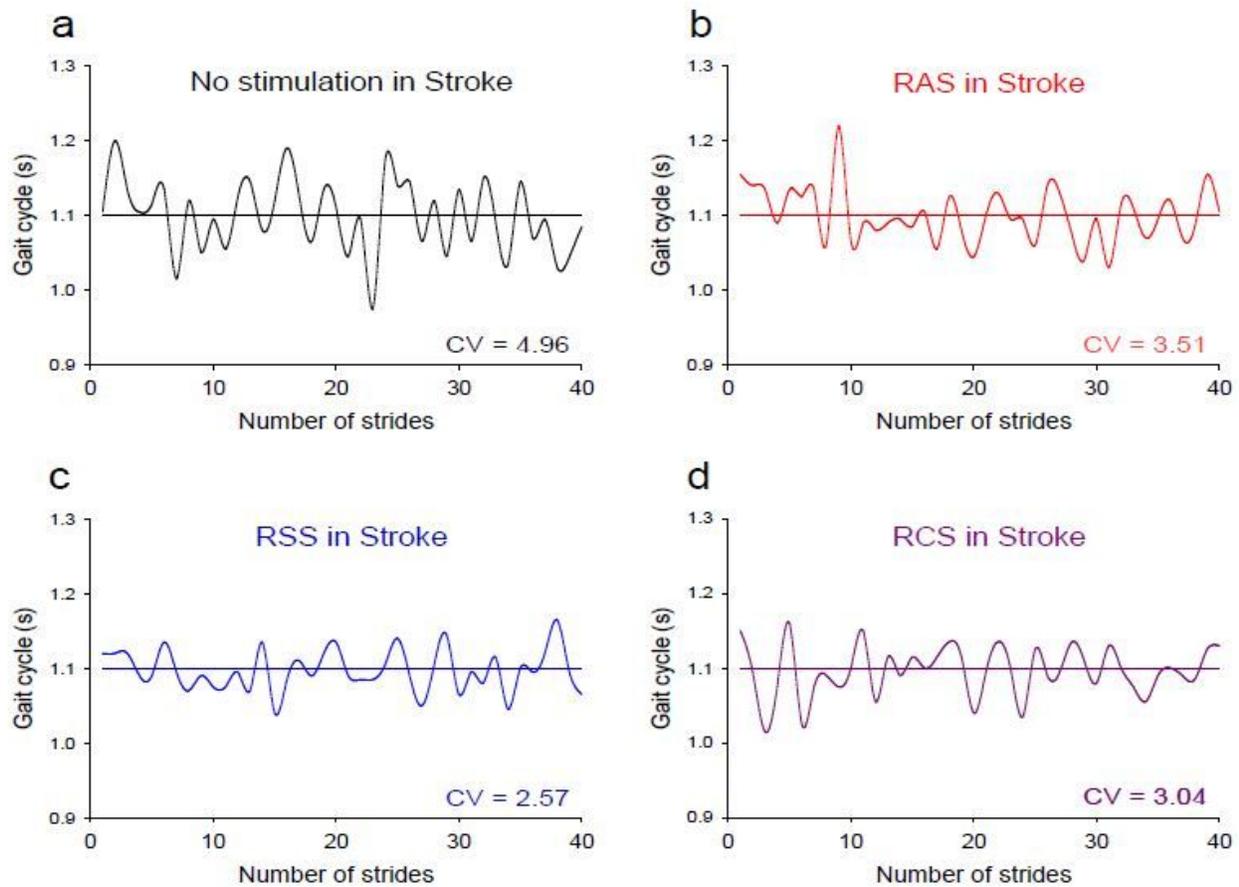


Figure 3

Examples of gait cycle from a patient with stroke in all walking conditions. a) No stimulation: normal walking; b) RAS: rhythmic auditory stimulation; c) RSS: rhythmic somatosensory stimulation; and d) RCS: rhythmic combined stimulation (RAS + RSS). The horizontal lines are the average gait cycle of patients with stroke. CV: Coefficient of variation.

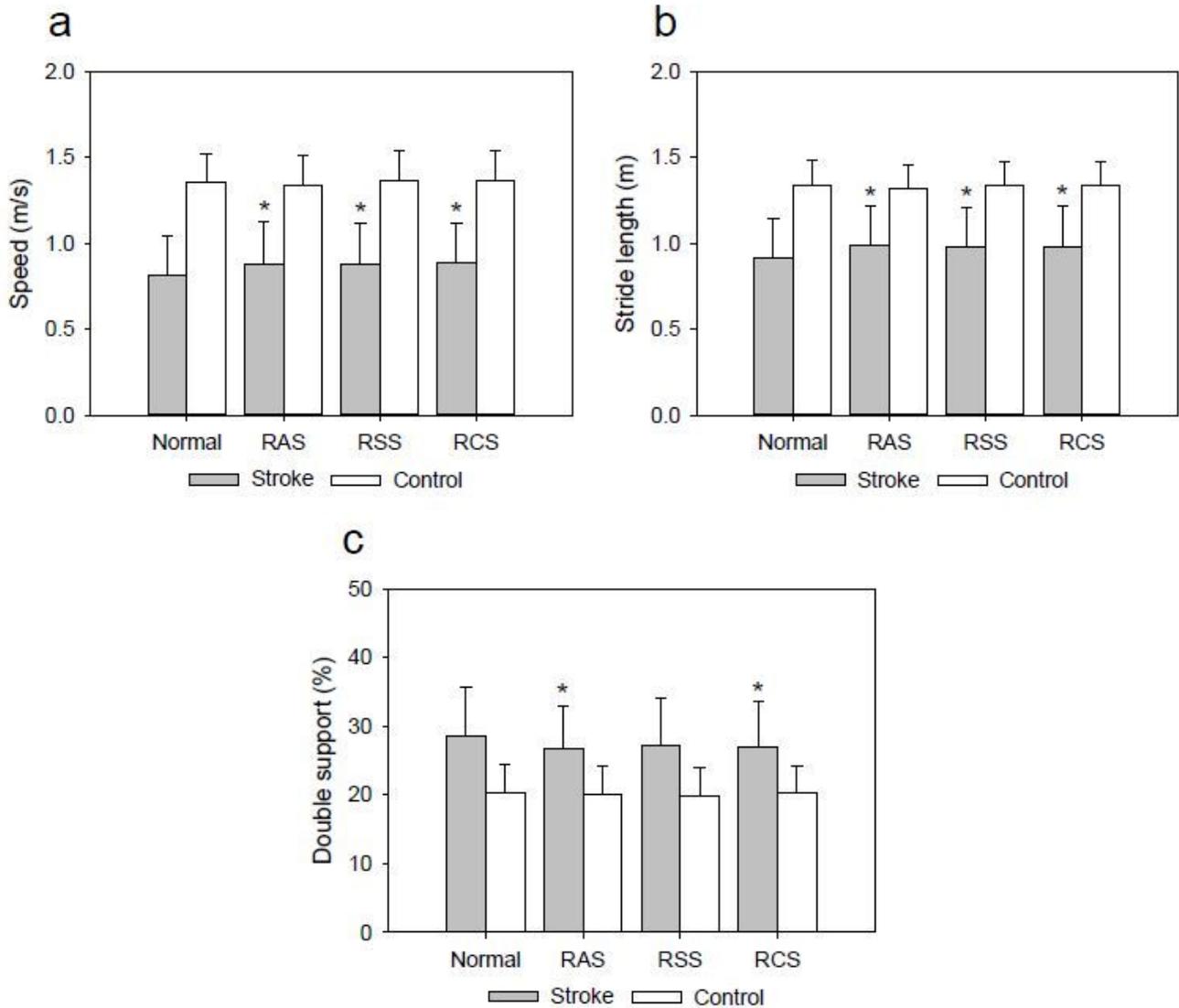


Figure 4

Effects of rhythmic sensory stimulation on gait performance compared to normal walking. a) speed; b) stride length; and c) double support. Normal: no stimulation; RAS: rhythmic auditory stimulation; RSS: rhythmic somatosensory stimulation; RCS: rhythmic combined stimulation (RAS + RSS). * indicates significant difference compared to normal condition ($P < 0.008$).

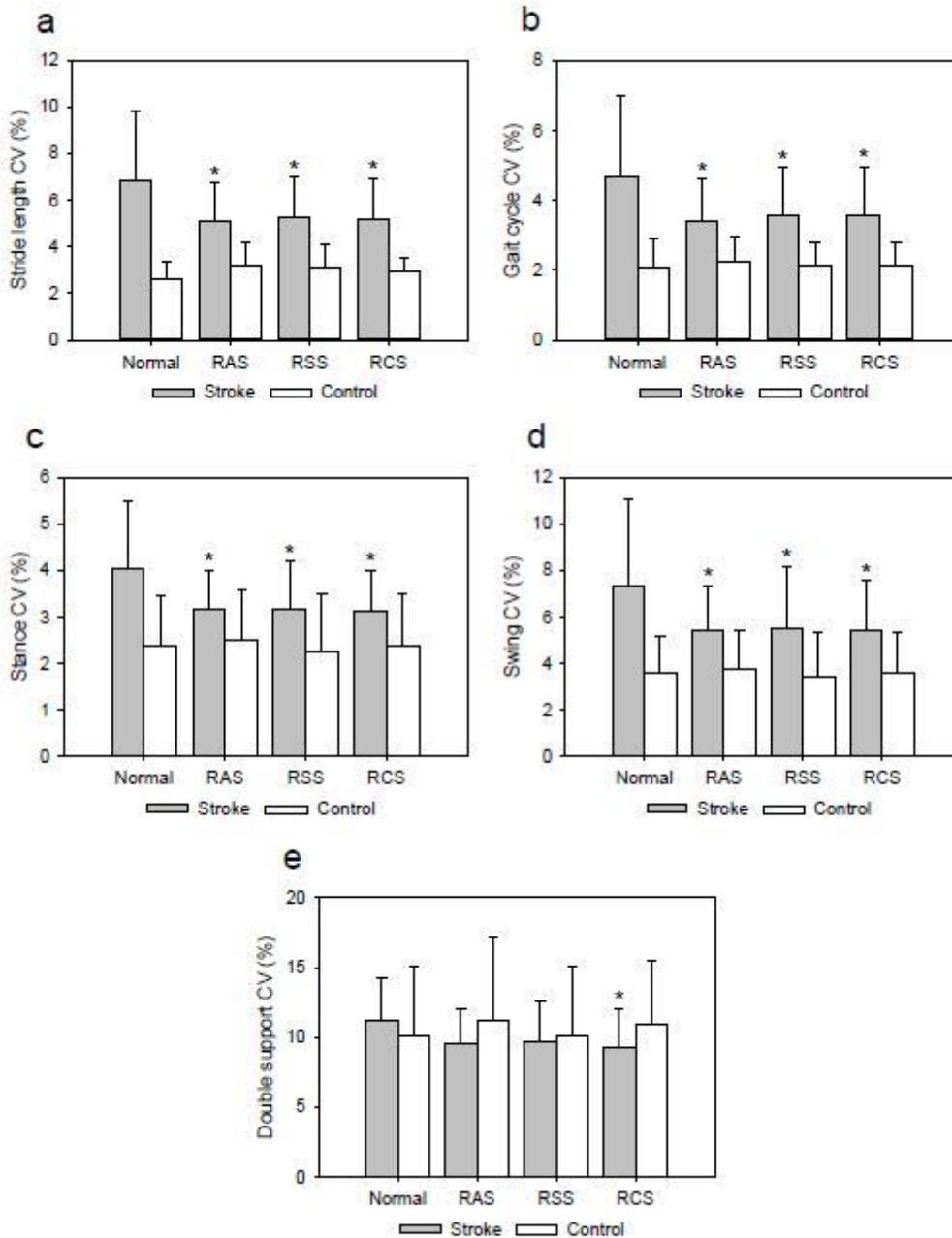


Figure 5

Effects of rhythmic sensory stimulation on gait variability compared to normal walking. Coefficients of variation (CV) of a) stride length; b) gait cycle; c) stance; d) swing; and e) double support. Normal: no stimulation; RAS: rhythmic auditory stimulation; RSS: rhythmic somatosensory stimulation; RCS: rhythmic combined stimulation (RAS + RSS). * indicates significant difference compared to normal condition ($P < 0.008$).

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