

1 **Title: Effects of dual-task training on balance and executive functions in**  
2 **Parkinson's disease: A pilot study**

3

4 **Short-Title: Dual-task training in Parkinson's**

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17

1 **Abstract**

2 The aim of this study was to analyze the efficacy of cognitive-motor dual-task training  
3 compared with single-task training on balance and executive functions in individuals  
4 with Parkinson's disease.

5 15 subjects, aged between 39 and 75 years old were randomly assigned to the dual-  
6 task training group (n=8) and single-task training (n=7) groups. The training was run  
7 twice a week for six weeks. The single-task group received balance training, and the  
8 dual-task group performed cognitive tasks simultaneously with the balance training.

9 There were no significant differences between the two groups at baseline. After the  
10 intervention, the results for mediolateral sway with eyes closed were significantly  
11 better for the dual-task group and anteroposterior sway with eyes closed was  
12 significantly better for the single-task group. The results suggest superior outcomes  
13 for the dual-task training compared to the single-task training for static postural  
14 control, except in anteroposterior sway with eyes closed.

15

16 **Keywords**

17 Parkinson's disease, dual-task training, executive functions, balance

18

## 1 **Introduction**

2 Parkinson's disease (PD) is considered to be the second most common  
3 neurodegenerative disorder affecting currently about 1% of the world population  
4 (Andlin-Sobocki, Jonsson, Wittchen, & Olesen, 2005; Campenhausen et al., 2005;  
5 Rodrigues de Paula, Teixeira-Salmela, Faria, Brito, & Cardoso, 2006). Some  
6 projections point to a large increase in this prevalence over the next decades  
7 (Campenhausen et al., 2005).

8 PD is clinically defined by motor symptoms such as tremor at rest, rigidity,  
9 bradykinesia, as well as postural and gait modifications (Giroux, 2007; Wielinski,  
10 Erickson-Davis, Wichmann, Walde-Douglas, & Parashos, 2005); and also by non-  
11 motor symptoms such as sleep disorders, cognitive impairment, depression and  
12 fatigue, some of which are adverse effects of the dopaminergic medication (Hubert &  
13 Fernandez, 2012). Another characteristic feature of PD is the difficulty to perform two  
14 tasks simultaneously. This difficulty is because the individuals have to focus on  
15 achieving normal movement patterns by activating the premotor cortex region without  
16 using the deficient basal ganglia circuit which is deficient in dopamine. Therefore, in  
17 dual-task situations that use the cortical resources to perform motor tasks, the  
18 performance of both the motor and cognitive components can be compromised  
19 (Brauer & Morris, 2010; Wu & Hallett, 2009). From this point of view, dual-task  
20 training should be considered as part of the rehabilitation process of these patients  
21 (Wu & Hallett, 2009), although until now no guidelines have been defined for this type  
22 of intervention. New paradigms have been studied concerning cognitive training,  
23 such as interventions of cognitive-motor dual-task. This type of intervention should be  
24 able to improve dual-task performance and/or improve motor and cognitive  
25 components individually (K. Baker, Rochester, & Nieuwboer, 2007; Montero-Odasso,

1 Verghese, Beauchet, & Hausdorff, 2012; Silsupadol, Siu, Shumway-Cook, &  
2 Woollacott, 2006; Yogev-Seligmann, Rotem-Galili, Dickstein, Giladi, & Hausdorff,  
3 2012).

4 Regarding specific dual-task training, recent studies have demonstrated its efficacy in  
5 various populations such as the elderly and individuals with neurological diseases,  
6 with the most notable improvements in gait and balance (Brauer & Morris, 2010;  
7 Sethi & Raja, 2012; Silsupadol, Lugade, et al., 2009; Silsupadol, Shumway-Cook, et  
8 al., 2009). This type of intervention for PD individuals has been focused mainly on  
9 gait (Brauer & Morris, 2010; Yogev-Seligmann, Giladi, Brozgol, & Hausdorff, 2011),  
10 and shows improvements in gait speed and gait variability during dual-task training.  
11 However, there is no evidence in the literature of the effects of this training on  
12 balance and executive functions evaluated independently for PD individuals. On the  
13 other hand, such separate evaluation of cognitive-motor dual-task training could be  
14 positive and enhance the meaningfulness of this type of training. Thus, considering  
15 the positive results of specific cognitive-motor dual-task training obtained in other  
16 populations and in other situations that could possibly be reproduced here, we  
17 conducted a randomized trial to study the efficacy of a cognitive-motor dual-task  
18 training program compared to a single-task program, and evaluated the cognitive and  
19 motor components independently, on PD individuals. Accordingly, we hypothesized  
20 that cognitive-motor dual-task training is more effective at improving balance and  
21 executive functions than single-task training in PD individuals.

22

## 23 **Materials and Methods**

### 24 **Participants**

1 Subjects with Parkinson's disease were recruited from the Portuguese Association of  
2 Parkinson's Patients. The inclusion criteria used were: capacity to walk ten meters  
3 without gait assistance, diagnosis of PD up to Stage 3 according to the modified  
4 Hoehn & Yahr scale. The exclusion criteria used were: cognitive deficit confirmed by  
5 the Mini Mental State Examination (Folstein, Folstein, & McHugh, 1975) using the  
6 following cut-off values according to the education level ( $\leq 22$  for 0-2 years of literacy;  
7  $\leq 24$  for 3-6 years; and  $\leq 27$  for  $\geq 7$  years (Morgado, Rocha, Maruta, Guerreiro, &  
8 Martins, 2009)), subthalamic neurosurgery, other neuromusculoskeletal and  
9 psychiatric disorders and illiteracy.

10 The subjects that voluntarily accepted to participate were randomized to either the  
11 dual-task or single-task training group. The random assignment procedure was  
12 performed with numbers generated by a computer program (Microsoft Office Excel  
13 2010), operated by an independent investigator. From a total of 23 eligible subjects,  
14 20 were included in the two groups. Before the intervention program started, there  
15 were 3 dropouts in the single-task training group (1 for surgery, 1 due to illness and 1  
16 who had various absences) and 2 dropouts in the dual-task training group (1 for  
17 personal reasons and 1 due to illness). Hence, 7 subjects were analyzed in the  
18 single-task training group and 8 subjects in the dual-task training group. These 15  
19 subjects made up the intervention program as shown in Figure 1.

20

21 < Insert Figure 1 about here >

22

23 The researcher that evaluated the results was not involved in the training program  
24 and had no knowledge to which group the subjects had been assigned, in order to  
25 prevent any possible critical judgment and manipulation of the results during the

1 evaluations. In addition, the participants were unaware of the two groups, making this  
2 a double-blind study.

3 The study was explained to each participant according to the intervention group in  
4 which they were randomly included. All participants gave their written informed  
5 consent in accordance to the Declaration of Helsinki, ensuring data confidentiality  
6 and freedom to withdraw from the program at any time. The study was approved by  
7 the ethics committee of “Instituto Politécnico do Porto – Escola Superior de  
8 Tecnologia da Saúde” and by the directive board of “Associação Portuguesa de  
9 Doentes de Parkinson”, in Portugal.

10

## 11 **Intervention**

12 All participants received balance training that was administered individually twice a  
13 week (60 min/session) for six weeks. All participants performed the same motor  
14 tasks; however, the participants of the dual-task group underwent the cognitive-motor  
15 dual-task training program and performed the cognitive tasks simultaneously with the  
16 motor tasks, while the participants of the dual-task group only underwent the single-  
17 task motor training program, and thus only performed the motor tasks. The  
18 intervention program was based on an existing training program (Silsupadol et al.,  
19 2006). The individual training sessions took place at the “Associação Portuguesa de  
20 Doentes de Parkinson” or at the “Instituto Politécnico do Porto – Escola Superior de  
21 Tecnologia da Saúde” according to each participant’s preference. Each session was  
22 organized into 4 stations of intervention, according to Gentile's taxonomy (Gentile,  
23 2000): stability without manipulation activities (e.g. to stand on top of a foam mattress  
24 with the eyes closed); gait without manipulation (e.g.: walk on a narrow path); stability  
25 with handling activities (e.g. rotate the waist holding a ball) and gait manipulation

1 activities (e.g. walking backwards around objects while holding a basket). The  
2 duration of the training sessions was the same for both groups. In the dual-task  
3 training, the cognitive activities included digit span (memorize a set of letters or  
4 numbers and repeat them in forward or reverse order), N-back (naming a preceding  
5 word, letter or number to the one given by the researcher), spelling words  
6 (researcher says words to be spelled in the correct order), stroop test (consists of two  
7 tasks, reading and naming colours. In both, the stimuli are colour names printed in an  
8 incongruent colour), image description (a picture is placed in front of the participant  
9 who should describe it with maximum detail), nomination (the participant must say  
10 names in a given category: flowers, animals, countries or beginning with a letter of  
11 the alphabet), counting (counting in forward and reverse order), description of daily  
12 activities and routines (describe the activities that they normally do during a weekday  
13 or weekend and describe how to do these activities, e.g. what are the stages of  
14 taking a shower).

15 All participants in the dual-task group performed the same cognitive activities, but not  
16 necessarily in the same order. The complexity of the exercises was increased as the  
17 sessions progressed. This increase was based on the addition of obstacles, reduction  
18 of the pause time, increasing the complexity of the cognitive task. Each participant  
19 received individual training by a professional for 12-15 minutes at each station, which  
20 led to a total of 60 minutes per session. Between stations, the participants performed  
21 a transition exercise, which was getting up from and sitting down on a chair 15 times.  
22 Before beginning the exercises, all procedures were explained to the participant. No  
23 reference was made to the tasks the participant should give more importance to.

24

## 25 **Outcome Measurements**



1 All outcome measurements were evaluated at baseline and after the intervention for  
2 all participants by a clinician who was blinded to the participant's group.

3 The outcome measurements of motor performance were obtained by Time Up and  
4 Go test (TUG), Unified Parkinson's Disease Rating Scale-part III (UPDRS-III) and  
5 pressure platform.

6 The Timed Up and Go test was used to assess the time the participant took to get up  
7 from a chair, walk 3 meters and return to the same chair (the total distance walked  
8 was 6 meters) and sit down again. The time value chose for each participant was the  
9 best, i.e. the lowest value, of three trials performed (Podsiadlo & Richardson, 1991).

10 The test-retest reliability and inter-rater reliability were ICC = 0.80 and  $r = 0.99$ ,  
11 respectively (Lim et al., 2005). UPDRS (Goetz et al., 2003) assesses the signs,  
12 symptoms and perception of individuals concerning their performance of activities of  
13 daily living (ADLs), based on a self-report and clinical observations; it should be  
14 noted that only the motor exploration (UPDRS-III) was applied. This assessment had  
15 a high internal consistency (Cronbach's alpha = 0.96) and a satisfactory inter  
16 reliability (all items had  $k > 0.40$ ) (Martínez-Martín et al., 1994). The pressure  
17 platform used was an Emed, from Novel (Germany), model AT 25A, with a sensorial  
18 area of  $380 \times 240 \text{ mm}^2$  and sensor resolution equal to 2 sensors/cm<sup>2</sup>. As a  
19 stabilometric measurement, the centre of pressure (COP) was evaluated in terms of  
20 the mediolateral direction (COP<sub>x</sub>), the anteroposterior direction (COP<sub>y</sub>), and the total  
21 velocity (V<sub>t</sub>) (Błaszczuk & Orawiec, 2011; Ganesan, Pal, Gupta, & Sathyaprabha,  
22 2010; Holmes, Jenkins, Johnson, Adams, & Spaulding, 2010). The participants were  
23 instructed to stand on the platform and remain in a self-selected comfortable upright  
24 position. The pressure data was taken twice: first, the subjects were instructed to  
25 remain standing on the platform and look towards a fixed point at a distance of 2

1 meters for 60 seconds with their eyes open (EO); second, the subjects were  
2 instructed to remain on the same platform for the same time but now with their eyes  
3 closed (EC) (Ebersbach & Gunkel, 2011). The EO/EC order was randomized in order  
4 to avoid any possible learning effect. The acquisition frequency of 25 Hz and  
5 normalized relative to each subject's body base of support.

6 The outcome measurements of cognitive performance were obtained by Rule Shift  
7 Cards Test (RSCardsT) and Trail Making Test (TMT) A and B. The RSCardsT is  
8 used to evaluate perseverance trends and the ability to switch from one pattern to  
9 another, by taking into account the errors and the time taken to complete the task  
10 (Golden, Espe-Pfeifer, & Wachsler-Felder, 2000). The TMT (Reitan, 1992) is a test  
11 divided into two parts: Part A evaluates attention and processing speed; and part B  
12 that assesses the cognitive flexibility and sequential alternation. In each part, the final  
13 score is the total time needed to complete the task (Reitan, 1992).

14 As in other similar studies with this type of population, all tests were carried out when  
15 the participants were taking the prescribed medication, denoted as "ON" medication  
16 (Conradsson, Löfgren, Ståhle, Hagströmer, & Franzén, 2012; Kelly, Eusterbrock, &  
17 Shumway-Cook, 2012).

18

## 19 **Statistical Analysis**

20 According to the nature of the variables under study, descriptive statistical analysis  
21 was performed using proportions for the variable gender, and measures of central  
22 tendency and dispersion for the variables age, education, hour of physical activity,  
23 height, weight, years of disease and intervention outcomes.

24 For the inferential analysis, the Kolmogorov-Smirnov test was used to assess data  
25 normality. Since the normality of the data distribution could not be assumed, we

1 chose to use non-parametric tests. The Mann-Whitney test for independent samples  
2 was used to verify the differences between the two groups at baseline and after  
3 intervention. In order to analyze which of the interventions was more effective, the  
4 changed scores (after the interventions relative to baseline) were used. Two-tailed  
5 tests were used in all analyses and were considered statistically significant when  
6  $p < 0.05$ . The training effect was calculated using the Cohen's d rule of thumb (Cohen,  
7 1988): low,  $0.20 \leq d < 0.50$ ; medium,  $0.50 \leq d < 0.80$ ; and high,  $d \geq .80$ . The data  
8 collected was conducted using IBM SPSS Statistics 22.0 (SPSS, Inc., Chicago, IL,  
9 USA).

10

## 11 **Results**

12 The values in Table 1 reveal that there were no significant differences between the  
13 two groups in terms of age, gender, education level, weight, height, years of illness  
14 and number of falls. Concerning the cognitive performance, there were no significant  
15 differences between groups at baseline on the RSCardsT, TMT A and B. As to the  
16 motor performance, there were no differences between groups on UPDRS-part III,  
17 TUG and COPx, COPy and Vt with eyes open and with eyes closed.

18

19 < Insert Table 1 about here >

20

21 In order to analyze which of the interventions was more effective, the differences  
22 between the two groups were statistically analyzed after the interventions relative to  
23 baseline, Table 2. In terms of the motor performance, the only differences were found  
24 in COPx and COPy with eyes closed. As to the COPx, the difference between  
25 baseline and after intervention was significantly higher for the dual-task group than

1 for the single-task group,  $U=7.5$ ,  $p=0.026$ , with high effect size,  $d=1.094$ . The  
2 difference between baseline and after intervention in terms of the COPy was  
3 significantly lower for the dual-task group than for the single-task group,  $U=7.5$ ,  
4  $p=0.029$ , with high effect size,  $d=1.43$ . Nevertheless, the total velocity ( $V_t$ ) with eyes  
5 open and with eyes closed revealed a high effect size ( $d=0.922$  and  $d=0.902$ ,  
6 respectively), and the remaining variables had a medium effect size.  
7 No significant differences were found between the two groups in terms of the  
8 executive functions performed. However, the TMT B had a high effect size ( $d=0.839$ ),  
9 the RSCardsT presented a medium effect size ( $d =0.590$ ) and the TMT A had a small  
10 size effect ( $d=0.324$ ).

11

12

< Insert Table 2 about here >

13

## 14 **DISCUSSION**

15 Studies have reported the positive influence of targeted interventions for motor  
16 training, whether for different cognitive components, including level of attention,  
17 processing speed, flexibility and alternating sequential, or for neuromotor issues,  
18 mainly in terms of muscle resistance, coordination, balance and agility (L. Baker et  
19 al., 2010; Davis et al., 2013; Mirelman et al., 2011; Moher, Liberati, Tetzlaff, &  
20 Altman, 2009; Tabak, Aquije, & Fisher, 2013; Tanaka et al., 2009). Our research has  
21 demonstrated that in a cognitive-motor dual-task training program with 12 sessions,  
22 the dual-task training was only statistically more effective than the single-task training  
23 for the COPx with eyes closed. A lower oscillation, i.e. smaller COP displacements,  
24 corresponds to a higher postural stability (Mochizuki, Duarte, Amadio, Zatsiorsky, &  
25 Latash, 2006) and thus, in agreement, our results suggested a better balance after

1 the intervention program in the dual-task training group. As to COPy with closed  
2 eyes, significant differences were also found, but the dual-task training group  
3 presented values worse than the single-task training group. This fact can be  
4 explained by the number of years of the disease that was higher in the dual-task  
5 training group. The centre of pressure of these participants was shifted to a more  
6 posterior position in order to compensate the usual postural deformities caused by  
7 high muscular rigidity (Jankovic, 2008; Matinolli, 2009). This body position, together  
8 with the loss of postural reflexes, age-related sensory changes, as well as other  
9 features, leads to greater instability in the anteroposterior component (Jankovic,  
10 2008).

11 COPx and COPy values with eyes open did not show significant differences between  
12 the two groups, but these variables had lower values after intervention in both. Some  
13 authors as, for example, (Oie, Kiemel, & Jeka, 2002; Tjernström, Fransson,  
14 Hafström, & Magnusson, 2002) defend that vision provides important feedback to  
15 the subjects about the physical environment, their spatial interactions and body sway,  
16 which complements the information provided by other sensorial receivers. Thus, the  
17 eyes open provides important information about postural orientation and helps to  
18 optimize the balance control, which may explain the better results found for COP  
19 displacement under this condition.

20 With regard to the Vt, it was found that the results were not statistically significant,  
21 but the effect size was high, as in previous studies with elderly individuals (Li et al.,  
22 2010; Plummer-D'Amato et al., 2012). Mochizuki et al. (2006) suggested that the  
23 lower values of velocity correspond to higher postural stability; however, in our study,  
24 the Vt with eyes closed increased in the dual-task training group, which may be a  
25 mechanism to compensate for the lower oscillation.

1 Based on the Timed Up and Go Test as well as the UPDRS-III test, the difference in  
2 terms of mobility was higher in the dual-task training group, with medium effect size,  
3 which indicates an improvement of the functional mobility of the individuals. These  
4 findings are consistent with other studies in which the average values were better in  
5 dual-task training programs, but with no significant results (Her et al., 2011; Jiejiao et  
6 al., 2012; Plummer-D'Amato et al., 2012; Vaillant et al., 2006).

7 Regarding the cognitive components, the TMT A, TMT B and RSCardsT results  
8 showed a tendency for improvement in both groups after intervention, likewise in a  
9 previous study by Hiyamizu et al. (2011) with healthy elderly individuals. These  
10 findings are also in agreement with other studies where visible improvements after  
11 dual-task interventions were found, although without statistical significance  
12 (Makizako et al., 2012; Pedroso et al., 2012; Pellecchia, 2005; Silsupadol, Lugade, et  
13 al., 2009).

14 The present study, as far as the authors' know, is innovative as it is the first study to  
15 assess the outcomes of a dual-task intervention on balance and executive functions  
16 in subjects with Parkinson's disease. Nonetheless, there are some limitations that  
17 should be discussed. The small size of the studied sample can limit the results,  
18 particularly regarding the significance of the statistical tests performed and the  
19 generalization of the findings. Hence, this work should be considered as a pilot study  
20 that has added knowledge concerning the effects of dual-task training on balance  
21 and executive functions in patients with PD. All participants involved were "ON"  
22 cholinergic medication, but the effect of the medication on the participants'  
23 performance was not taken into account. Therefore, although the intervention  
24 adopted was selected based on other closely related studies (Silupadol, Lugade, et  
25 al., 2009; Silsupadol, Shumway-Cook, et al., 2009), it is suggested that future studies

1 should also include a cognitive training before or after the balance training in the  
2 group that undergo the single-task training.

3 In conclusion, as was hypothesized for this study, our findings revealed a more  
4 positive response with the dual-task intervention compared to the single-task  
5 intervention. The motor training with a cognitive task performed simultaneously  
6 improved the performance of some parameters related to balance and executive  
7 functions of individuals with Parkinson's disease. These observations highlight the  
8 strength of rehabilitative interventions based on dual-task training.

9

#### 10 **Declaration of interest**

11 The authors report no conflicts of interest. The authors alone are responsible for the  
12 content and writing of the paper.

13

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43

1 **TABLE CAPTIONS**

2

3 Table 1. Comparison at baseline between the single- and dual-task groups.

4

5 Table 2. Comparison between the single- and dual-task groups after the intervention  
6 relatively to baseline.

1 **TABLES**

2 Table 1

	<b>Single-task Group</b>		
	<b>(n=8)</b>	<b>Dual-task Group (n=7)</b>	<b>p-value</b>
Age (years)	62.3 (12.9)	63.4 (9.5)	0.862
Gender, male (%)	6 (85.7%)	5 (62.5%)	0.310 <sup>a</sup>
Education (years)	10.4 (5.1)	8.6 (6.4)	0.288
Physical activity (hours per week)	1.9 (1.3)	1.3 (0.3)	0.208
Body weight (kg)	67.3 (13.5)	66.8 (13.2)	0.817
Height (cm)	168.3 (8.0)	163.9 (7.4)	0.121
Years of disease	7.7 (7.5)	8.8 (4.3)	0.115
Time Up and Go	11.8 (4.4)	11.3 (3.8)	0.798
UPDRS-part III	14.8 (3.9)	14.3 (4.2)	0.795
Eyes opened			
Mediolateral sway (COPx - cm)	0.938 (0.457)	0.813 (0.249)	0.848
Anteroposterior sway (COPy - cm)	1.084 (0.351)	1.120 (0.527)	0.655
Total velocity (Vt-cm/s)	0.513 (0.426)	0.337 (0.082)	0.898
Eyes closed			
Mediolateral sway (COPx - cm)	0.671 (0.248)	0.813 (0.171)	0.949
Anteroposterior sway (COPy - cm)	1.187 (0.473)	1.133 (0.434)	0.137
Total velocity (Vt - cm/s)	0.578 (0.315)	0.538 (0.447)	0.491
RSCardsT	1.71 (1.38)	2.25 (1.49)	0.475
TMT A	86.33 (69.92)	68.75 (28.40)	0.948
TMT B	186.50 (98.78)	168.75 (55.81)	0.439

Results are: mean and (standard deviation) or (%)

3 <sup>a</sup> Chi-square test

1 Table 2

	<b>Single-task Group (n=8)</b>	<b>Dual-task Group (n=7)</b>	<b>p-value</b>	<b>Size Effect</b>
Time Up and Go	-1.800 (1.127)	-2.900 (3.318)	0.620	0.480
UPDRS-part III	-4.833 (3.764)	-7.000 (2.204)	0.345	0.792
Eyes opened				
Mediolateral sway (COPx - cm)	-0.273 (0.325)	-0.145 (0.093)	0.535	0.581
Anteroposterior sway (COPy - cm)	-0.096 (0.366)	-0.273 (0.257)	0.848	0.605
Total velocity (Vt-cm/s)	-0.148 (0.208)	-0.012 (0.091)	0.128	0.922
Eyes closed				
Mediolateral sway (COPx - cm)	0.112 (0.370)	-0.165 (0.114)	0.026*	1.094
Anteroposterior sway (COPy - cm)	-0.341 (0.465)	0.286 (0.479)	0.029*	1.430
Total velocity (Vt - cm/s)	-0.130 (0.365)	0.096 (0.176)	0.181	0.902
RSCardsT	0.286 (0.489)	1.125 (2.031)	0.336	0.590
TMT A	-11.833 (43.190)	-2,750 (15.416)	0.950	0.324
TMT B	-31.333 (48.980)	-0.250 (32.115)	0.345	0.839

Results are: mean and (standard deviation)

\* p-value<0.05

2

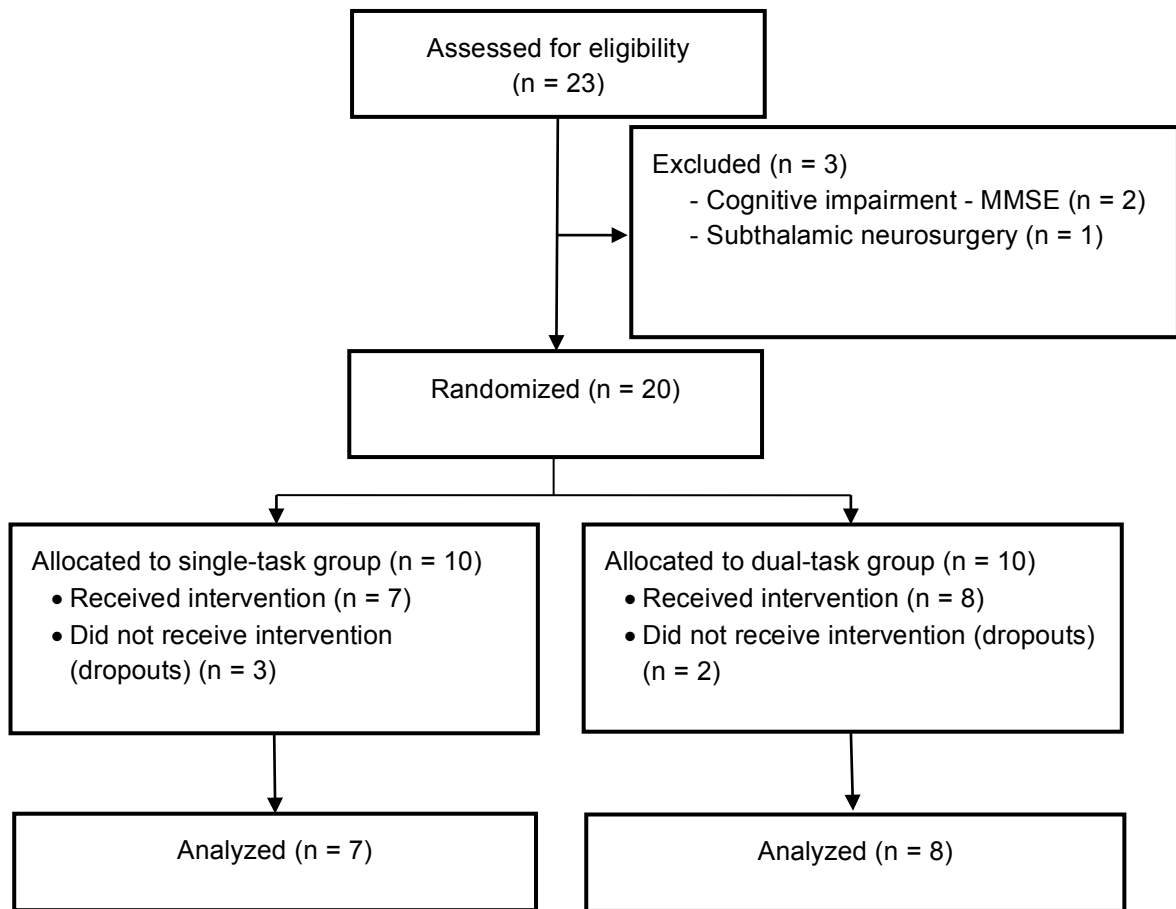
1 **FIGURE CAPTION**

2 Figure 1. CONSORT (Schulz, Altman, & Moher, 2010) diagram of the recruitment  
3 process adopted.

4

1 **FIGURES**

2 **Figure 1**



3