

# Supplementary Material

## A Randomized, Double-Blinded Crossover Trial of Short Versus Conventional Pulse Width Subthalamic Deep Brain Stimulation in Parkinson's Disease

**Supplementary Table 1. Adverse Events**

Event	Number of AE in PW60	Number of AE in PW30	Total number of subjects in both conditions
<i>Adverse Events associated with stimulation</i>			
Dyskinesia	8	8	9
Speech impairment	2	1	3
Worsening of symptom control	1	3	4
<b>Total</b>	<b>11</b>	<b>12</b>	<b>13/30</b>
<i>Adverse Events probably associated with stimulation</i>			
Numbness in a limb	1	0	1
Spasms in fingers or toes	2	1	2
Impairment of balance	1	0	1
<b>Total</b>	<b>4</b>	<b>1</b>	<b>4/30</b>
<i>Adverse Events not associated with stimulation</i>			
Neckpain	1	1	1
Pain in a limb	2	1	3
Dorsal pain	0	2	2
Mild depressive symptoms	5	3	5
Disrupted sleep	0	1	1
<b>Total</b>	<b>8</b>	<b>8</b>	<b>11/30</b>
<i>Serious Adverse Event not associated with stimulation</i>			
DBS system infection	1	0	1/30

Adverse events as reported at follow-up visits or at dropout. AE, adverse event; DBS, deep brain stimulation; PW30, trial condition with 30  $\mu$ s; PW60, trial condition with 60  $\mu$ s.

**Supplementary Table 2. Stimulation Settings at Follow-up Visits**

Subject	Stimulation Setting	Current PW60 [mA]	Current PW30 [mA]
1	R: C+, 3- (100%), 119 Hz	R: 1.8*	R: 3.7
	L: C+, 9- (25%), 10- (25%), 11- (25%), 119 Hz	L: 8.1	L: 8.4
2	R: C+, 1- (10%), 2- (10%), 3- (10%), 4- (24%), 5- (23%), 6- (23%), 130 Hz	R: 2.7	R: 4.0*
	L: C+, 9- (10%), 10- (10%), 11- (10%), 12- (24%), 13- (23%), 14- (23%), 130 Hz	L: 2.3	L: 4.0
3	R: C+, 0- (60%), 1- (20%), 2- (20%), 130 Hz	R: 1.8*	R: 2.9
	L: C+, 8- (60%), 9- (18%), 10- (22%), 130 Hz	L: 2.1	L: 3.1
4	R: C, 1+ (50%), 2- (100%), 3+ (50%), 130 Hz	R: 2.0	R: 3.0*
	L: C+, 9- (30%), 11- (30%), 12- (40%), 174 Hz	L: 1.7	L: 3.0
5	R: C+, 1- (34%), 2- (33%), 3- (33%), 116 Hz	R: 2.2	R: 4.3*
	L: C+, 11- (50%), 14- (50%), 116 Hz	L: 3.1	L: 5.0
6	R: C+, 1- (34%), 2- (33%), 3- (33%), 130 Hz	R: 3.0*	R: 4.3
	L: C+, 9- (34%), 10- (33%), 11- (33%), 130 Hz	L: 3.4	L: 4.9
7	R: C+, 1- (34%), 2- (33%), 3- (33%), 130 Hz	R: 2.5	R: 4.0*
	L: C+, 12- (34%), 13- (33%), 14- (33%), 130 Hz	L: 1.5	L: 2.8
8	R: C+, 7- (100%), 130 Hz	R: 0.2*	R: 0.2*
	L: C+, 15- (100%), 130 Hz	L: 1.2	L: 1.0
9	R: C+, 4- (34%), 5- (33%), 6- (33%), 130 Hz	R: 0.7	R: 1.1*
	L: C+, 12- (34%), 13- (33%), 14- (33%), 130 Hz	L: 0.7	L: 0.7
10	R: C+, 4- (34%), 5- (33%), 6- (33%), 119 Hz	R: 2.9*	R: 5.8
	L: C+, 12- (50%), 13- (50%), 119 Hz	L: 2.9	L: 6.3
11	R: C+, 4- (100%), 130 Hz	R: 0.7	R: 1.4
	L: C+, 9- (100%), 130 Hz	L: 1.9	L: 3.3
12	R: C+, 4- (33%), 5- (34%), 6- (33%), 130 Hz	R: 2.1*	R: 4.6
	L: C+, 12- (10%), 13- (10%), 14- (10%), 15- (70%), 130 Hz	L: 2.6	L: 4.5
13	R: C+, 1- (16%), 2- (16%), 4- (33%), 5- (35%), 185 Hz	R: 3.2	R: 4.9*
	L: C+, 10- (13%), 11- (15%), 13- (36%), 14- (36%), 185 Hz	L: 2.7	L: 3.8
14	R: C+, 4- (31%), 5- (42%), 6- (27%), 130 Hz	R: 2.3	R: 3.3*
	L: C+, 12- (42%), 13- (42%), 14- (16%), 130 Hz	L: 2.3	L: 3.3
15	R: C+, 4- (34%), 5- (33%), 6- (33%), 130 Hz	R: 2.2*	R: 2.8
	L: C+, 12- (34%), 13- (33%), 14- (33%), 130 Hz	L: 2.4	L: 3.3
16	R: C+, 3- (100%), 204 Hz	R: 2.7	R: 4.0*
	L: C+, 12- (34%), 13- (33%), 14- (33%), 204 Hz	L: 0.8	L: 1.0
17	R: C+, 4- (33%), 5- (34%), 6- (33%), 198 Hz	R: 1.5	R: 2.6*
	L: C+, 12- (34%), 13- (33%), 14- (33%), 198 Hz	L: 1.3	L: 1.9
18	R: C+, 1- (34%), 2- (33%), 3- (33%), 130 Hz	R: 1.2*	R: 2.4
	L: C+, 9- (50%), 10- (50%), 130 Hz	L: 1.9	L: 2.9
19	R: C+, 1- (18%), 2- (16%), 3- (16%), 4- (18%), 5- (16%), 6- (16%), 130 Hz	R: 2.6*	R: 3.4
	L: C+, 12- (34%), 13- (33%), 14- (33%), 130 Hz	L: 2.6	L: 3.1
20	R: C+, 2- (34%), 3- (33%), 4- (33%), 130Hz	R: 2.4	R: n.a.
	L: C+, 10- (34%), 11- (33%), 12- (33%), 130Hz	L: 2.9	L: n.a.
21	R: C+, 4- (100%), 130Hz	R: n.a.	R: 1.1
	L: C+, 9- (100%), 130Hz	L: n.a.	L: 1.7
22	R: C+, 5- (34%), 6- (33%), 7- (33%), 179Hz	R: n.a.	R: 4.6
	L: C+, 13- (50%), 15- (50%), 179Hz	L: n.a.	L: 6.4
23	R: C+, 5- (34%), 6- (33%), 7- (33%), 130Hz	R: n.a.	R: 3.1
	L: C+, 13- (34%), 14- (33%), 15- (33%), 130Hz	L: n.a.	L: 4.0
24	R: C+, 5- (34%), 6- (33%), 7- (33%), 174Hz	R: n.a.	R: 0.6
	L: C+, 10- (18%), 11- (16%), 12- (16%), 13- (18%), 14- (16%), 15- (16%), 174Hz	L: n.a.	L: 3.5

Asterisk marks the patient's preferred stimulation setting. C, Case; L, left hemisphere; n.a., not available due to dropout; PW30, trial condition with 30  $\mu$ s; PW60, trial condition with 60  $\mu$ s; R, right hemisphere.

**Supplementary Table 3. Motor Diary Outcomes per period**

Motor States [h]	Period A N = 24	Period B N = 19	LME Period A vs. Period B	
	Mean (SD), [90%-CI]	Mean (SD), [90%-CI]	Estimate (SE)	p
“On”	11.95 (4.34) [10.4 to 13.5]	10.65 (4.92) [8.7 to 12.6]	-1.39 (0.68)	0.2
“Off”	2.18 (3.14) [1.1 to 3.3]	2.76 (3.51) [1.4 to 4.2]	0.97 (0.52)	0.3
“Dyskinesia”	1.57 (3.19) [0.5 to 2.7]	2.14 (3.71) [0.7 to 3.6]	0.36 (0.38)	1.0
“Sleep”	8.41 (1.29) [7.9 to 8.9]	8.39 (1.64) [7.7 to 9.0]	-0.04 (0.18)	1.0

Comparison of motor diary outcomes between Period A and Period B (see Fig. 1).  
90%-CI, 90% confidence interval; SD, standard deviation; SE, standard error.

**Supplementary Table 4. Motor Diary Outcomes for paired data only**

Motor States [h]	PW60 N = 19	PW30 N = 19	PW60 vs. PW30 N = 19	
	Mean (SD), [90%-CI]	Mean (SD), [90%-CI]	Estimate (SE)	p
“On”	11.68 (4.19) [10.0 to 13.3]	11.04 (5.05) [9.0 to 13.1]	-0.39 (3.21)	1.0
“Off”	2.02 (3.67) [0.9 to 3.0]	1.93 (3.55) [1.1 to 3.8]	1.16 (2.27)	1.0
“Dyskinesia”	1.95 (2.60) [0.6 to 3.5]	2.49 (3.38) [0.5 to 3.3]	1.89 (1.67)	1.0
“Sleep”	8.30 (1.39) [7.7 to 8.9]	8.53 (1.57) [7.9 to 9.2]	-1.37 (0.67)	0.2

Comparison of motor diary outcomes between PW60 and PW 30 only including patients completing the trial. 90%-CI, 90% confidence interval; SD, standard deviation; SE, standard error; PW30, trial condition with 30  $\mu$ s; PW60, trial condition with 60  $\mu$ s.

**Supplementary Table 5. Subgroup analysis: Comparison of Non-preferred Settings**

	npPW60 (N = 9)		npPW30 (N = 9)		p
	Mean	SD	Mean	SD	
<i>Motor states [h]</i>					
“On”	11.95	4.38	10.35	4.30	1.00
“Off”	1.96	3.00	3.83	3.85	0.955
“Dyskinesia”	2.20	3.03	0.83	1.65	1.00
“Sleep”	7.78	1.20	8.98	2.01	0.562
<i>Secondary outcomes</i>					
VAS “ability to speak”	3.77	3.06	5.07	2.28	1.00
Intelligibility	8.26	1.40	7.79	1.10	1.00
VAS “ability to walk”	5.12	2.60	5.16	2.77	1.00
10-m timed walk [s]	8.20	1.56	9.78	1.37	0.80
UPDRS-I	1.22	1.20	1.89	2.42	1.00
UPDRS-II	8.44	3.88	8.22	2.91	1.00
UPDRS-III	18.67	7.52	16.33	6.61	1.00
PDQ39-SI	58.12	40.42	74.36	62.74	1.00
NMSQ	6.78	3.15	8.22	5.02	1.00
<i>Energy indices</i>					
Change in current (PW60→PW30) [%]	38.23	41.05	31.87	55.53	1.00
Battery charge index [ $\mu$ J]	40.09	24.06	42.49	29.79	1.00
Charge per pulse [nC]	226.00	87.05	254.00	130.06	1.00
Total electrical energy delivered [ $\mu$ J/s]	76.41	54.05	83.39	64.20	1.00

npPW60, 60  $\mu$ s condition of patients preferring 30  $\mu$ s setting; npPW30, 30  $\mu$ s condition of patients preferring 60  $\mu$ s setting; SD, standard deviation