

## Supplementary document

### Beneficial nonmotor effects of subthalamic and pallidal neurostimulation in Parkinson's disease

#### 1. Results

##### 1.1. *Baseline characteristics in the original cohort*

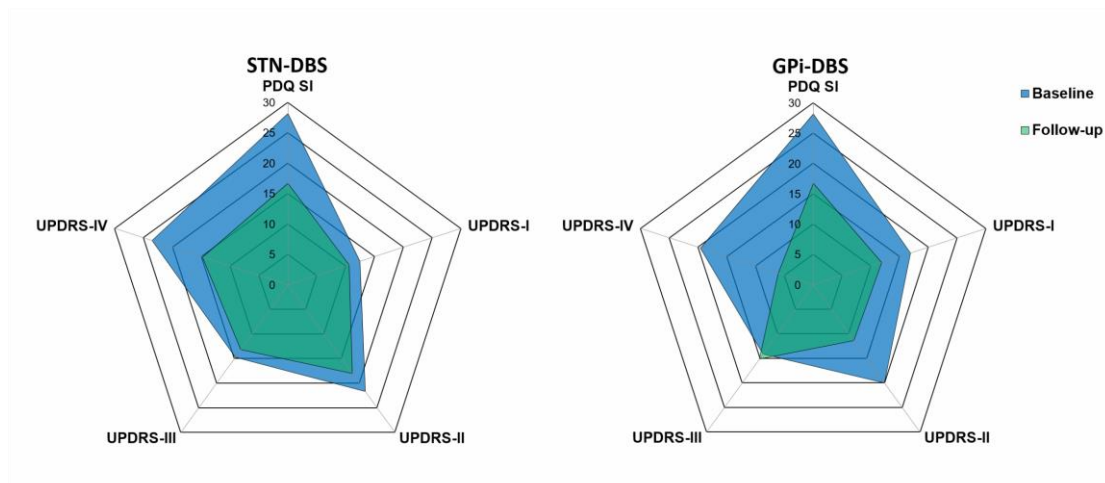
In the original cohort 60 patients with PD were included. Demographic characteristics were similar, Hoehn and Yahr Staging was slightly higher in the GPi-DBS group than in the STN-DBS group, and preoperative UPDRS-III MedON scores were significantly higher in the STN-DBS compared to the GPi-DBS group ( $p=0.023$ ).

##### 1.2. *Changes of outcomes at follow-up in the original cohort*

In both groups of the original cohort, PDQ SI, UPDRS-II, UPDRS-IV, Schwab and England Scale, and NMSS improved significantly from baseline to follow-up (for STN-DBS all  $p<0.001$ , for GPi-DBS all  $p<0.01$ ) (Table e-1). As expected, LEDD reduction was approximately 51% in the STN-DBS group ( $p<0.001$ ) and only approximately 10% in the GPi-DBS group ( $p=0.357$ ). This difference of LEDD reduction was significant (corrected  $p<0.001$ ). LEDD reduction was not significantly correlated to other outcome parameters. Post-hoc analyses of NMSS domain scores in the STN-DBS group, resulted in significant improvements of sleep/fatigue ( $p=0.002$ ), mood/cognition ( $p=0.017$ ), perceptual problems/hallucinations ( $p=0.007$ ), urinary symptoms ( $p=0.012$ ), sexual function ( $p=0.017$ ), and the miscellaneous domain ( $p=0.003$ ), which was driven by the items pain (baseline:  $4.5 \pm 4.3$ ; follow-up:  $2.4 \pm 3.6$ ;  $p=0.032$ ) and olfaction (baseline:  $3.9 \pm 3.7$ ; follow-up:  $1.7 \pm 2.8$ ;  $p=0.017$ ). In the GPi-DBS group, sleep/fatigue ( $p=0.004$ ), mood/cognition ( $p=0.001$ ), and sexual function ( $p=0.015$ ) improved significantly at follow-up.

In the STN-DBS group, effect sizes (Table e-2) were 'small' for UPDRS-I and -III, 'moderate' for UPDRS-II, -IV, and NMSS, and 'large' for PDQ SI, Schwab and England Scale, and LEDD reduction. In the GPi-DBS group, effect sizes were 'small' for UPDRS-I, Hoehn and Yahr scale, and LEDD reduction, 'moderate' for Schwab and England Scale, and 'large' for PDQ SI, UPDRS-II, -IV, and NMSS. NNT results were favorable for STN-DBS regarding LEDD reduction, UPDRS-I, -III, and Schwab and England Scale (Table e-3). On the other hand, NNT results were better in GPi-DBS for UPDRS-II, -IV, Hoehn and Yahr Scale, PDQ SI, and NMSS. In both targets, PDQ SI change scores were significantly correlated to UPDRS-II and Schwab and England Scale change scores (all  $p < 0.05$ ) (Table e-4) and in STN-DBS, a significant correlation was also observed for NMSS change scores ( $p = 0.002$ ).

**Supplementary figure – PDQ SI and UPDRS at baseline and follow-up in the matched cohort for subthalamic and pallidal stimulation**



**Abbreviations:** GPi-DBS = Pallidal deep brain stimulation; PDQ SI = Parkinson's Disease Questionnaire; STN-DBS = Subthalamic stimulation, UPDRS = Unified Parkinson's Disease Rating Scale

**Supplementary figure** illustrates PDQ SI and UPDRS domains at baseline (blue) and follow-up (green) for the STN-DBS and GPi-DBS groups in radar charts.