

Follow-up Programming and Troubleshooting

Jinyoung Youn, MD, PhD

Department of Neurology, Samsung Medical Center, Sungkyunkwan
University School of Medicine, Seoul, Korea

Disclosure

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1. Follow-up Programming in PD patients with DBS
2. Approach to the troubleshooting cases in PD patients with DBS

Routine follow-up patients

- Checking the system
 - Impedance
 - Battery and charging
 - Usage of programs
- Identify problems with history-taking and examination
 - Adjusting medication vs. DBS parameters
 - For DBS parameters: amplitude, frequency, pulse width, contact
 - Always consider the adverse effect of DBS

Device issues

- Impedance: abrupt worsening of parkinsonian symptoms which could be unilateral or bilateral
- Battery: especially to refer to the surgeon at the right time for the battery replacement
- Charging: to confirm whether the patients charged well without any problem
- Usage of programs: to figure out which stimulation parameter is better

Routine follow-up patients

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 - Adjusting medication vs. DBS parameters
 - For DBS parameters: amplitude, frequency, pulse width, contact
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Identify problem

Doctor's	Patient's	Care-giver's
<ol style="list-style-type: none">1. Wearing Off2. Off freezing3. MCI4. Dyskinesia	<ol style="list-style-type: none">1. Off pain2. Depression3. Constipation4. Wearing off	<ol style="list-style-type: none">1. ICD2. Depression3. MCI4. Dyskinesia

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Great response from DBS

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Good response from DBS

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Minimal response from DBS

Identify problem

- Motor symptoms
 - Appendicular sign, asymmetry
 - Axial symptoms and gait disorders
 - Fluctuations and dyskinesia
 - Non-motor symptoms
 - Hypomania
 - Impulse control disorders and DDS
 - Cognitions
- Parkinsonian symptoms (levodopa-responsive or not) vs. stimulation-induced side effects vs. unrelated symptoms

Response to Levodopa

- If the patient's problem is responsive to levodopa, increasing stimulation (usually amplitude) can be beneficial
- Also, fluctuating symptoms with medication can be better with stimulation
- Although tremor did not respond to levodopa, tremor can be better with stimulation. Especially high frequency stimulation at the dorsal contact could be helpful

Identify problem

Doctor's	Patient's	Care-giver's
<ol style="list-style-type: none">1. Wearing Off2. Off freezing3. MCI4. Dyskinesia	<ol style="list-style-type: none">1. Off pain2. Depression3. Constipation4. Wearing off	<ol style="list-style-type: none">1. ICD2. Depression3. MCI4. Dyskinesia

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Adjust DBS or medication

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Adjust DBS or medication
Antidepressant, laxatives

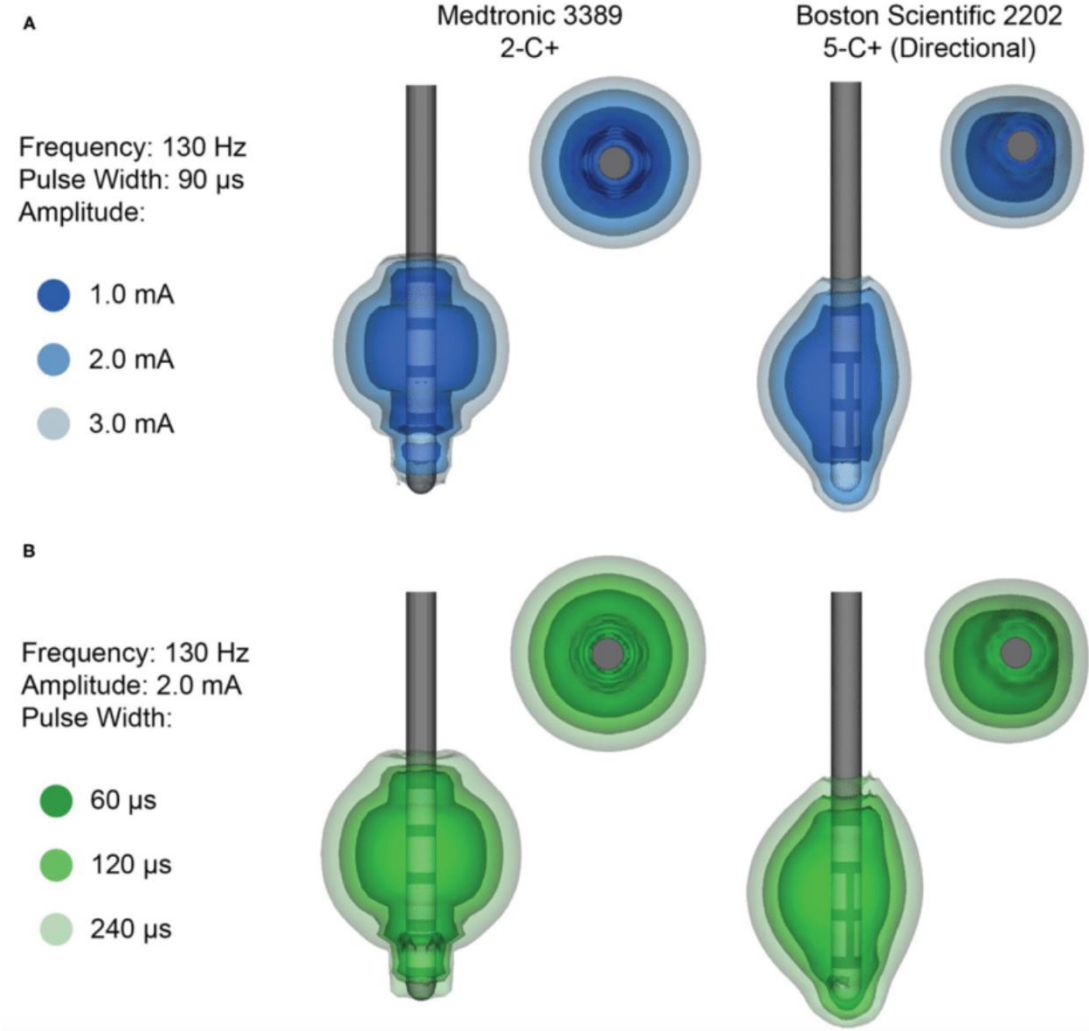
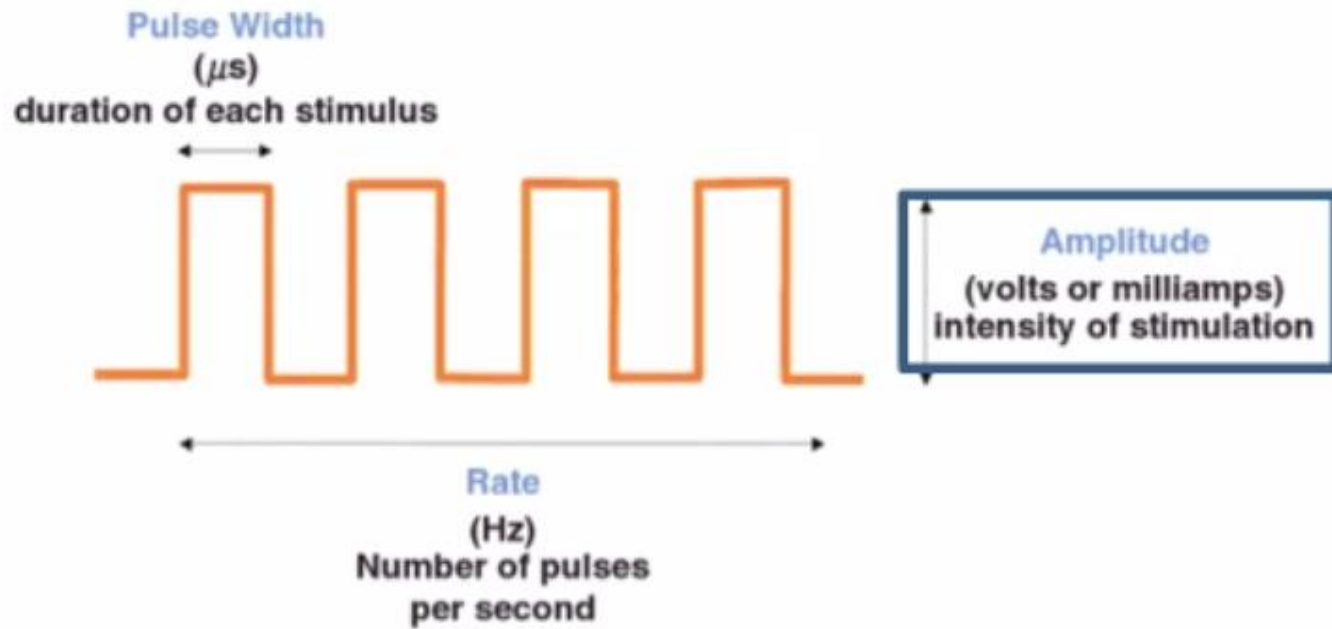
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Decrease DA or other dopaminergic medication

Increasing medication vs. stimulation

- There is no absolute rule to decide increasing medication or stimulation
- For tremor, increasing stimulation can be a better option
- For fluctuation, increasing stimulation with/without decreasing medication can be better than medication adjustment
- For asymmetry, adjusting stimulation is preferred
- Decisions should be made based on age, side effects, battery consumption, ADL and QoL
- Also, the target for the stimulation amplitude or LEDD can vary depending on the patients

Adjusting stimulation



Frey J et al. 2022

Take Home Message (follow-up patients)

- Check the device always, but especially in patients with aggravated parkinsonism
- Patient's complaints could be different from the problems from the physician's point of view
- Definite patient's complaints or problems based on motor or non-motor symptoms responsive or non-responsive to levodopa, stimulation-induced side effects and non-related symptoms to PD
- Adjust parameters amplitude > frequency > pulse width > contacts in outpatients clinic, but you can review the best contact based on image-guided program or local field potential nowadays

Troubleshooting

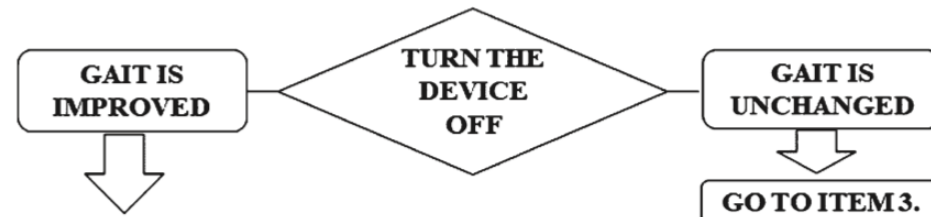
- Almost the same algorithm with managing follow-up patients, but more focused on dealing with chief complaints related to parkinsonian symptoms or stimulation-induced adverse effects

Lessons from gait problem after DBS

1.OBSERVE THE PATIENT AND IDENTIFY THE PROBLEM



2.MIND THE GAP: THE INTERNAL CAPSULE.

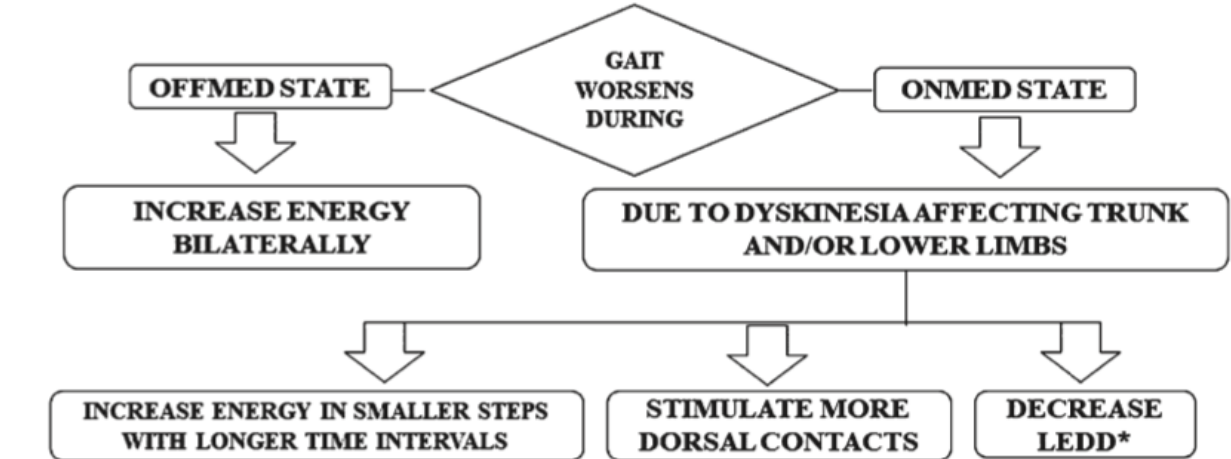


TURN ON EACH ELECTRODE SEPARATELY AND FIND THE "GUILTY" ONE

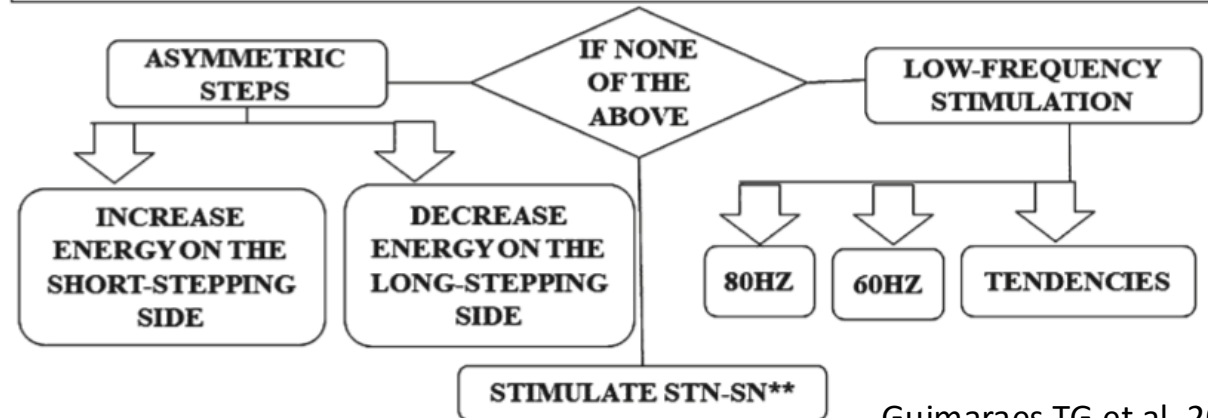
USE STRATEGIES TO AVOID THE SPREAD OF ENERGY OUTSIDE THE NUCLEUS

- Reduce amplitude
- Use bipolar stimulation
- Use interleaving stimulation

3.DETERMINE RESPONSIVENESS TO LEVODOPA



4.FINE TUNE ADJUSTMENTS



Guimaraes TG et al. 2022

Functional dystonia referred to DBS clinic

TABLE 1 Demographics and clinical classification of FMD patients who were referred for functional neurosurgery (DBS and ablative procedures)

Phenomenology	Functional dystonia	Functional tremor	Functional myoclonus	Functional tourettism	All phenomenologies
No. (% of referrals with similar phenomenology)	6 (9)	3 (1.8)	1 (50)	1 (17)	11 (4.7)
Total DBS referrals for similar phenomenology	67	161	2	6	236 ^a
Age (y ± SD)	44 ± 16.7	61 ± 2.2	65	23	48.8 ± 17.3
Sex (F:M)	4:2	1:2	1:0	1:0	7:3
Disease duration (y ± SD)	7.8 ± 12.2	9.3 ± 7.6	2	13	8.4 ± 10.3
Diagnostic classification ^b % (n)					
• Clinically definite	50% (3)			100% (1)	36% (4)
• Clinically established plus other features	33% (2)	33% (1)			27% (3)
• Laboratory supported definite ^c	17% (1)	67% (2)	100% (1)		36% (4)

Gorodetsky C et al. 2022

Step 3. Review where we are stimulating

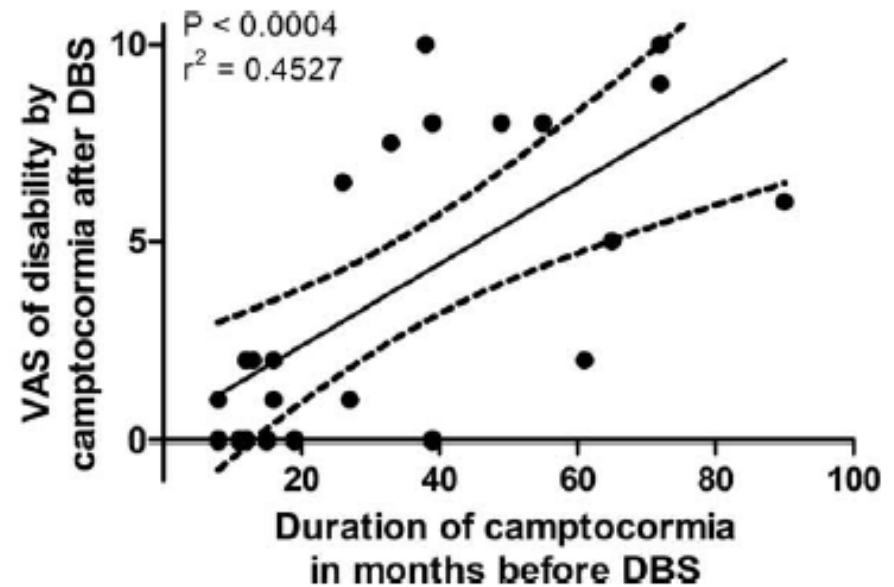
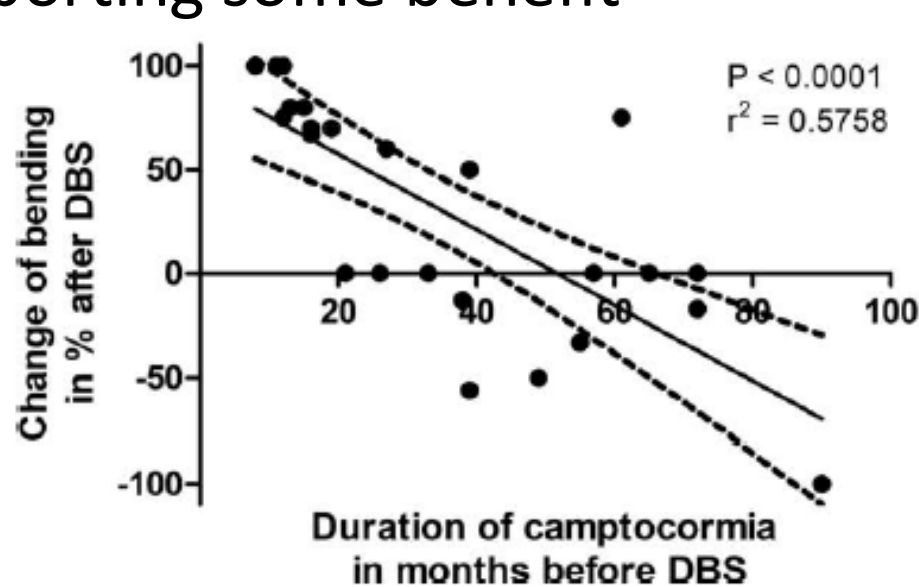
- To offer optimal spots for the stimulation, it is important to know where we are stimulating
- Directional stimulation could be useful to avoid sour spot and stimulate sweet spot

Step 4. Check response to levodopa

- When the problem responds to levodopa, there's more chance to improve with stimulation adjustment
- Also, reviewing where we stimulate is important and Image-guided programming or review of local field potential could be helpful to find the sweet spot

Response to Levodopa

- Axial symptoms, which may not respond to levodopa, usually do not improve with increasing stimulation, even with some studies reporting some benefit



Schulz-Schaeffer WJ et al. 2015

Step 5. Apply Advanced Stimulation

- When the problem responds to levodopa, there's more chance to improve with stimulation adjustment
- Also, reviewing where we stimulate is important and Image-guided programming or review of local field potential could be helpful to find the sweet spot

Low frequency stimulation

Low frequency stimulation

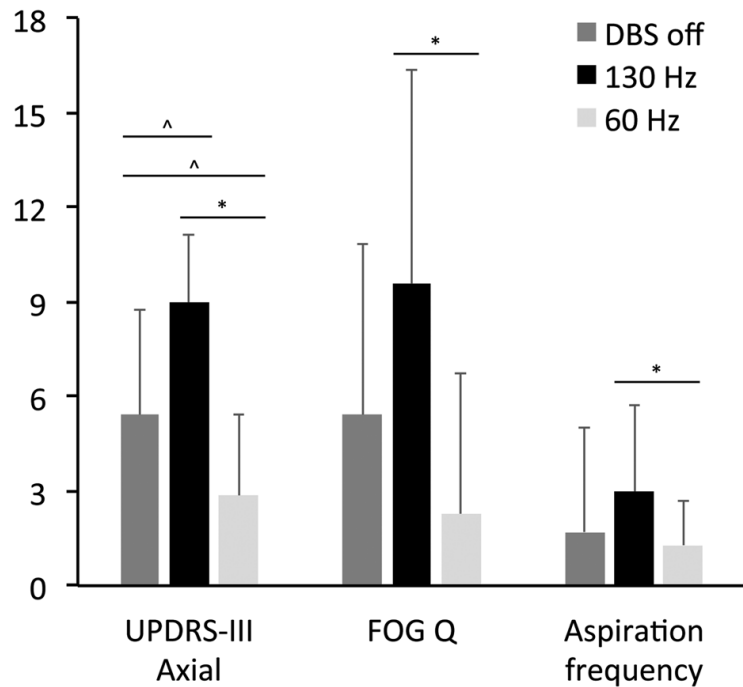
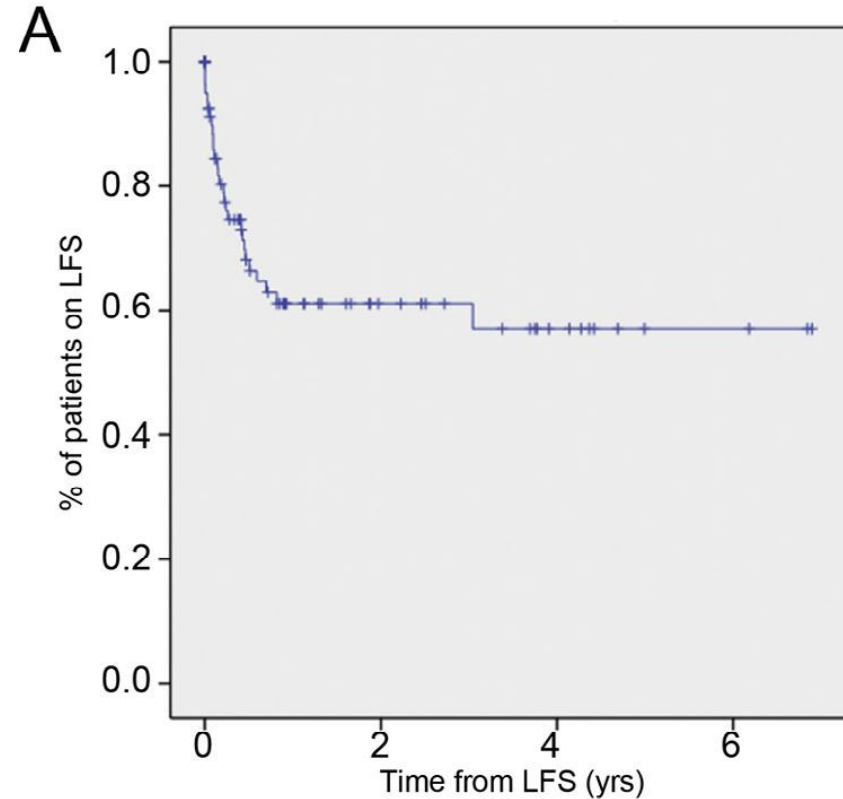


FIG. 1. Data from Xie et al.²² presented as mean and standard deviation during DBS off, HFS (130 Hz) and LFS (60 Hz). Abbreviations: * = $p < 0.05$; ^ = $p < 0.05$ (Bonferroni correction); FOG Q freezing of gait

Watch out appendicular sign



Biase L, Fasano A. 2016, Zibetti M et al. 2016

Combined with Nigral Stimulation

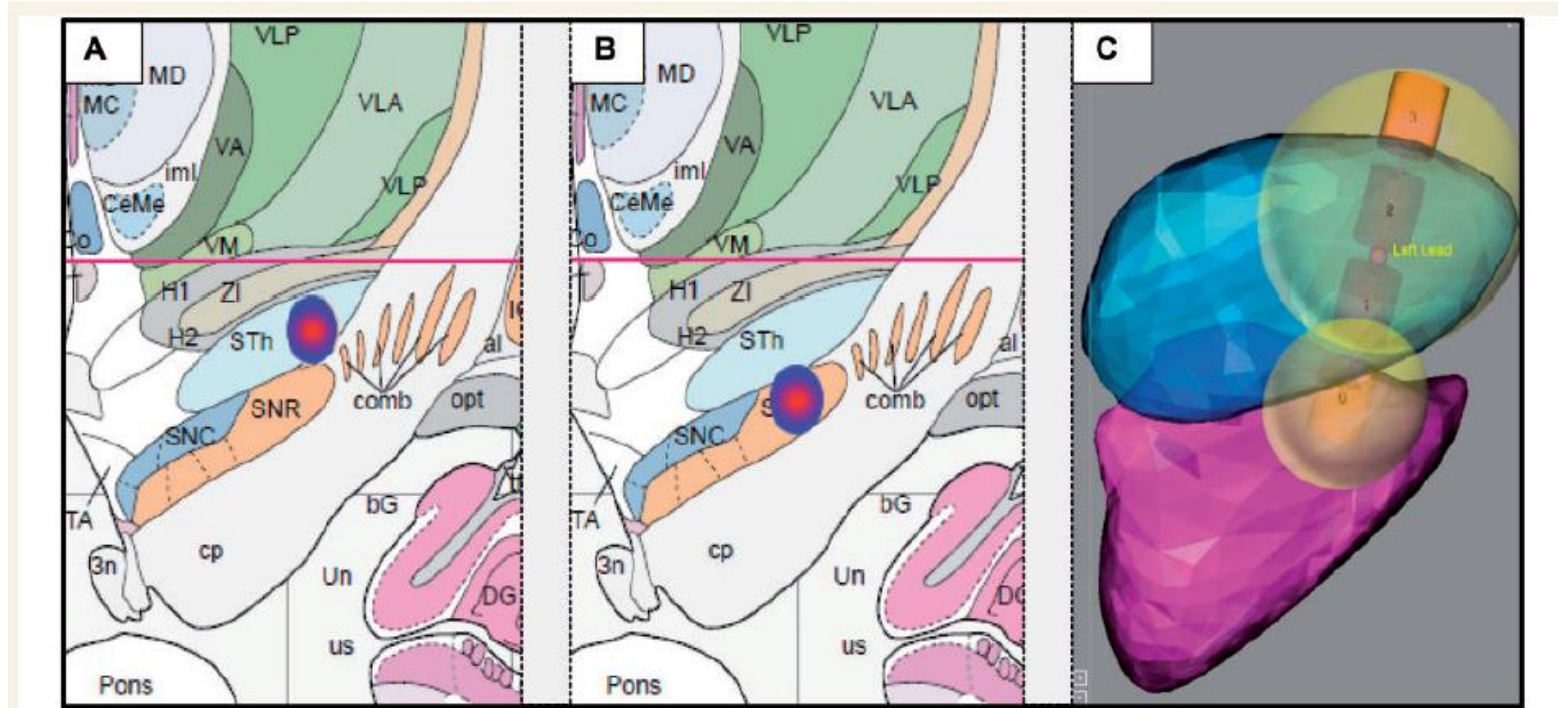


Figure 2 Localization of active electrode contacts of (A) dorsolateral STN and (B) dorsolateral SNr. Coordinates relative to the mid-commissural point (MCP) were: left STN -11.4 ± 0.8 , -0.9 ± 2.0 , -3.0 ± 1.7 ; right STN 13.5 ± 1.1 , -0.5 ± 1.7 , -2.2 ± 1.5 ; left SNr -10.0 ± 0.9 , -3.4 ± 2.1 , -6.4 ± 1.8 ; right SNr 12.1 ± 1.3 , -3.3 ± 1.7 , -5.8 ± 1.5 (x, y, z; x = medio-lateral, y = antero-posterior, z = rostro-caudal). Electrode coordinates (mean \pm standard deviation in x- and y-direction) are visualized in coronal view on the Atlas of the Human Brain with permission (Mai *et al.*, 2007). (C) An additional illustrative image of electrode localization including a simulation on volume of tissue activated was kindly provided by Medtronic based on work by Yelnik *et al.* (2007) (atlas) and D'Haese *et al.* (2012) (atlas and algorithms).

Weiss D et al. 2013

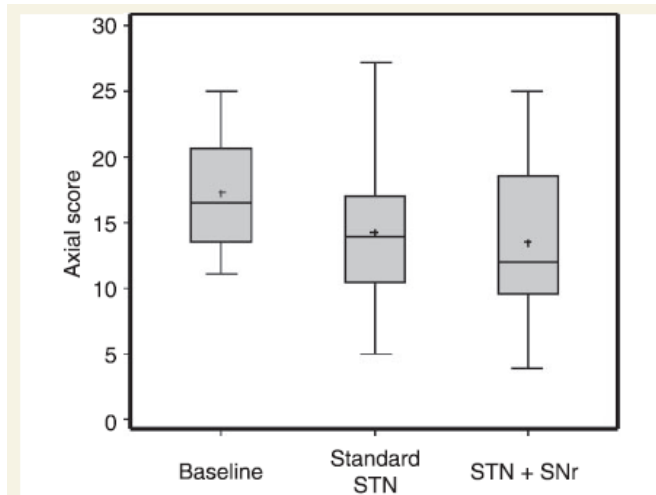


Figure 3 Primary endpoint at '3-week follow-up'. Results are given as box plots. x-axis: therapeutic condition; y-axis: axial score. [STN + SNr] = combined STN + SNr stimulation.

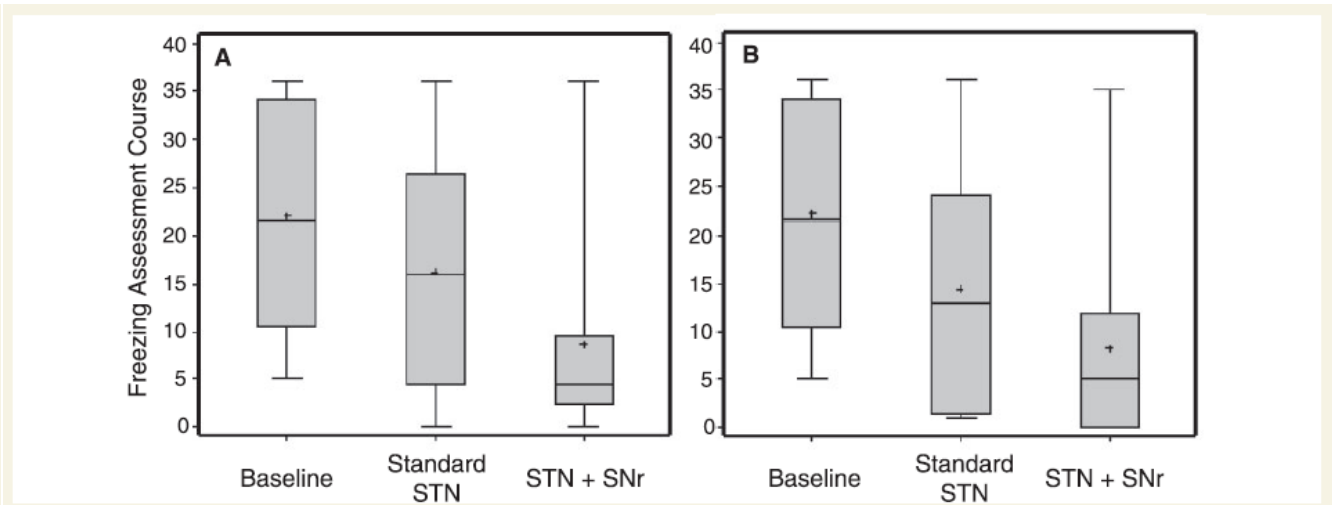


Figure 4 Secondary endpoint: results at (A) 'immediate testing' and at (B) '3-week follow-up' are given for the Freezing of Gait Assessment Course. Results are given as box plots. x-axis: therapeutic condition; y-axis: score of the Freezing of Gait Assessment Course. [STN + SNr] = combined STN + SNr stimulation.

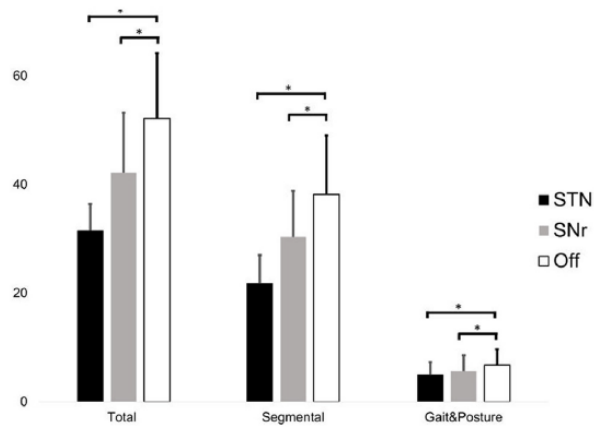


FIGURE 1 | Score of the total UPDRS III (left), segmental score (sum of items 20–26 + 31, only upper and lower limbs), and gait and posture subscore (sum of items 27–30) during subthalamic nucleus (STN), substantia nigra pars reticulata (SNr), and Off stimulation. Significant differences ($p < 0.05$) are denoted by horizontal square brackets. Both STN and SNr stimulations could significantly improve the segmental and gait and posture subscore.

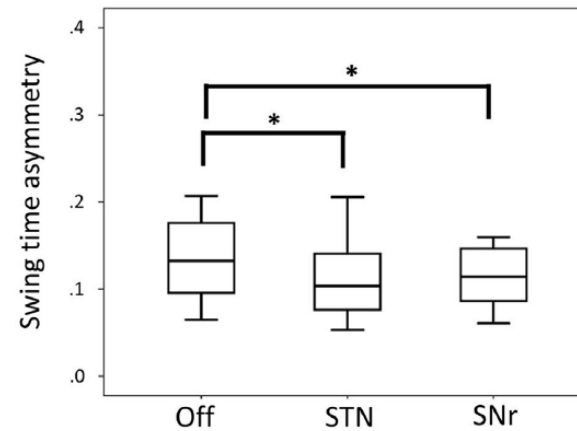
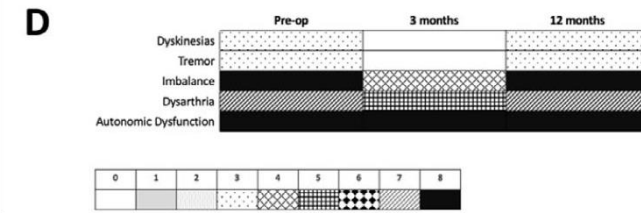
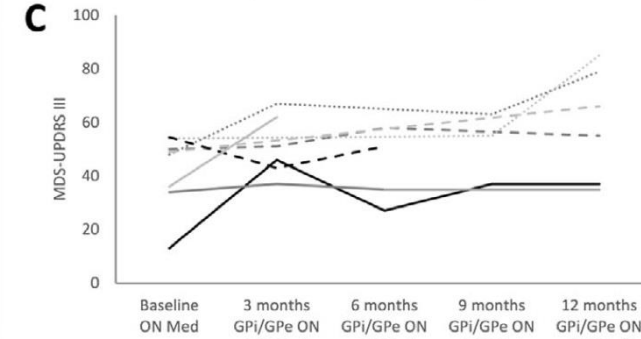
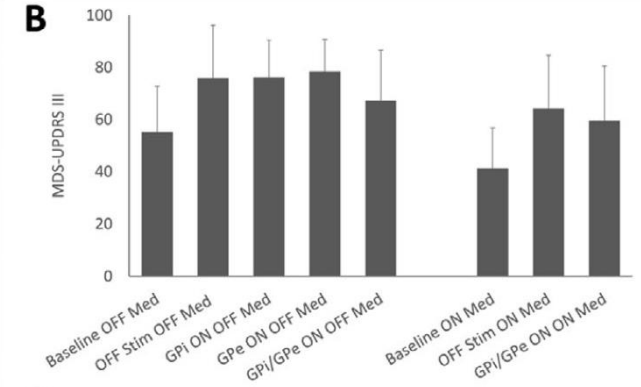
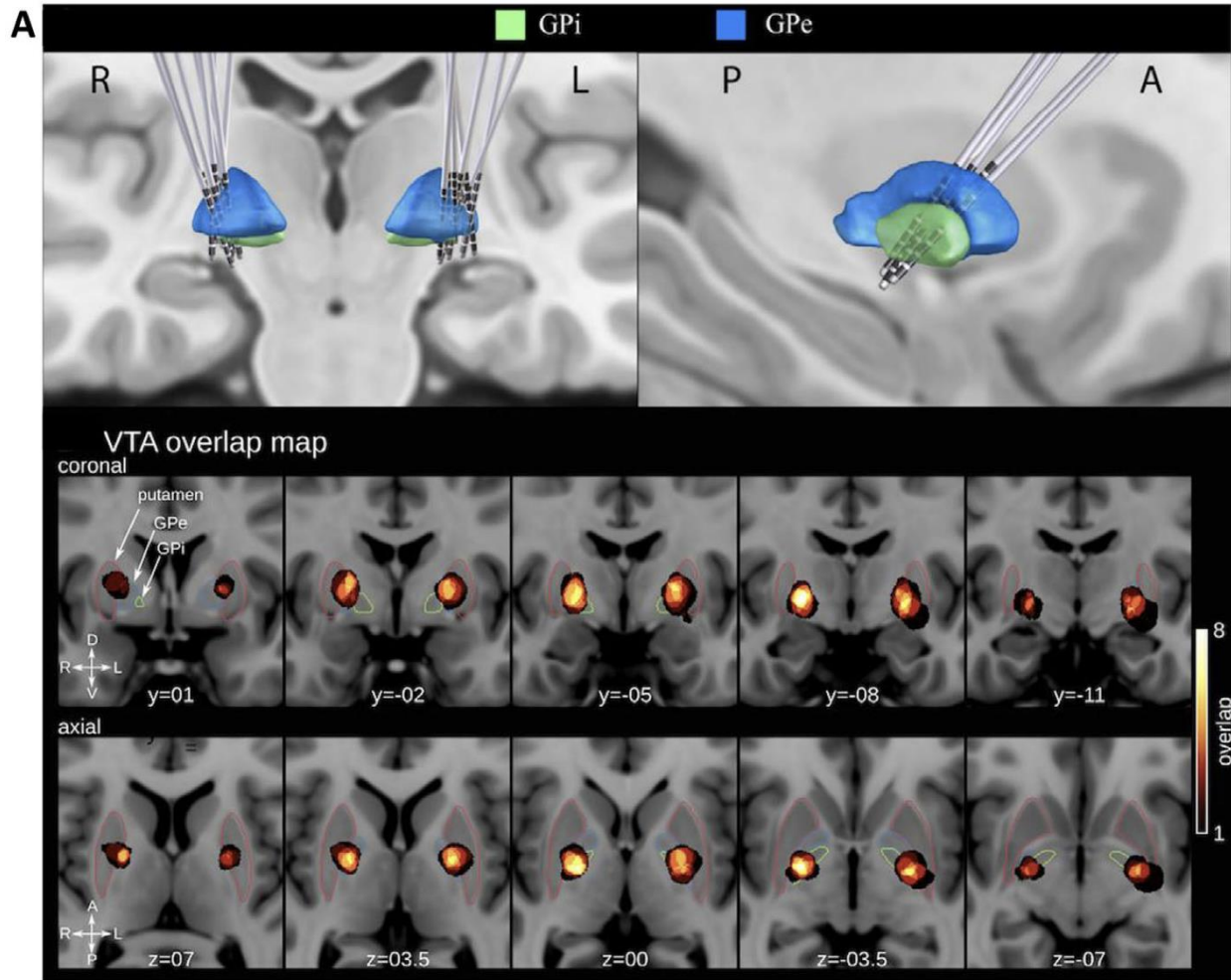


FIGURE 2 | Boxplot representing median values, 25–75% range (box) and min–max range (bars) of swing time asymmetry. Differences were computed with the Wilcoxon signed rank test and are denoted by horizontal square brackets. Abbreviations: STN, subthalamic nucleus deep brain stimulation; SNr, substantia nigra deep brain stimulation; ll, leg length; CV, coefficient of variation.

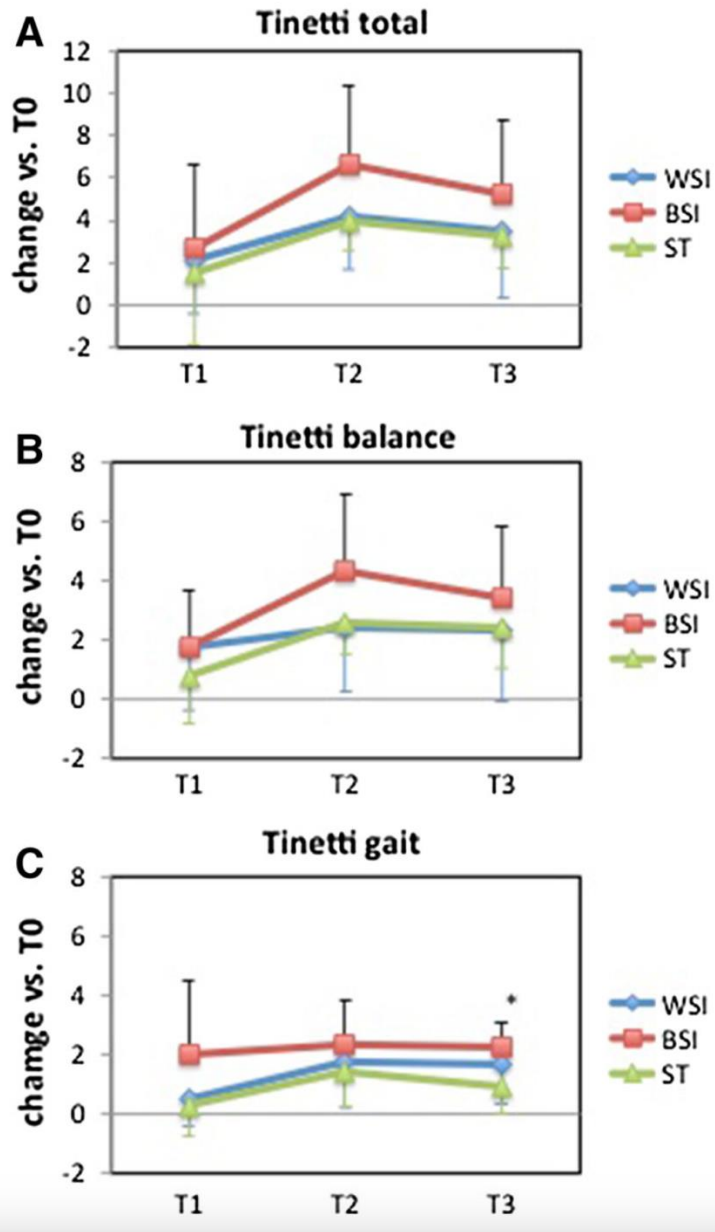
Weiss D et al. 2013, Scholten N et al. 2017

Additional GPe stimulation

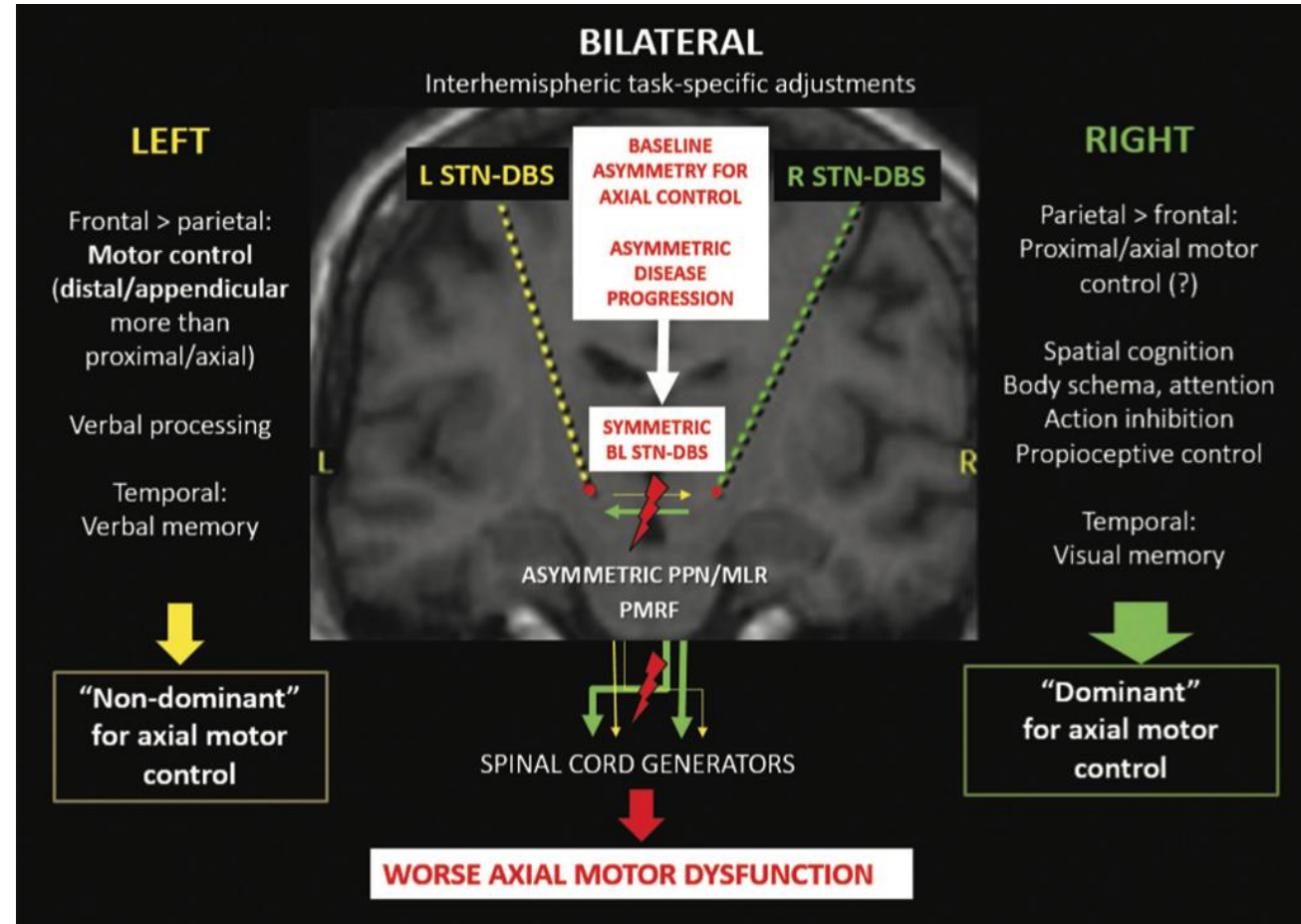


Di Luca D et al. 2023

Asymmetry



Right side stimulation



Ricciardi L et al. 2015, Lizzaraga et al. 2017

Take Home Message (Troubleshooting)

- Identify the problem, first
- Always consider capsular side effects
- Response to more stimulation and levodopa is important
 - Some symptoms can be seen with peak-dose (ON FOG, uncontrolled dyskinesia)
- Changing contacts based on the review of stimulation site
 - Image-guided programming or the review of local field potentials can be helpful and maximize the benefit of directional stimulation
- Low frequency stimulation could be helpful for the axial symptoms
 - There is a risk for worsening appendicular sign
- For gait issues, asymmetry or focusing on right hemisphere can be also considered