

# Results of Deep Brain Stimulation for Parkinson's Disease after 30 Cases

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## ABSTRACT

**Introduction:** Deep brain stimulation of the subthalamic nucleus (STN) is an effective therapy for medically refractory Parkinson's disease leading to significant improvement of Parkinsonian symptoms through functional inhibition of the STN.

**Aim of this study:** To analyze the outcome of bilateral subthalamic nucleus, deep brain stimulation in advanced Parkinson's disease patients. This is a clinical observational study.

**Material and methods:** This is the result of bilateral subthalamic nucleus-deep brain stimulation (STN-DBS) done in 30 patients for advanced Parkinson's disease in the Institute of Neurosciences, Kolkata, during the past 7 years (2013 to 2019, August) by the authors team. Outcome had been analyzed.

**Results:** Excellent outcome was found after the required programming. Ninety percent patients have shown excellent result. The dosage of antiparkinsonian medications was significantly reduced, with a consequent reduction of dyskinesias.

**Conclusion:** The effect of the STN-DBS on the motor fluctuations and on the levodopa-induced dyskinesias led to a significant improvement of motor part of Unified Parkinson Disease Rating Scale (UPDRS [III]) rating.

**Keywords:** Deep brain stimulation, Parkinson's disease, Subthalamic nucleus-deep brain stimulation.

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## OBJECTIVES

Hyperactivity of subthalamic nucleus (STN) plays an important role in the pathophysiology of Parkinson's disease. Through chronic high-frequency electrical stimulation, it is possible to achieve functional inhibition of STN leading to improvement of Parkinsonian symptoms and significant reduction of dopaminergic drugs with an improvement of drug-induced dyskinesias.<sup>1-5</sup>

The aim of this study was to analyze the outcome of the bilateral subthalamic nucleus-deep brain stimulation (STN-DBS) for Parkinson's disease during 2013 to 2019 in the Institute of Neurosciences, Kolkata, by the authors team.

## MATERIALS AND METHODS

Total number of patients—30 (advanced Parkinson's disease).

Sex—20 males, 10 females.

Age—51–70 years.

Average duration of disease—10 years.

### Preoperative UPDRS (Part III)

Average score—19 (on medicine), 56 (off medicine).

Average preoperative levodopa dosage—956 mg/day.

Average duration of motor fluctuations—5 years.

Average duration of dyskinesia—4 years.

### Predominant Symptoms

Rigidity and hypokinesia—12.

Tremor—8.

Severe dyskinesia—10.

### Patient Selection Methods

There is a 10-point criteria chart which needs to be "YES" for all except points 8 and 9 (Table 1).<sup>4</sup>

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**Conflict of interest:** None

**Table 1:** 10-point criteria for case selection

1	Age < 75 years	Yes
2	Idiopathic PD (no PSP/MSA/CBD/LBD, etc.)	Yes
3	Levodopa responsive	Yes
4	Poor/adverse response to drug	
	(a) Increased off period	Yes
	(b) Disabling dyskinesia	Yes
	(c) Disabling motor fluctuations	Yes
5	Degree of disability (UPDRS Part III score) > 25	Yes
6	Neuropsychology, MMSE > 24	Yes
7	Levodopa challenge response positive (30% improvement in UPDRS after 12-hour off medication)	Yes
8	Advanced comorbidity	No
9	Long-term anticoagulation	No
10	Willing for surgery and programming	Yes

PSP, progressive supranuclear palsy; UPDRS, unified Parkinson's disease rating scale; MSA, multiple system atrophy; CBD, corticobasal degeneration; LBD, Lewy body dementia

## Surgical Procedure

No antiparkinsonism medicine was given in the morning after the last previous night dose to see the clinical effects during awake surgery (Figs 1 to 14).

- Preoperative DBS protocol MRI was under general anesthesia (done day before if surgery planned awake, but done on the same day if surgery planned under general anesthesia)
- Fixation of stereotactic frame (we use Leksell frame) under scalp block if surgery planned awake.
- Planning in StealthStation (computer software)—
  - Anatomical STN targeting,
  - Trajectory planning,
  - Selection of entry point on the skull,
  - Getting the stereotactic frame settings.
- Burr hole was done at the entry point selected
- Microelectrode recording (MER) to locate the STN
- Microstimulation to see clinical effects and side effects
- Final electrode placement and confirmed by C-arm fluoroscopy
- Same procedure was repeated in opposite side.
- Pacemaker (battery) placement at subclavicular subcutaneous space.
- Impedance check and programming

Antiparkinsonian drugs to be started as early as possible through Ryle's tube. Pacemaker started "ON" after 48 hours in the low setting.

Patients were usually discharged after 7 or 8 days. The next programming was done after 2 weeks periodically according to clinical effects.

Programming parameters:

- Contact selection,
- Intensity of current (voltage),
- Pulse width (microsecond),
- Frequency (Hz),
- Mode of stimulation (monopolar, bipolar, and tripolar).

Postoperative CT scan of brain and preoperative MRI had been merged in StealthStation to see the best contacts and stimulated accordingly.

Voltage, pulse width, and frequency had been increased and adjusted according to clinical response.

## RESULTS

Hypokinesia, tremor, rigidity, and dyskinesia of 27 patients had improved significantly.

One patient expired due to neurolept malignant syndrome, followed by pneumonia and septicemia.

One patient developed infection of battery location, required wound debridement, but recovered.

Two patients had small hematoma along the lead track, resolved with conservative treatment. One of them developed hemiparesis.

Twenty-seven patients had bilateral monopolar cathodic stimulation. Two patients had unilateral bipolar stimulation, and one patient had bilateral bipolar stimulation.

Mean stimulation voltage was 2.8 (ranging from 1 to 3), pulse width was 60 microsecond (ranging from 60 to 90), and rate ranged from 130 to 180 Hz.

Levodopa was stopped in patients with severe dyskinesia with stimulation ON and few other antiparkinsonian medications ON (Tables 2 to 4).

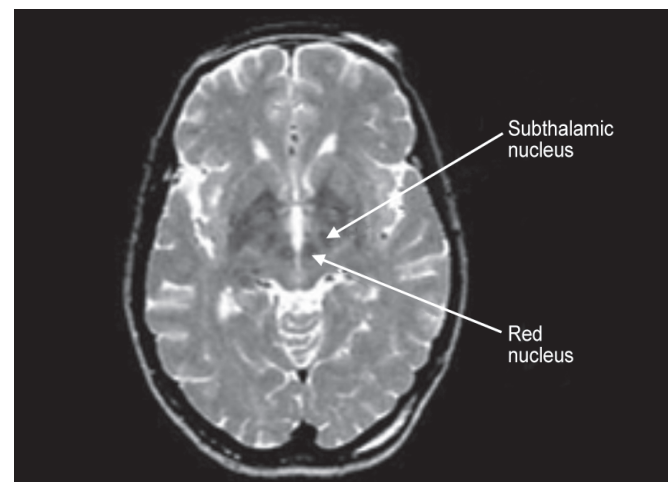


Fig. 2: Location of STN and red nucleus in T2-weighted MRI



Figs 1A and B: (A) Leksell stereotactic frame fixed with skull; (B) CT scan was being done with stereotactic frame



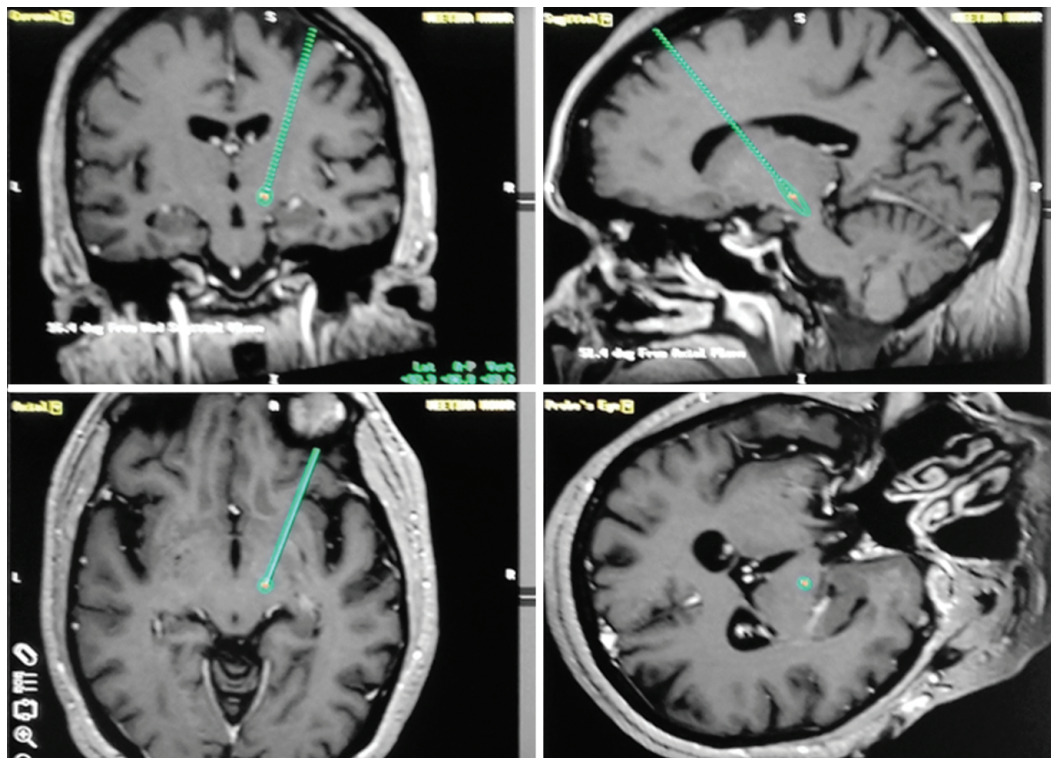


Fig. 3: STN targeting, trajectory and entry point selection at computer planning station from preoperative DBS protocol MRI—right side

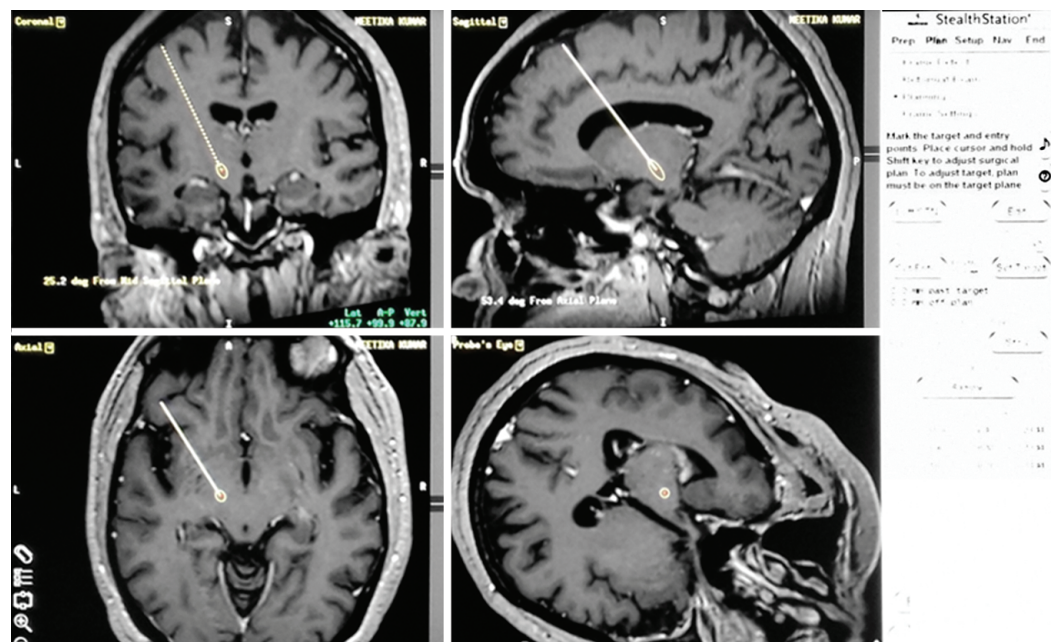


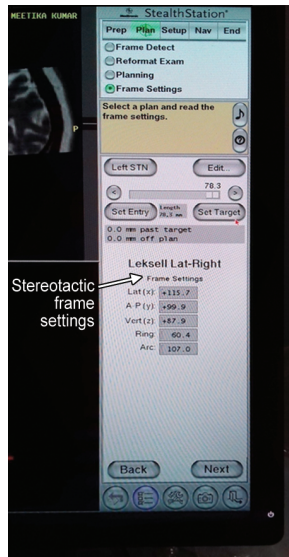
Fig. 4: STN targeting, trajectory and entry point selection at computer planning station from preoperative DBS protocol MRI—left side

## CONCLUSION

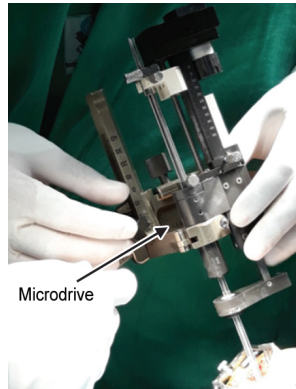
The results of STN-DBS in this small series, therefore, seem to be good, effective, and safe for the treatment of select medically refractory Parkinson's disease with a overall 5% risk of complications, which is comparable to the existing literature.

## ACKNOWLEDGMENTS

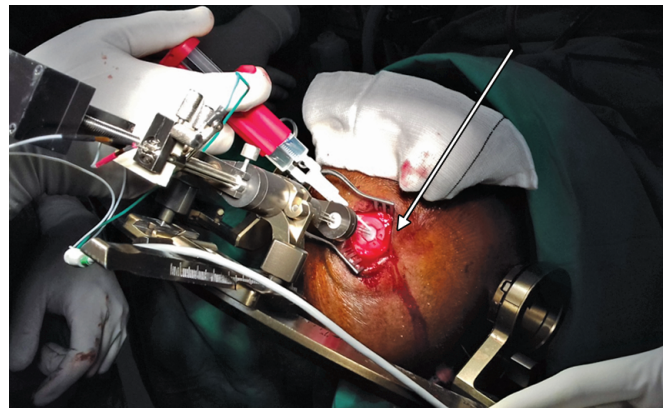
Dr Hrishikesh Kumar and Dr Purba Basu (Department of Neurology, Movement Disorder), Department of Psychiatry and Psychology, Department of Anaesthesiology and Critical Care, Department of Radiology, Institute of Neurosciences, Kolkata, West Bengal, India.



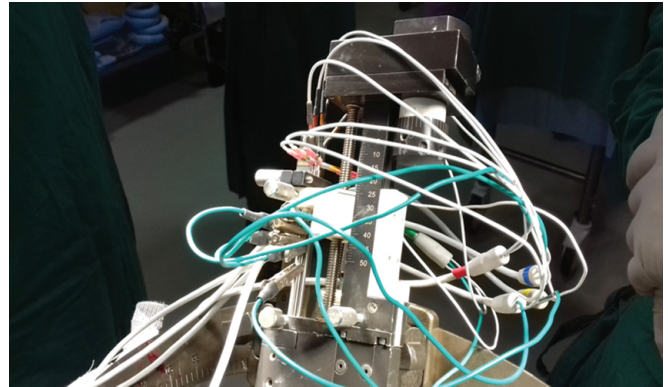
**Fig. 5:** After selecting target, trajectory and entry point in the computer station based on preoperative MRI, stereotactic coordinates have been generated



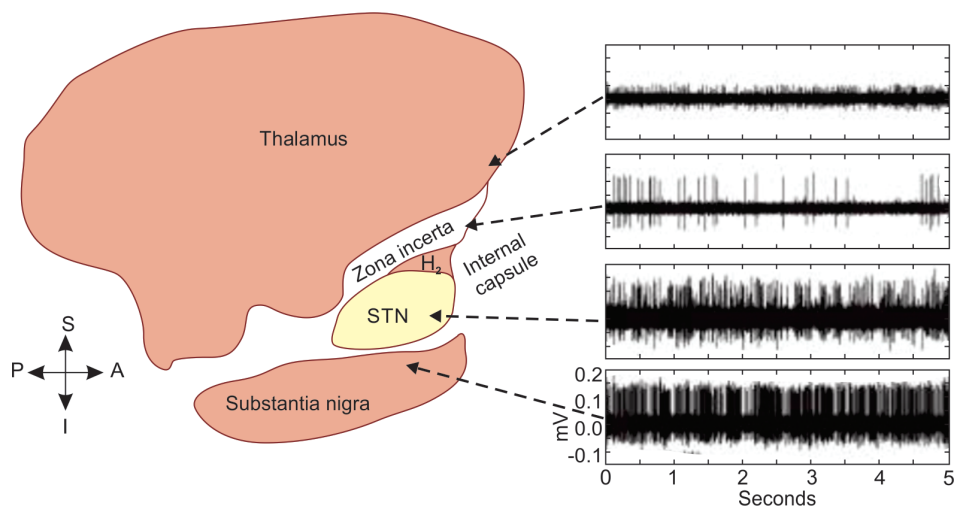
**Fig. 7:** Microelectrode drive was attached with stereotactic frame for microelectrode recording



**Fig. 6:** Burr hole was made at the selected entry point, dura was opened and fibrin glue was given to restrict the CSF egress as CSF egress can cause brain shift resulting malpositioning of lead



**Fig. 8:** Microelectrode recording (MER) and microstimulation is going on through microelectrode drive and wires



**Fig. 9:** As the microelectrode (red line) passes through thalamus, zona incerta subthalamic nucleus, and substantia nigra, different electrophysiological graphs will appear, guiding us to know location of the electrode





Fig. 10: Intraoperative fluoroscopy to see the lead position

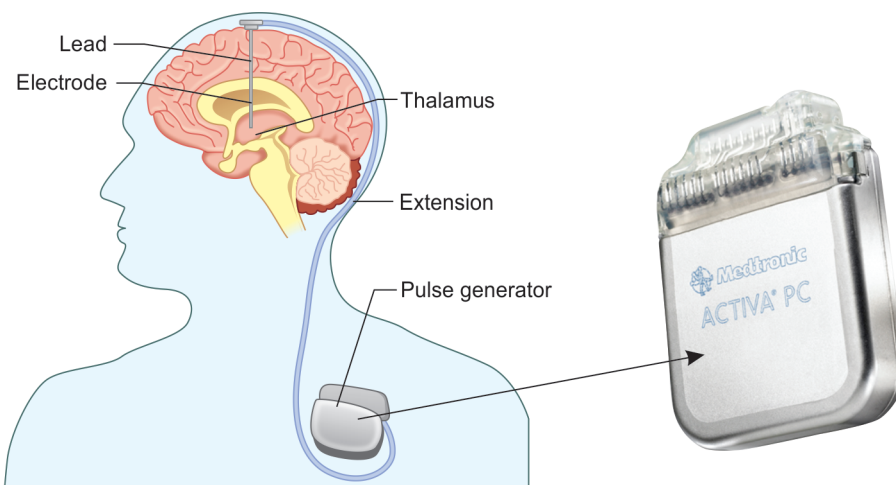


Fig. 11: Components of DBS system (lead, extension wire and pulse generator)

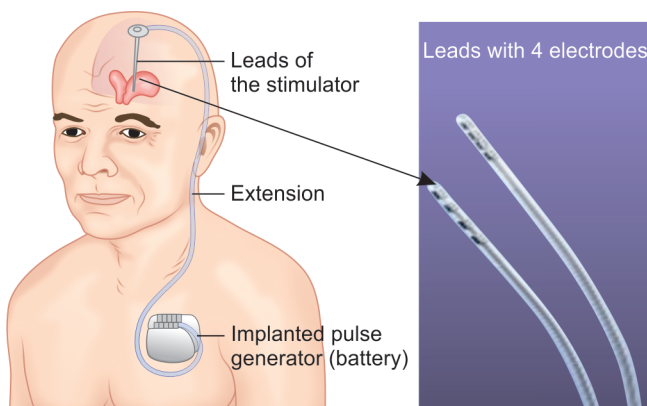
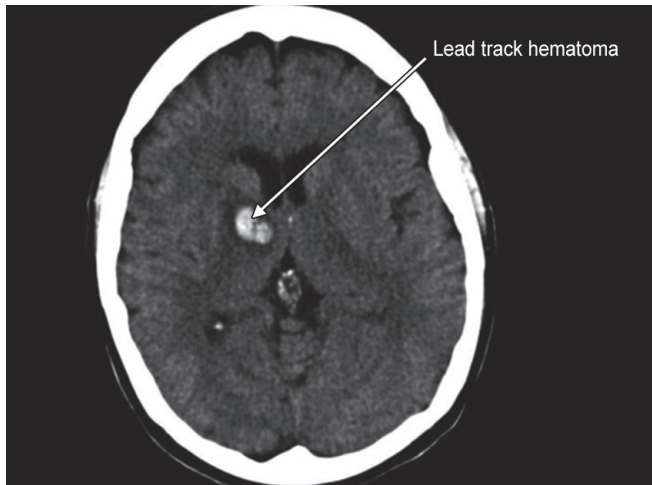


Fig. 12: Leads with four electrodes which remain within STN



Fig. 13: Programmer for postoperative programming of pacemaker (pulse generator)



**Fig. 14:** Postoperative CT scan showing small hemorrhage as complication

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**Table 2:** Comparison between the preoperative and postoperative conditions (mean values)

Test	Before surgery		Stimulation ON	
	Medication OFF	Medication ON	Medication OFF	Medication ON
UPDRS part III (overall average)	59	20	25	14
Rigidity (item 22)	4	2	2	1
Akinesia (item 31)	4	2	1	1
Tremor (items 20 and 21)	8	4	1	1
Postural stability (item 29)	3	2	1	1
Gait (item 30)	3	1	1	0.8
Speech (item 18)	2	1	1	0 → 1
Part IV dyskinesia (LID)		11 (levodopa ON)		2 (levodopa OFF)
Stand-walk-sit test				
Seconds	48	17	20	15
No. of steps	73	30	34	27
Clinical fluctuations (items 36, 37, 38, and 39)		4	0	0

**Table 3:** Clinical outcome after deep brain stimulation from different studies<sup>1</sup>

Studies	No. of patients	Follow-up (years)	Improvement in UPDRS III (%)	Decrease in OFF time (%)	Increase in ON time without dyskinesia
Krack et al. (1997)	15	1	71		
Kumar et al. (1998)	7	1	58	80	200
Limousin et al. (1998)	20	1	60	72.7	
DBSPGSG (2001)	96	0.5	51	61	270–229
Volkman et al. (2001)	16	1	67		
Pahwa et al. (2003)	19	2.3	28	61	
Krack et al. (2003)	49	5	66–54		
Rodriguez-Oroz et al. (2005)	49	3	50–39	56–43	260–265
Fraix et al. (2006)	95	1	57		192
Deuschl et al. (2006)	156	0.5	41	64	237
Weaver et al. (2009)	255	0.5	29	42	171
Hamani et al. (2005)	471	5	56–49		
Kleiner-Fisman et al. (2006)	921	>0.5	52	68.2	
Our study (Ghosh et al. (2020))	30	0.5	75		100



**Table 4:** Complication after DBS related to surgery and hardware according to various studies<sup>1</sup>

<i>Studies</i>	<i>No. of leads</i>	<i>Follow-up (months)</i>	<i>Hemorrhage (%)</i>	<i>Infection (%)</i>	<i>Hardware complication (%)</i>
Binder et al. (2003)	357	60	3.1		
Temel et al. (2004)	178	60		3.8	
Blomstedt and Hariz et al. (2005)	161	40		3	17.3
Deuschl et al. (2006)	156	6	1.9	3.8	1.3
Goodman et al. (2006)	181	4	2	4.7	11.5
Voges et al. (2006)	352	56	0.2	5.7	13.9
Seijo et al. (2007)	252	37	6.9		3.84
Kenney et al. (2007)	507	10	1.5	4.4	4
Tir et al. (2007)	206	1	5.8	6.8	3.9
Sillay et al. (2008)	759	6			4.5
Weaver et al. (2009)	242	6	0.8	9.9	6.6
Hamani et al. (2005)	471		2		9
Hamani and Lozano (2006)	922		2.8	6.1	11.4
Kleiner-Fishman et al. (2006)	921		3.9	3.6	4.5
Videnovic and Metman (2008)	2205		3.8	2.9	5
Our study (Ghosh et al. (2020))	60	6	3.3 (2 patients)	1 (battery location)	1 (kinking of wires)