

SYLLABUS

Marrakesh, Morocco, November 12-17, 2011

XXth WORLD CONGRESS OF NEUROLOGY



SOCIÉTÉ MAROCAINE
DE NEUROLOGIE

WCN Education Program

Thursday, 17 November, 2011

09:00-12:30

FUNCTIONAL NEUROSURGERY AND DBS

Chairperson: **Tipu Aziz, USA**

HOW CAN WE USE FUNCTIONAL NEUROSURGERY IN EPILEPSY

Christian Elger, Germany

CURRENT INDICATIONS AND RESULTS OF DEEP BRAIN STIMULATION FOR MOVEMENT DISORDERS

Badih Adada, Lebanon

FUNCTIONAL NEUROSURGERY, CURRENT AND FUTURE DIRECTIONS

Tipu Aziz, USA

10:30-11:00 *Coffee Break*



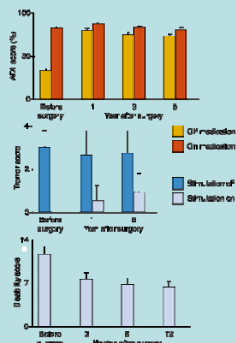
Functional Neurosurgery, Current and Future Directions

Professor Tipu Z Aziz, D.Med.Sci
Nuffield Department of Surgery
University of Oxford



Established DBS Indications

- Parkinson's Disease
- Tremor
- Dystonia



Parkinson's Disease

- Pre-op
- Unpredictable OFF periods
- Severely disabled in OFF state by tremor & akinesia



Parkinson's disease treated

- L Dopa
- On-Off Effects
- Dyskinesias
- Psychiatric effects



Parkinson's disease

- No understanding of the neural mechanisms
- 1979 MPTP case report
- 1983 primate MPTP model
- Electrophysiology
- 2-DG studies

Frozen Addicts

Fetal nigral brain
cell transplantation
in
Parkinsonism

STN lesion in MPTP primate

- Parkinsonian Primate
- L-DOPA responsive
- Unilateral STN lesion
- Total Reversal
- 2-DG studies reduced uptake in PPN

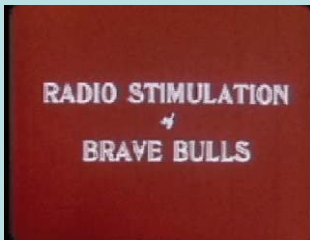


Surgical Technique

- Co-ordinates & entry point set



Deep Brain Stimulation



Deep Brain Stimulation

- Reversible
- Safer bilateral surgery
- Expensive
- Multiple surgeries over patient lifetime



Models of DBS Mechanisms

- Depolarisation block
- Activation of neuronal terminals that inhibit/excite efferent outputs (synaptic modulation)
- Depletion of efferent transmitters
- Network jamming
- Enhancement of transmitter release

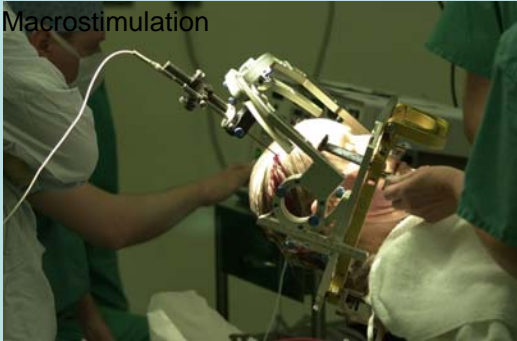
Surgical Technique

- Twist drill



Surgical Technique

- Macrostimulation



Surgical Technique



2-DG studies in MPTP primate

- Over active STN
- Excess inhibition of PPN
- Could STN surgery be therapeutic?
- DeLong-Ibotenic Acid lesions of STN
- Crossman- RF lesions of STN

STN Stimulation



PPN and Parkinson's Disease

- Degenerate in parkinsonian brain.
- Degenerate in akinetic non-parkinsonian disorders eg MSA and PSP.
- Also degenerate in DYT1 dystonia.

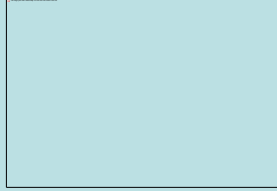
POST-MPTP

- Parkinsonian after MPTP

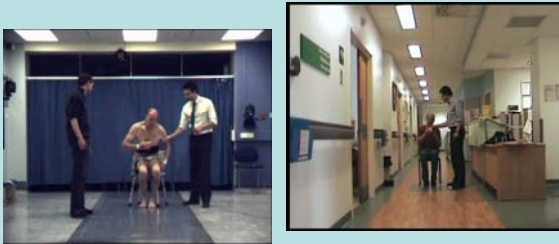


POST-MPTP + BICUCULLINE

- Note reversal of parkinsonian signs



PPN DBS Freezer-Fallers



Neuropathic Pain DBS

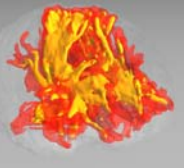
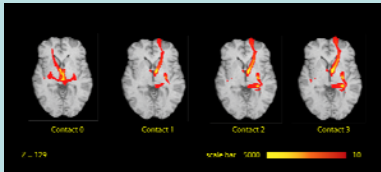
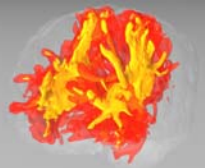


Cingulate DBS Depression



New Technology

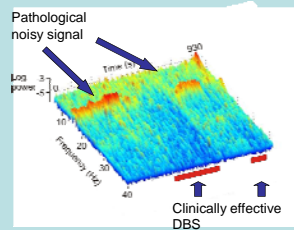
- Pre-planning with DTI
- Rechargeable pacemakers
- Closed loop stimulation
- Fully cranial implants



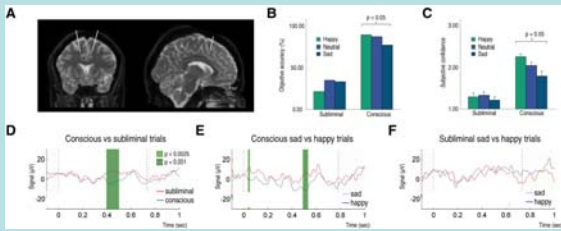
Owen et al (2008) JCON

Pathological Oscillations in PD

- Established the nature of the pathological 'noisy' signal in the brain in PD
- Demonstration that DBS works by suppressing local 'noise' in the brain
- Proof that one of the key functions of the human basal ganglia is the optimisation of motor programs given feedback on recent performance

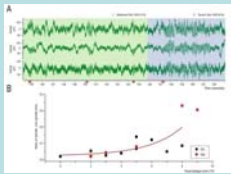
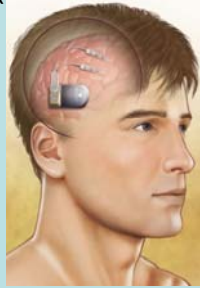


ACC DBS for Pain



Cranial Device (Neuropace)

- Closed loop
- Cranial implant
- Epilepsy
- Under trials



Green et al (2009) *Neurology*

Brain Machine Interface

- Rhesus Monkey
- Implanted electrodes



Courtesy of DEKA Research & Development and The Rehabilitation Institute of Chicago

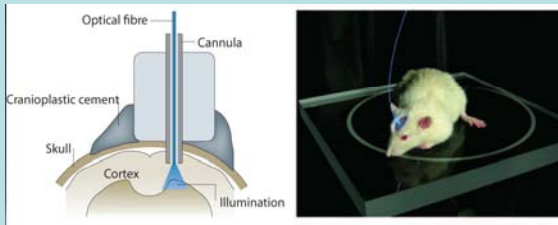
Optogenetics

Optical Deconstruction of Parkinsonian Neural Circuitry

Viviana Gradinaru,^{1,2*} Murtaza Mogri,^{1*} Kimberly R. Thompson,¹
Jaimie M. Henderson,³ Karl Deisseroth^{1,4†}

Deep brain stimulation (DBS) is a therapeutic option for intractable neurological and psychiatric disorders, including Parkinson's disease and major depression. Because of the heterogeneity of brain tissues where electrodes are placed, it has been challenging to elucidate the relevant target cell types or underlying mechanisms of DBS. We used optogenetics and solid-state optics to systematically drive or inhibit an array of distinct circuit elements in freely moving parkinsonian rodents and found that therapeutic effects within the subthalamic nucleus can be accounted for by direct selective stimulation of afferent axons projecting to this region. In addition to providing insight into DBS mechanisms, these results demonstrate an optical approach for dissection of disease circuitry and define the technological toolbox needed for systematic deconstruction of disease circuits by selectively controlling individual components.

Optogenetics



Optogenetics

- Hegemann, Bamberg, Nagel 2005:
- Channelrhodopsin-2 excites to Blue light
- Same folks: Pigment inhibits to Yellow light
- Deisseroth 2007: Lateral Hypothalamus
- CHR-2-wakes sleeping mice by release of Orexin

Optogenetics

Fiberoptic Control of Locomotion in ChR2 Mouse

Acknowledgements



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Functional
Neurosurgery

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Department of
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Genetics
