NonHuman Primate Research: Patient Advocacy Perspective

Joel S Perlmutter **Scientific Director Dystonia Medical Research Foundation** Elliot Stein Family Professor of Neurology, Radiology, Neuroscience, Physical Therapy, **Occupational Therapy** Washington University in St. Louis

Disclosures: Joel S Perlmutter

Research Support / Grants: NIH, Washington University,

American Parkinson Disease Association (APDA), Greater St. Louis Chapter of the APDA, McDonnell Center for Higher Brain Function, Barnes-Jewish Hospital Foundation, Huntington's Disease Society of America, CHDI, MJ Fox Foundation, Fixel Foundation, U Toronto, U Michigan, Riney Fund, N Grant Williams Fund, Jo Oertli Fund.

Stock/Equity/Consulting: None

Honoraria: U Rochester, American Academy of Neurology, American Neurologic Association, CHDI; Beth Israel - Harvard U, Stanford U, U Pennsylvania, U Illinois at Chicago (can of biscotti), Boston U

Speakers Bureau: None

US Patent/Other: None

NHP & Neurologic Disease: Critical Model Systems

Animal models with behaviors similar to dystonia in humans



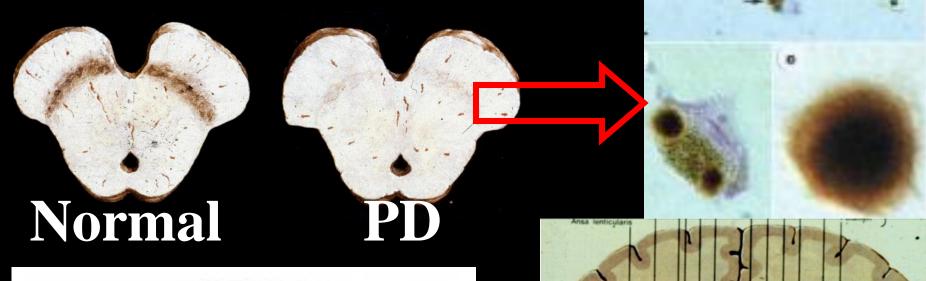


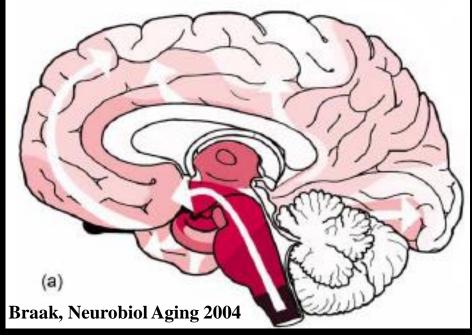
Current Problem

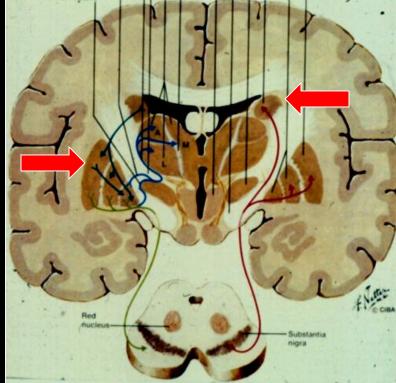
Cell based, lower animals, rodents and computer models do not have dystonia

Good for biochemical studies

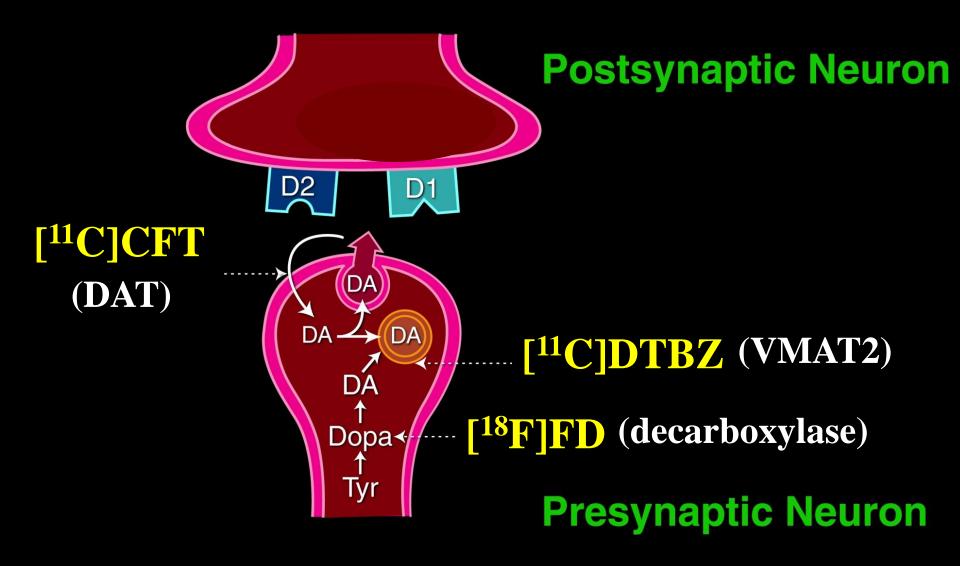
Parkinson Disease Lewy body







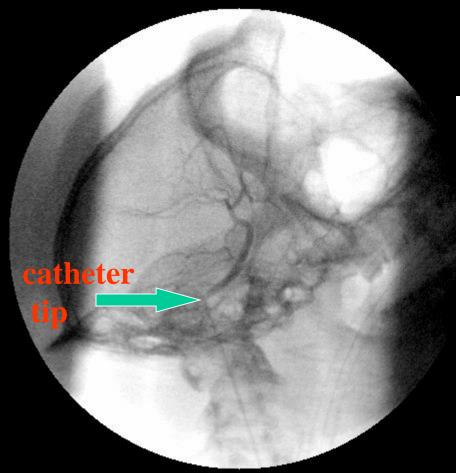
The Dopaminergic Synapse

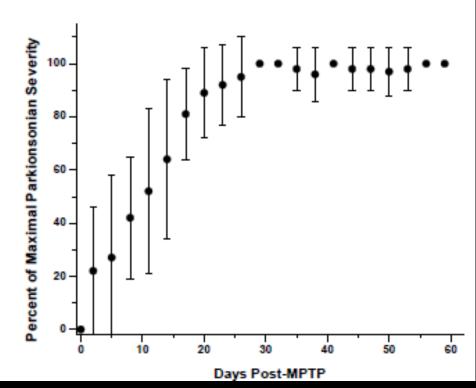


ELLDOPA: PSG, NEJM 2004 12 **Baseline DAT predicts** 10 Severity at 2 yrs (PPMI) Change in Total UPDRS from baseline Ravina et al Mov Dis 2012 8 **300** Placebo 6 150 P = .0364 2 150mg 0 -2 P < .001-4 6000 -6 -8 6 10 14 18 22 26 30 34 38 42 46 Week Medications Baseline withdrawn -Placebo --- 150mg --- 300mg --- 600mg

Unilateral ic MPTP

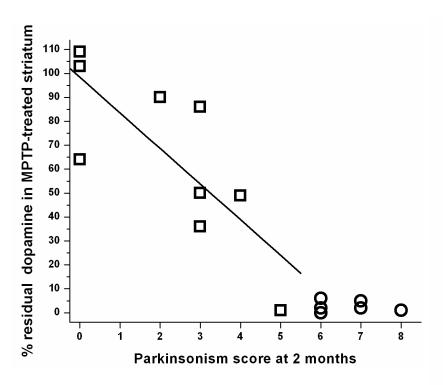


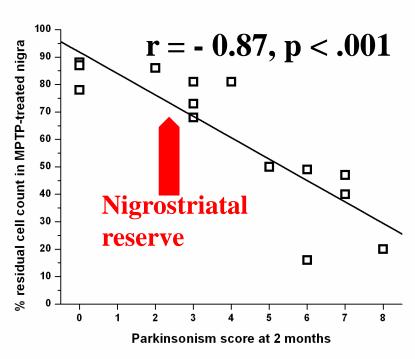




Todd R et al, 1996; Perlmutter JS et al, 1997 Tabbal S et al, 2006; Tabbal S et al, 2012

Parkinsonism vs striatal DA, SNpc





Ratings vs Dopamine

Ratings vs SNpc Cells





Tabbal S et al, Exp Neurol 2012

Current Needs

NHP model with obvious dystonia

For:

biomarker development/validation

DREADDS (Designer Receptor Exclusively

Activated by Designer Drugs)

therapeutic target identification metrics of therapeutic efficacy

NHP Critical

Pathophysiology vs Behavior