

Berenson-Allen Center for Noninvasive Brain Stimulation
Beth Israel Deaconess Medical Center
Harvard Medical School

Behavioral Intervention Research Using tDCS

Dylan J. Edwards PhD

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Professor of Neuroscience, ECU Australia

Adaptive

Plasticity

Maladaptive

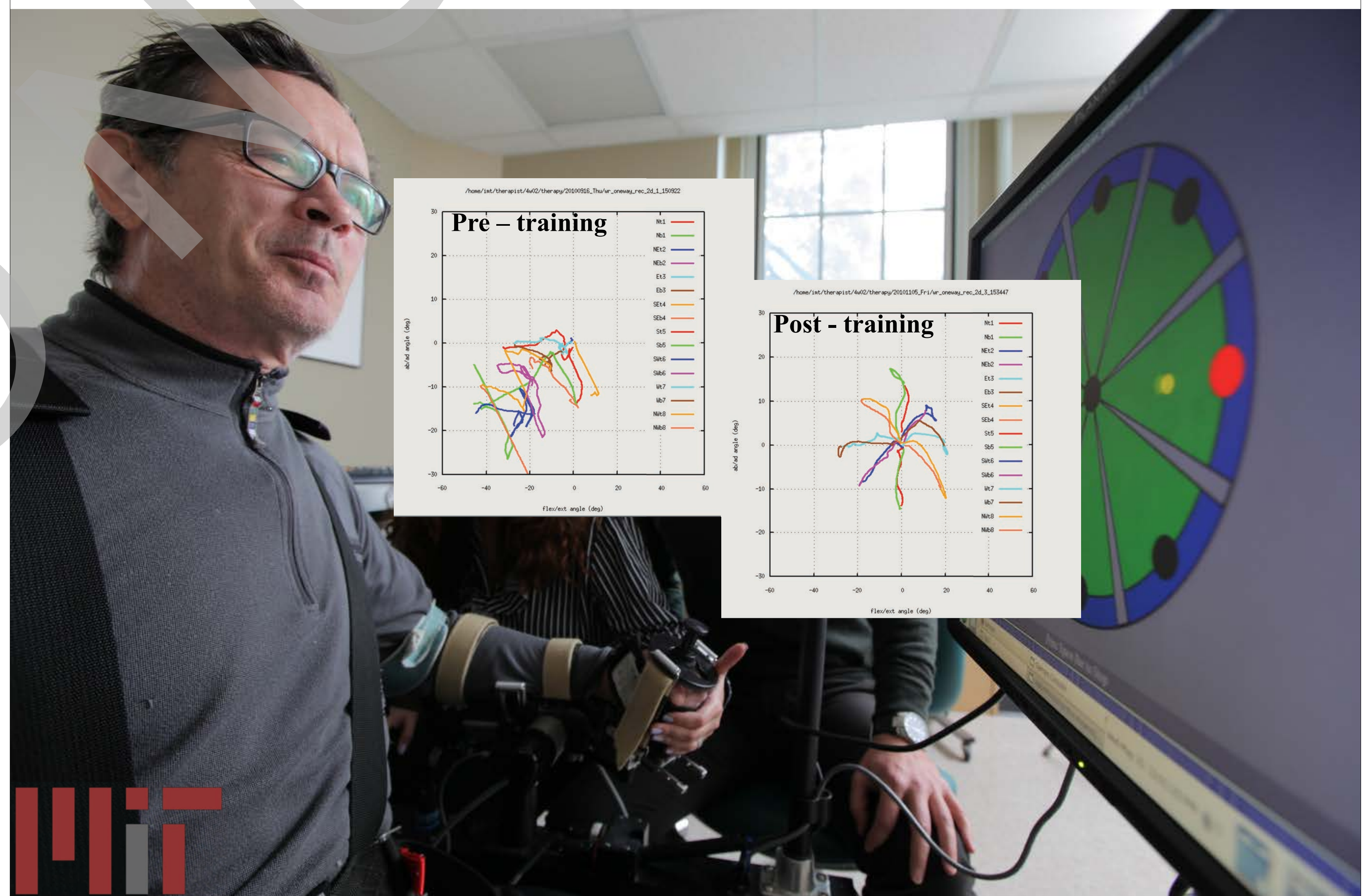
Promoting useful plasticity in motor cortex

Patient

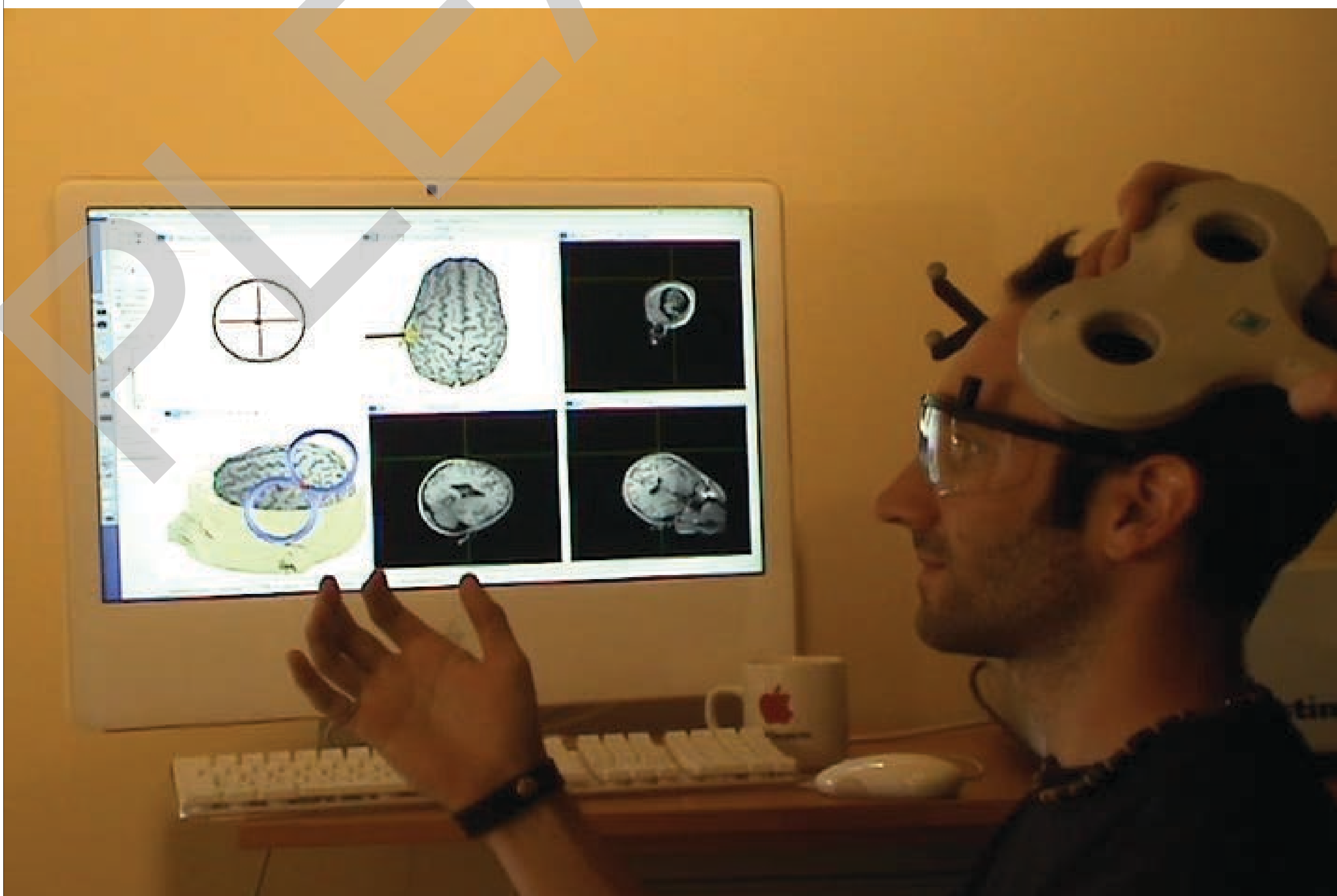
Intervention

Patient + Intervention

Robotics for assessment of performance kinematics



TMS Demonstration

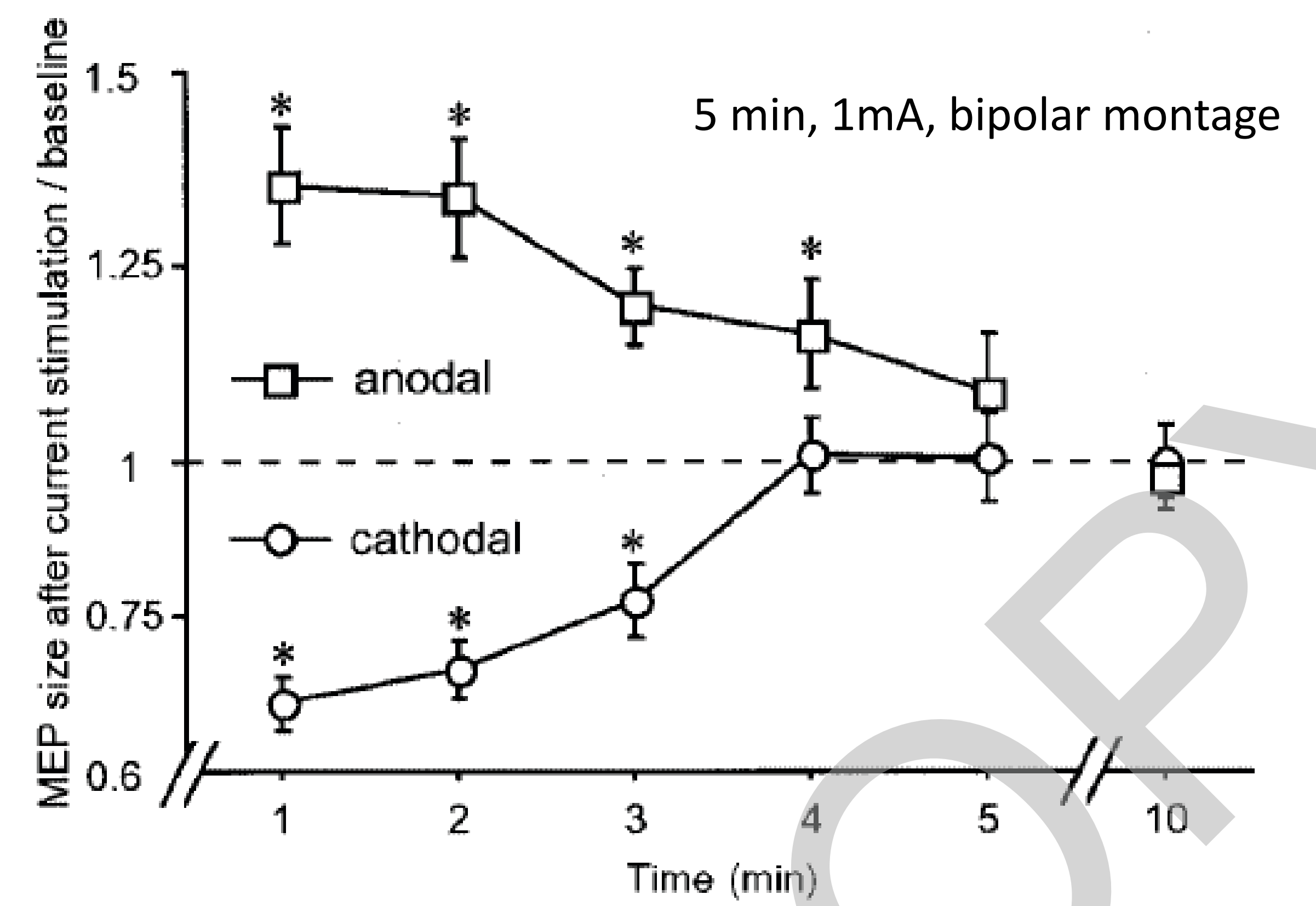
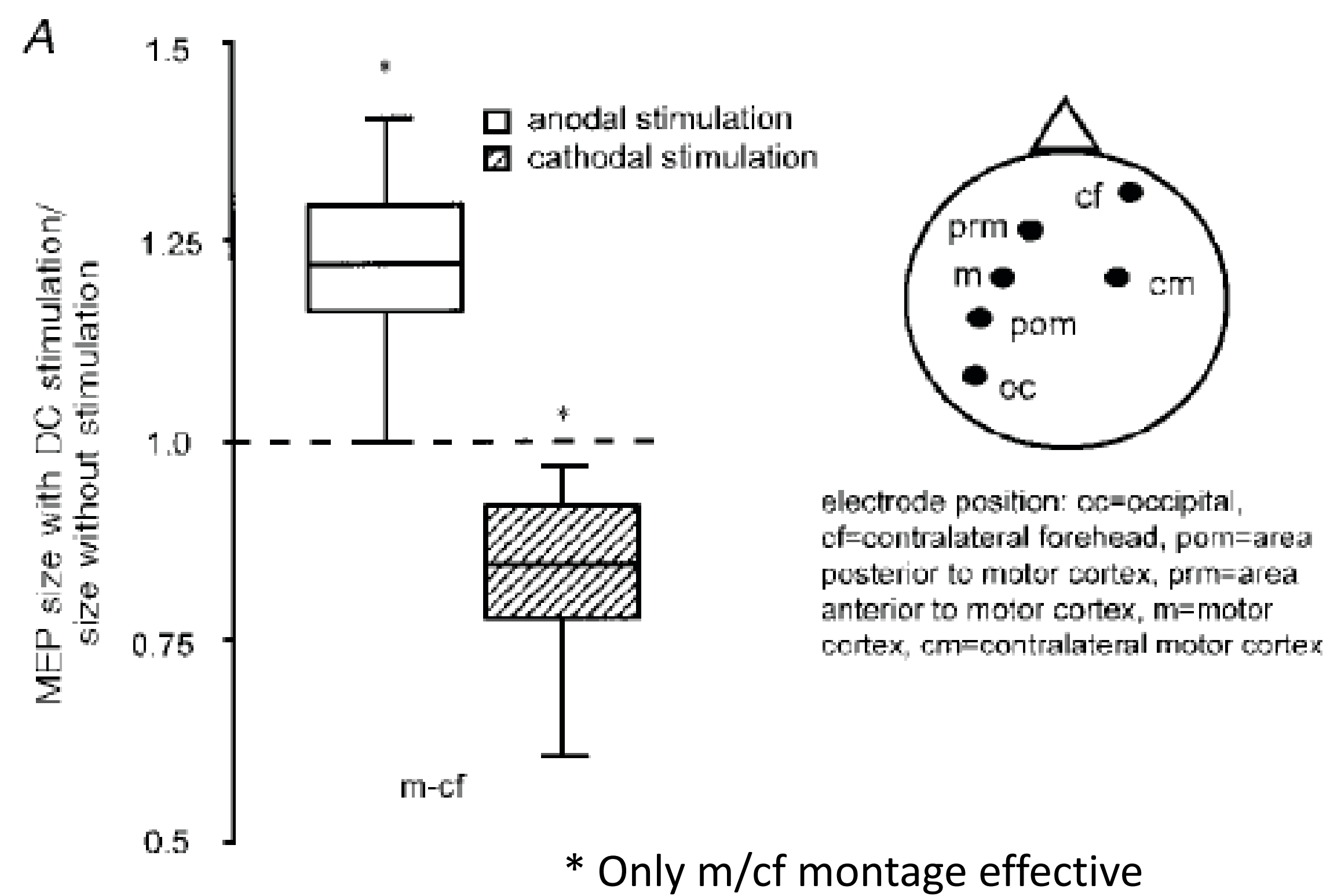


tDCS



Excitability changes induced in the human motor cortex by weak transcranial direct current stimulation

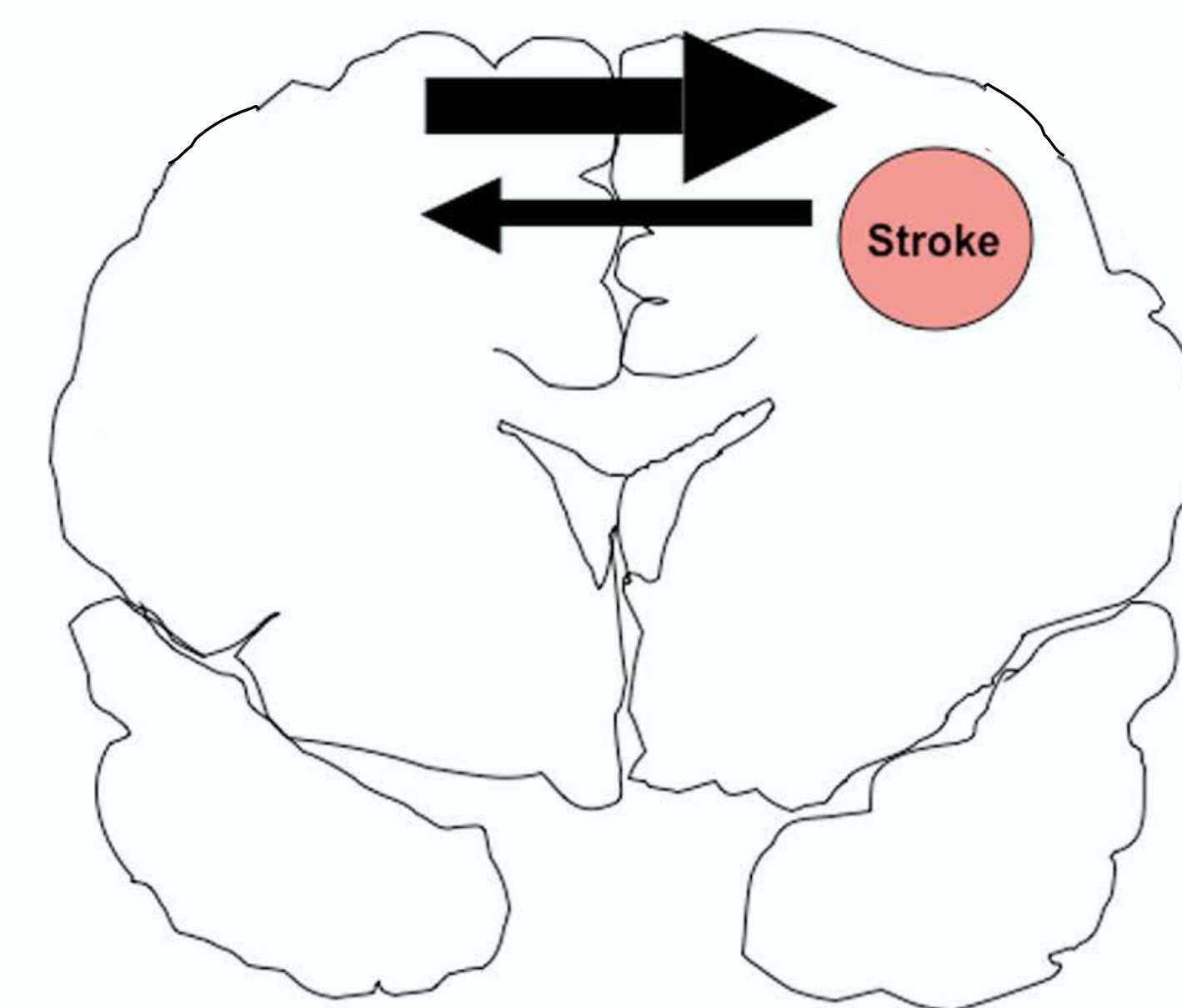
M. A. Nitsche and W. Paulus



"In order to induce aftereffects, a stimulus duration of at least 3 min at 1 mA or an intensity of 0.6 mA for 5 min"

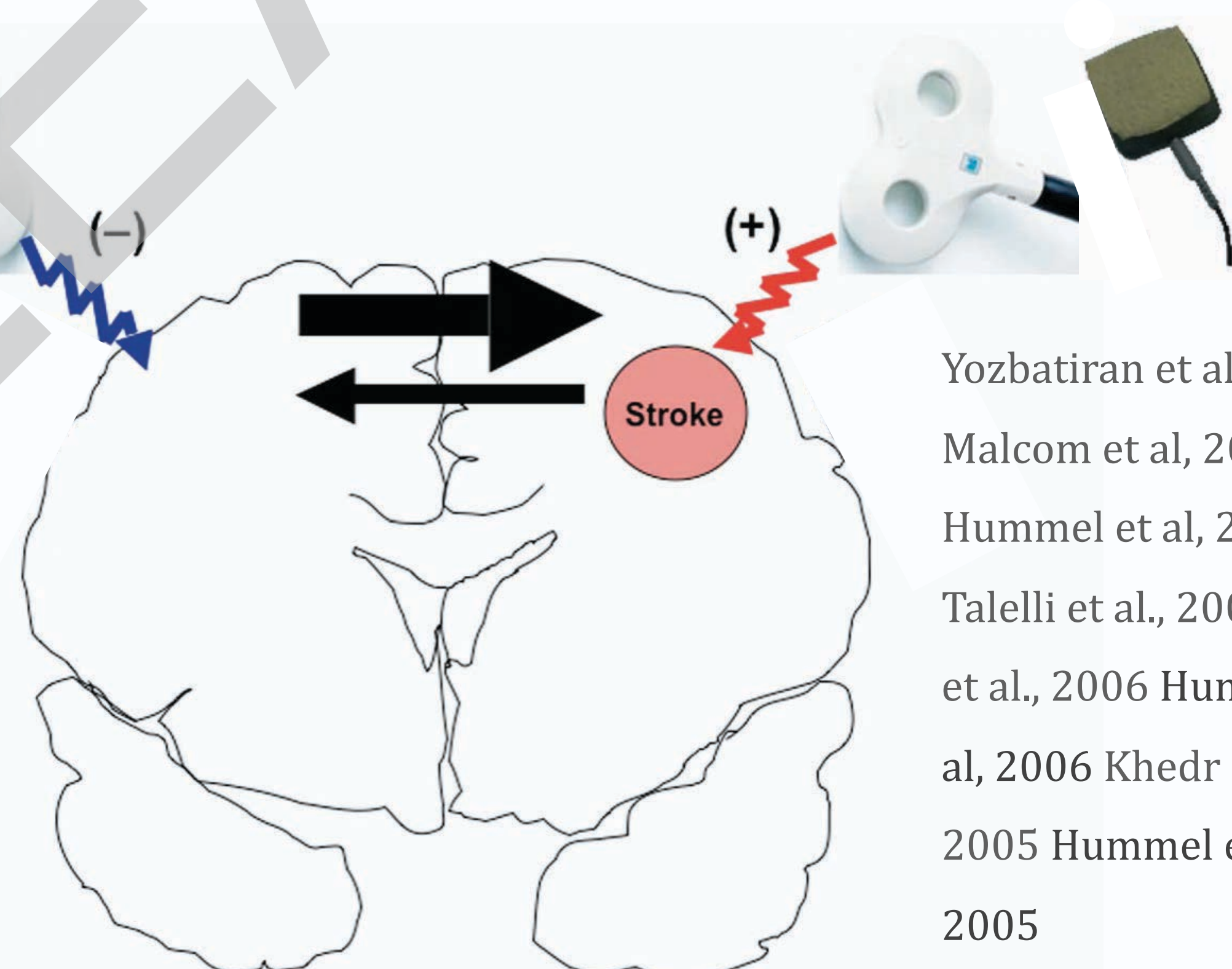
Nitsche et al, 2000

Corticomotor excitability in stroke



Webster et al (2006)

- Takeuchi, et al, 2008
- Boggio et al, 2007
- Fregni et al, 2006
- Fregni et al, 2005
- Mansur et al, 2005
- Takeuchi et al, 2005
- Boggio, et al., 2006
- Werhahn, et al., 2003



- Yozbatiran et al, 2009
- Malcom et al, 2007
- Hummel et al, 2007
- Talelli et al, 2007
- Kim et al, 2006
- Hummel et al, 2006
- Khedr et al, 2005
- Hummel et al, 2005

Webster et al (2006)

IMPROVED CORTICOMOTOR OUTPUT FROM IPSI-LESIONAL M1 & IMPROVED MOTOR BEHAVIOUR

Functional Improvements

- sRT/cRT
- Pinch force acceleration
- fingers/thumb AROM
- Movement accuracy
- Purdue Pegboard
- JTT

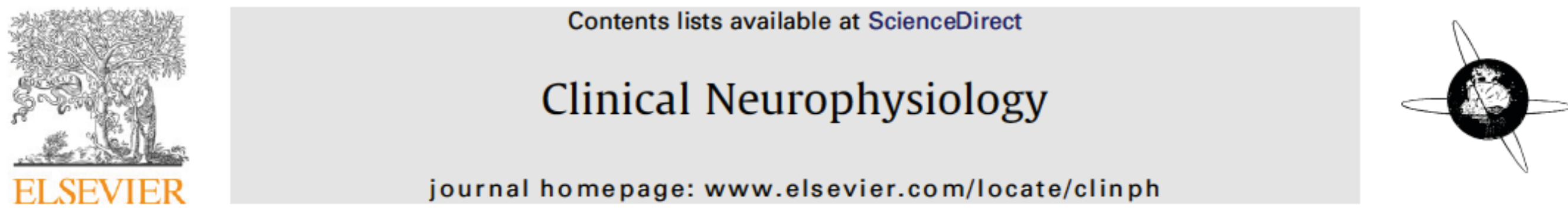
TMS correlates

- Resting MT
- Transcallosal Inhibition
- MEP Amplitude

Webster et al (2006)

IMPROVED CORTICOMOTOR OUTPUT FROM IPSI-LESIONAL M1 & IMPROVED MOTOR BEHAVIOUR

Anodal tDCS favors clinical improvement in stroke...



Invited Review

Does anodal transcranial direct current stimulation enhance excitability of the motor cortex and motor function in healthy individuals and subjects with stroke: A systematic review and meta-analysis

A. Bastani, S. Jaberzadeh*

Department of Physiotherapy, School of Primary Health Care, Faculty of Medicine, Nursing and Health Sciences, Monash University, Melbourne, Australia

SPECIAL ISSUE

A Meta-analysis of the Efficacy of Anodal Transcranial Direct Current Stimulation for Upper Limb Motor Recovery in Stroke Survivors

Butler et al.

Survivors

Level of Evidence: Level 1a.

J HAND THER. 2012; ■■■-■■■

Restorative Neurology and Neuroscience 25 (2007) 123–129

Repeated sessions of noninvasive brain DC stimulation is associated with motor function improvement in stroke patients

Paulo S. Boggio^{a,b,1}, Alice Nunes^{a,1}, Sergio P. Rigonatti^a, Michael A. Nitsche^c, Alvaro Pascual-Leone^{d,e} and Felipe Fregni^{d,*}

^aDepartment of Experimental Psychology and Department of Psychiatry, University of Sao Paulo, Sao Paulo, Brazil (address where the work was carried out)

^bJcleo Neurociências Mackenzie University, Sao Paulo, Brazil

^cDepartment of Clinical Neurophysiology, Georg-August-University, Goettingen, Germany

^dCenter for Noninvasive Brain Stimulation, Beth Israel Deaconess Medical Center, Harvard Medical School, Boston, USA

^eInstitute Gutmann for Neurorehabilitation, Barcelona, Spain

How does repetitive behavior affect motor cortex?

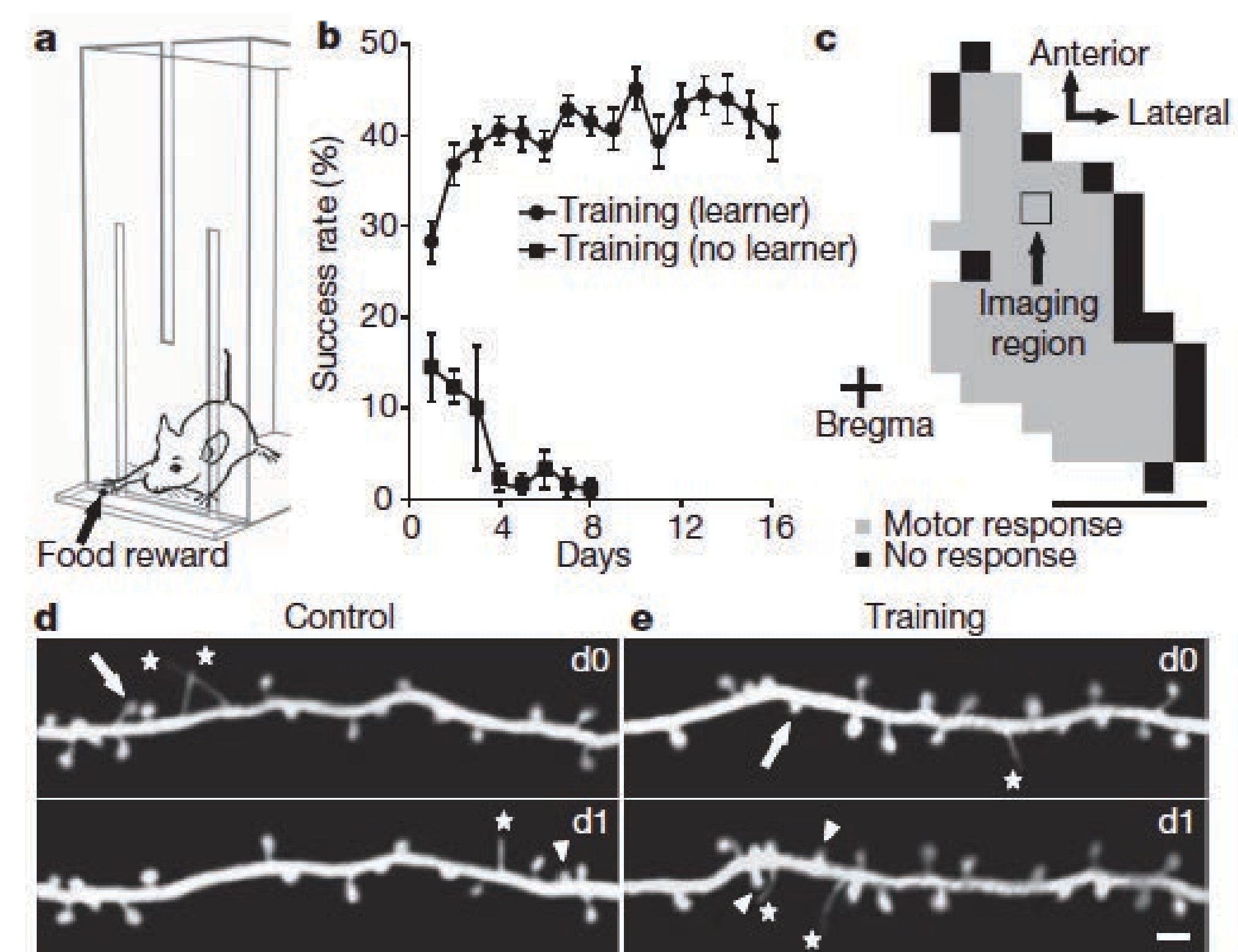
doi:10.1038/nature08389

nature

LETTERS

Rapid formation and selective stabilization of synapses for enduring motor memories

Tonghui Xu^{1*}, Xinzhu Yu^{1*}, Andrew J. Perlik¹, Willie F. Tobin¹, Jonathan A. Zweig¹, Kelly Tennant², Theresa Jones² & Yi Zuo¹



Neurobiology of Learning and Memory 74, 27–55 (2000)

Effects of Repetitive Motor Training on Movement Representations in Adult Squirrel Monkeys: Role of Use versus Learning

Erik J. Plautz*, Garrett W. Milliken,† and Randolph J. Nudo‡,§

*Department of Neurobiology and Anatomy, University of Texas–Houston, Houston, Texas 77030;

†Department of Psychology, College of Charleston, Charleston, South Carolina 29424; and

‡Department of Molecular and Integrative Physiology and §Center on Aging, University of Kansas Medical Center, Kansas City, Kansas 66160

Motor map does not change unless in skill context

Exp Brain Res (2006) 174: 199–209
DOI 10.1007/s00221-006-0440-8

RESEARCH ARTICLE

S. Koenke · K. Lutz · U. Herwig · U. Ziemann
L. Jäncke

Extensive training of elementary finger tapping movements changes the pattern of motor cortex excitability

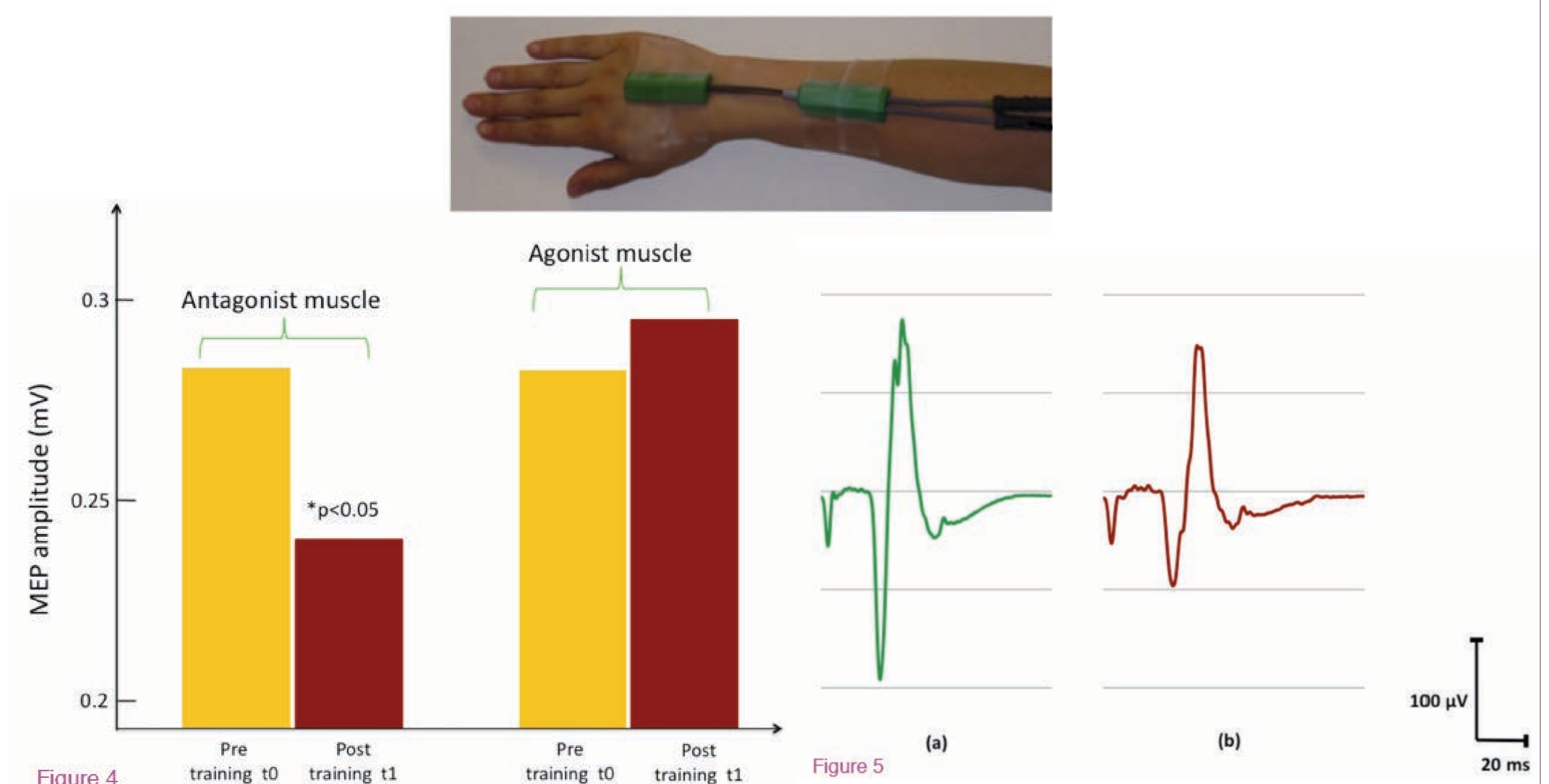
Simple repetitive finger movements increase excitability

Journal of NeuroEngineering and Rehabilitation

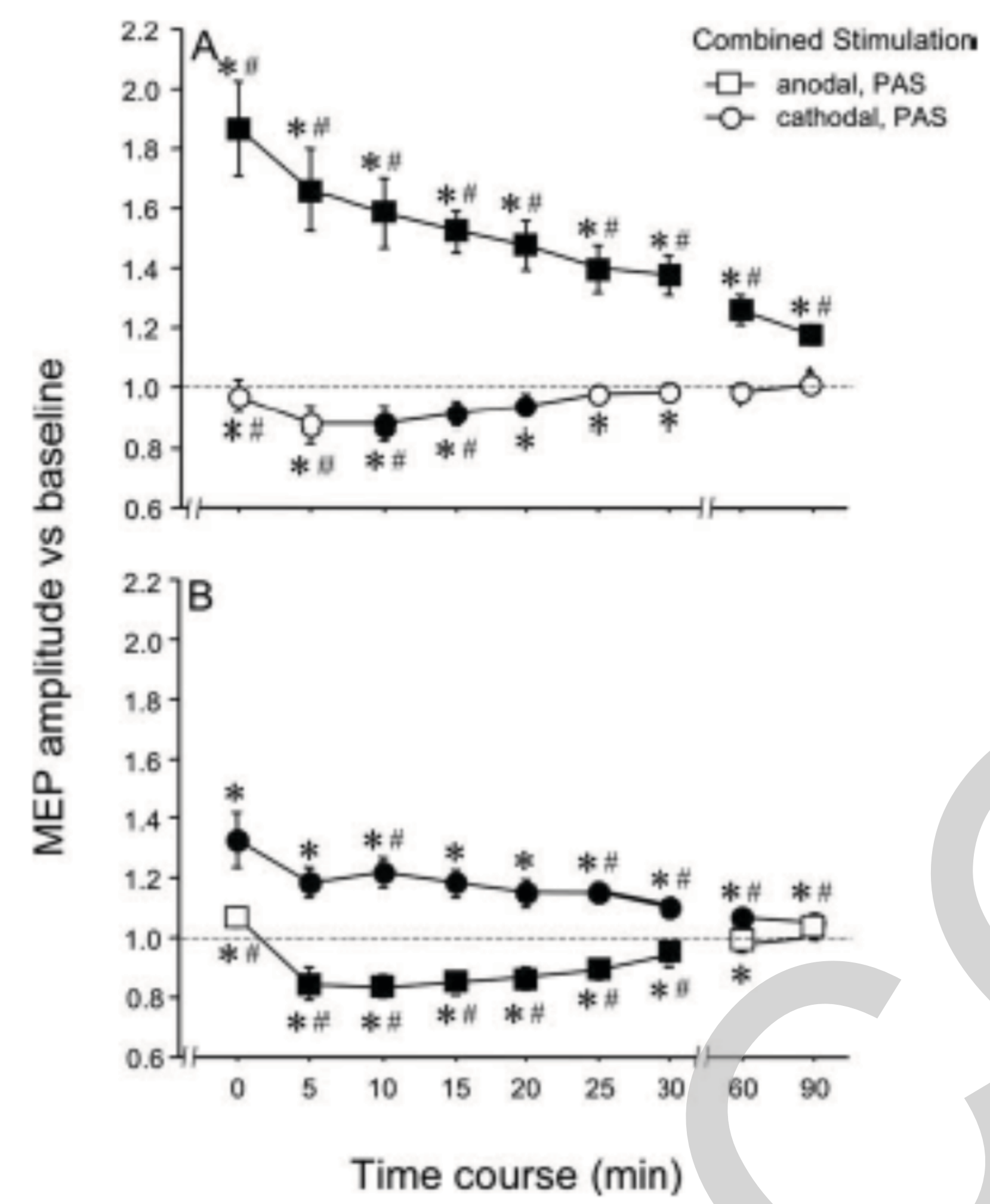
BioMed Central

Reversal of TMS-induced motor twitch by training is associated with a reduction in excitability of the antagonist muscle.

Viola Giacobbe^{1,2}, Bruce T. Volpe¹, Gary W. Thickbroom², Felipe Fregni^{3,6}, Alvaro Pascual-Leone^{3,5}, Hermano I. Krebs⁴, Dylan J. Edwards^{1,2,3}



How does combined intervention affect motor cortex?



Anodal tDCS prior to excitatory PAS further boosts excitability, while during tDCS reverses effect to reduced excitability

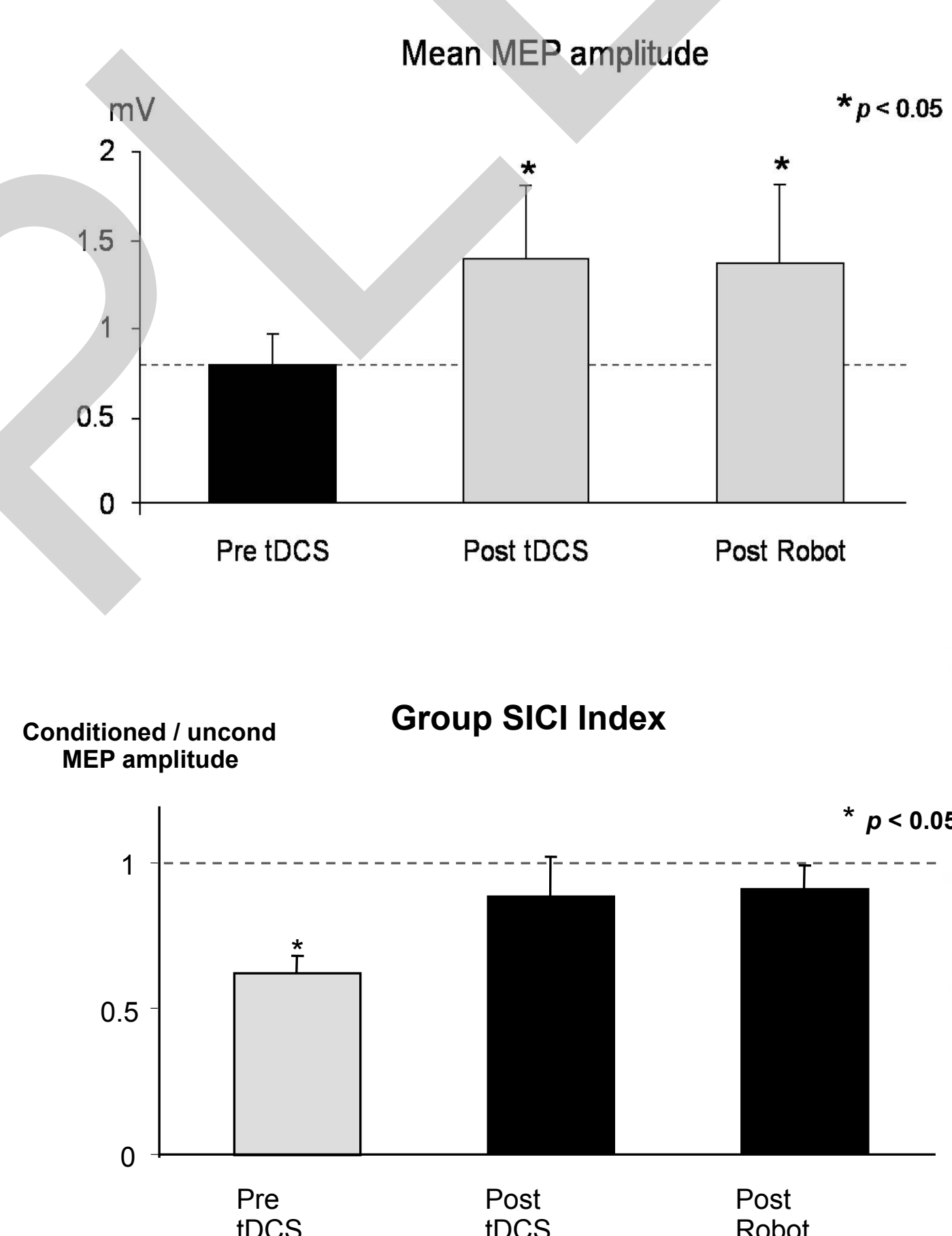
Nitsche et al (2007)

Is coupling tDCS with training good?

Anodal tDCS combined with robotic motor training



Anodal tDCS combined with robotic motor training



Edwards et al (2009)

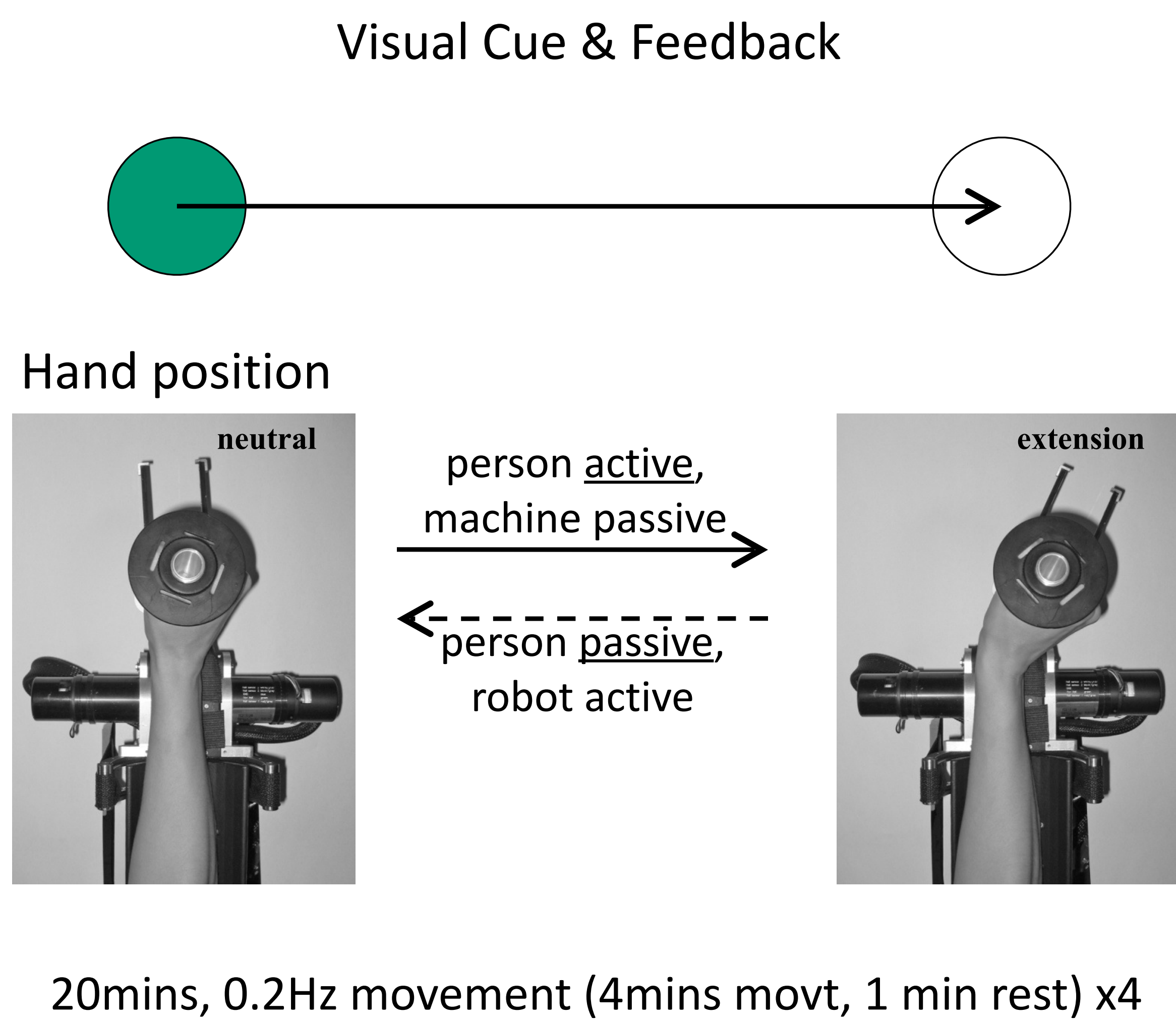
Relevance of kinematic measures to clinical function

- Movement Speed (peak, mean)
- Movement Smoothness
- Aim
- Deviation

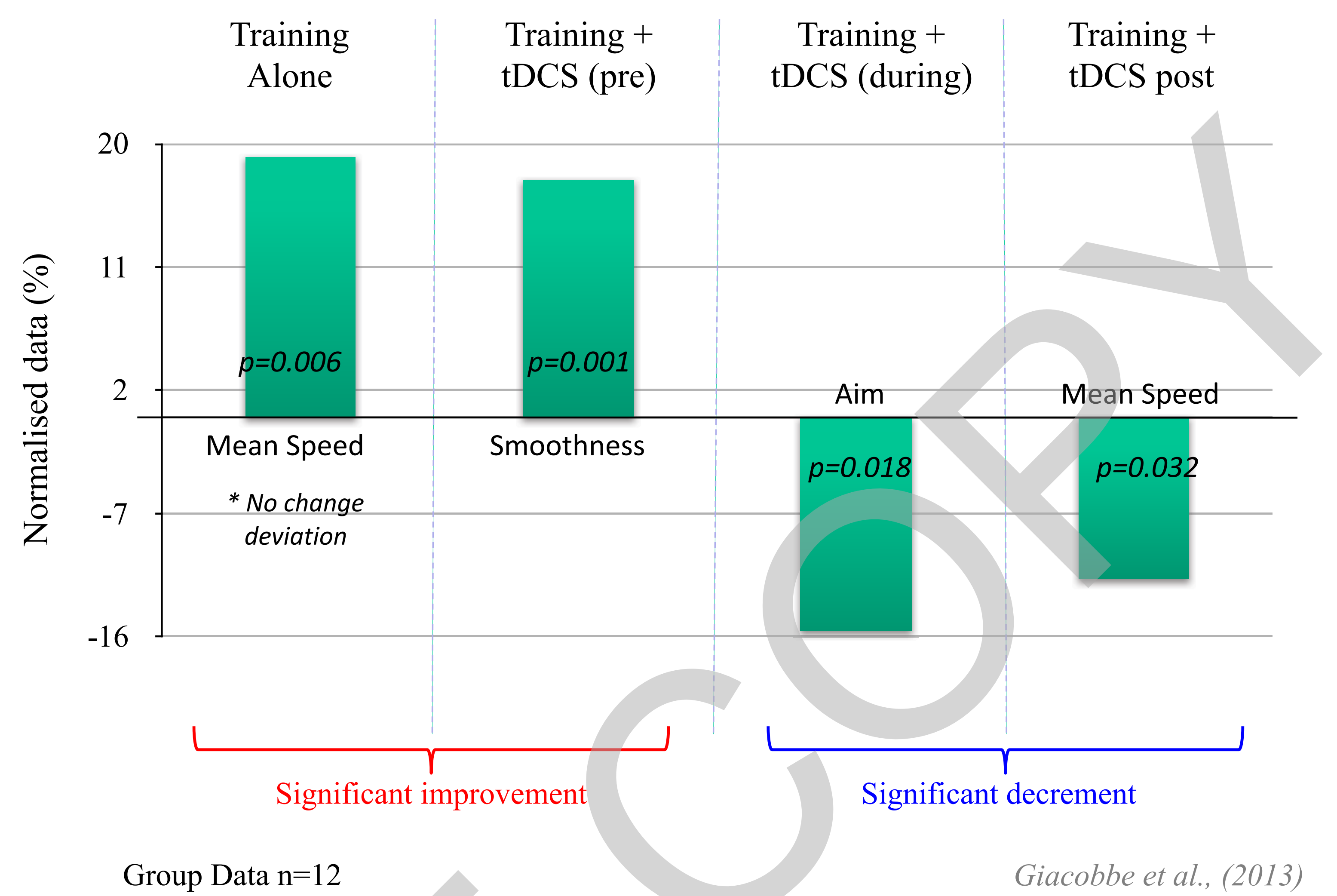
Highest correlation with clinical function

"Kinematic Robot-Based Evaluation Scales and Clinical Counterparts to Measure Upper Limb Motor Performance in Patients With Chronic Stroke" (Bosecker et al, 2009)

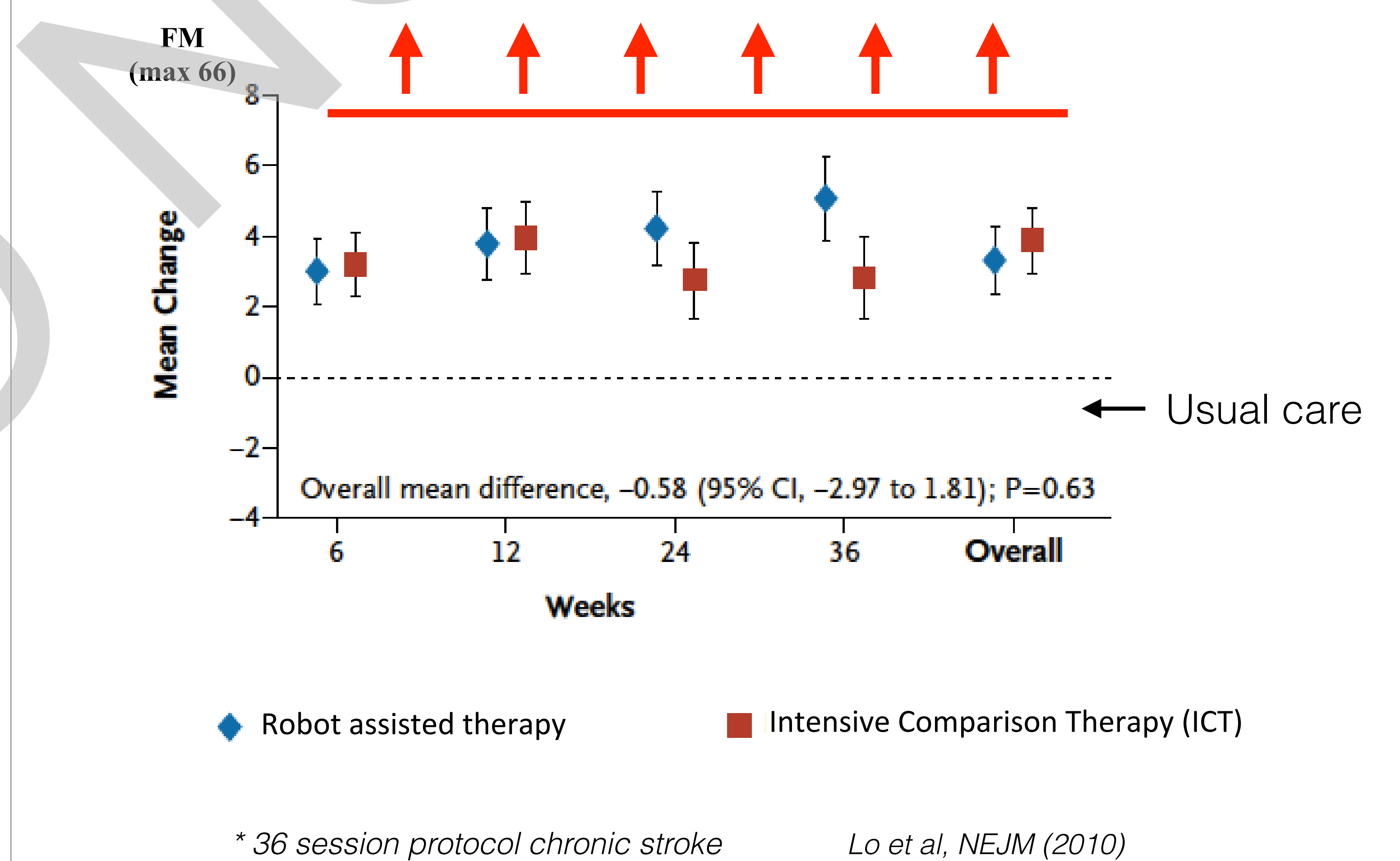
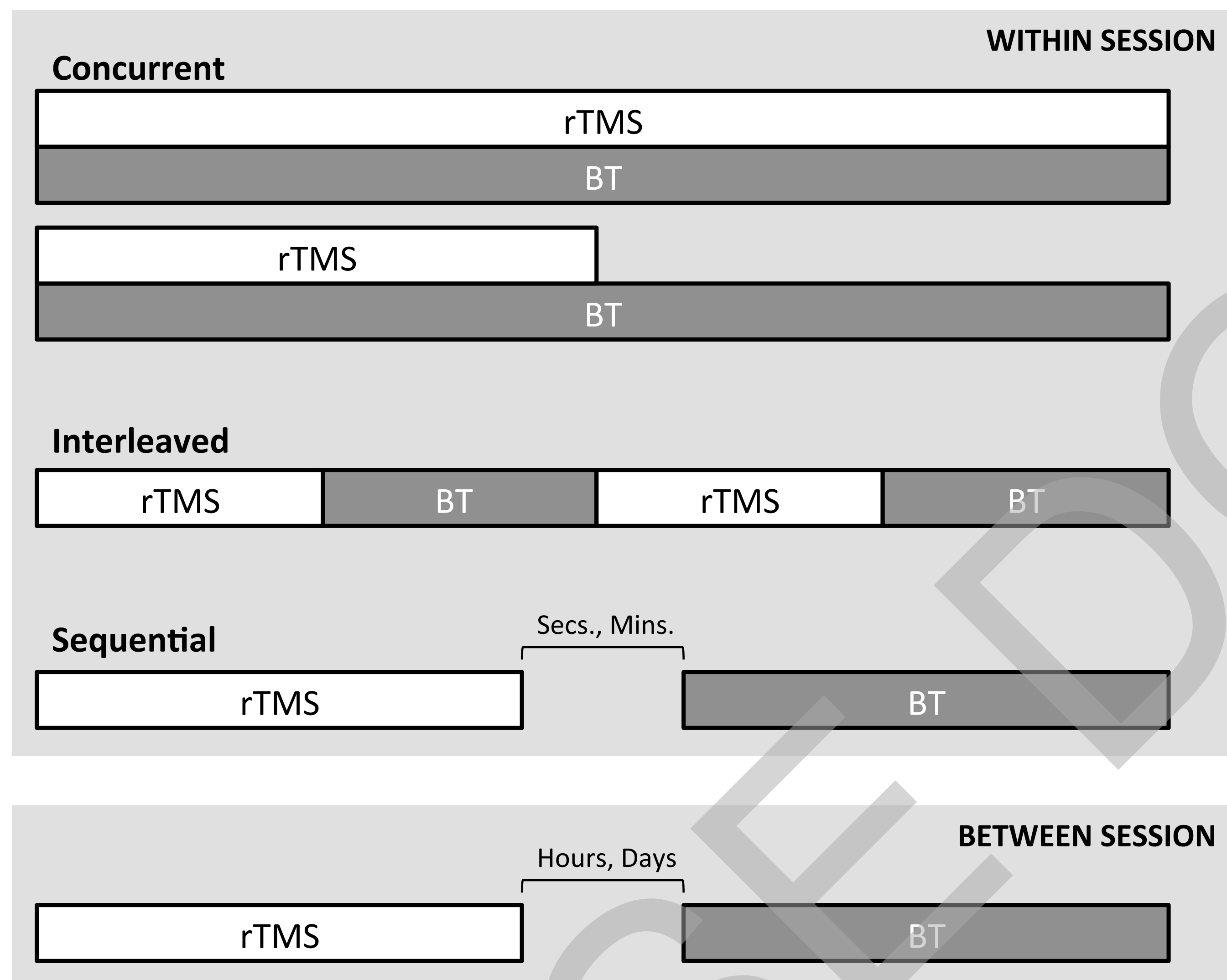
Movement Training Paradigm



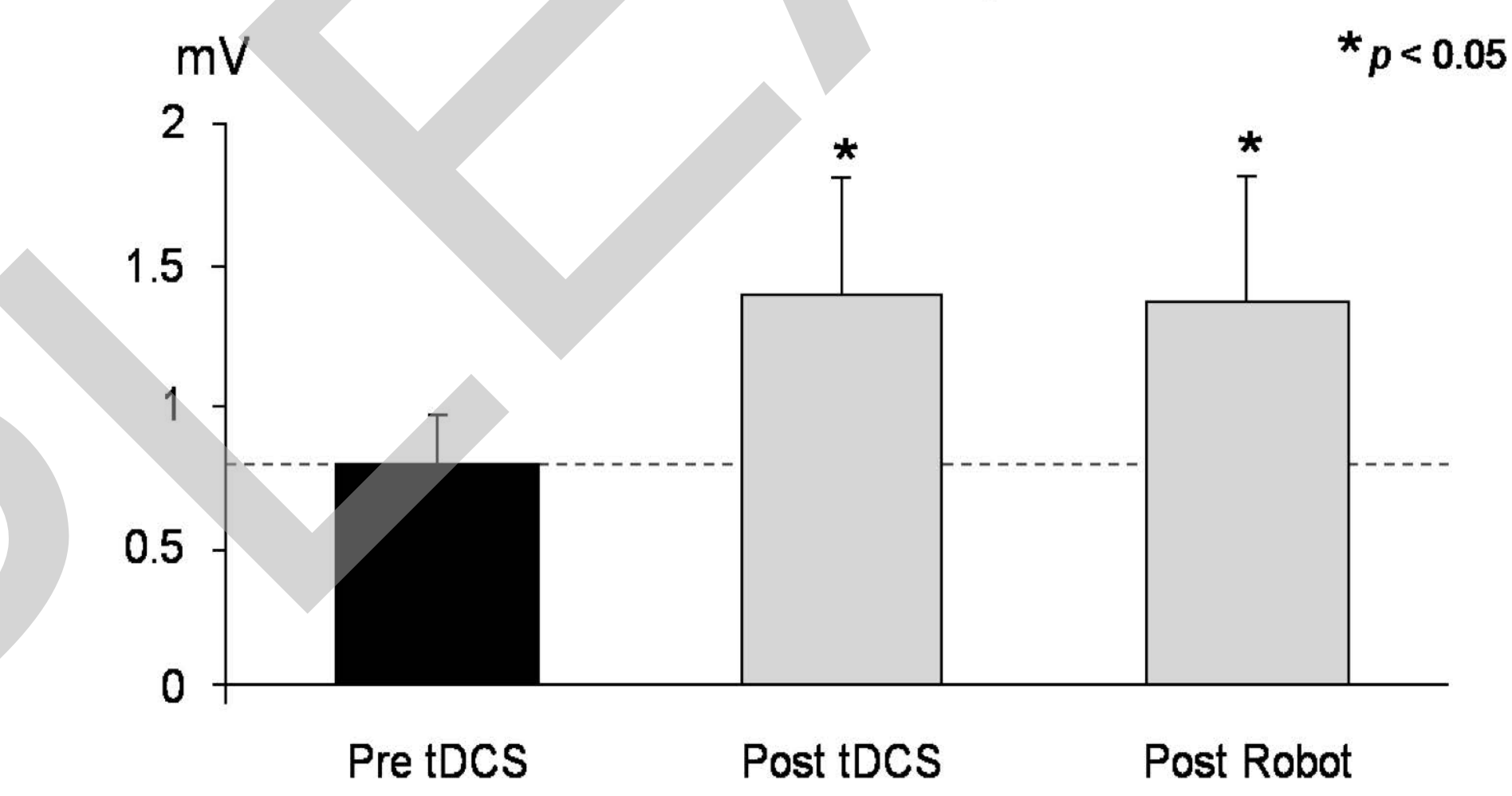
Key Findings: Effect of Intervention on Motor Performance



Timing of tDCS and behavioral therapy



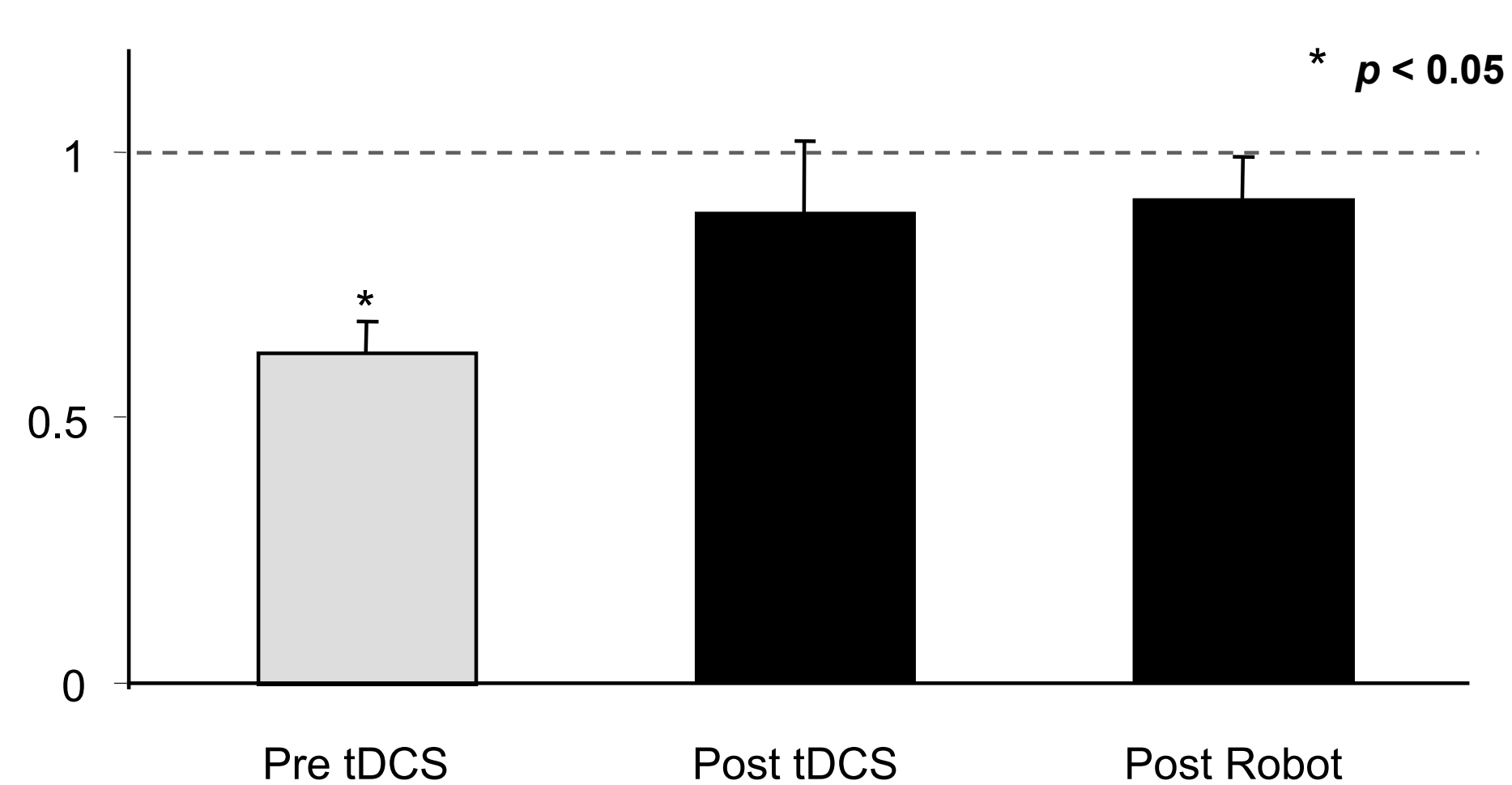
Mean MEP amplitude



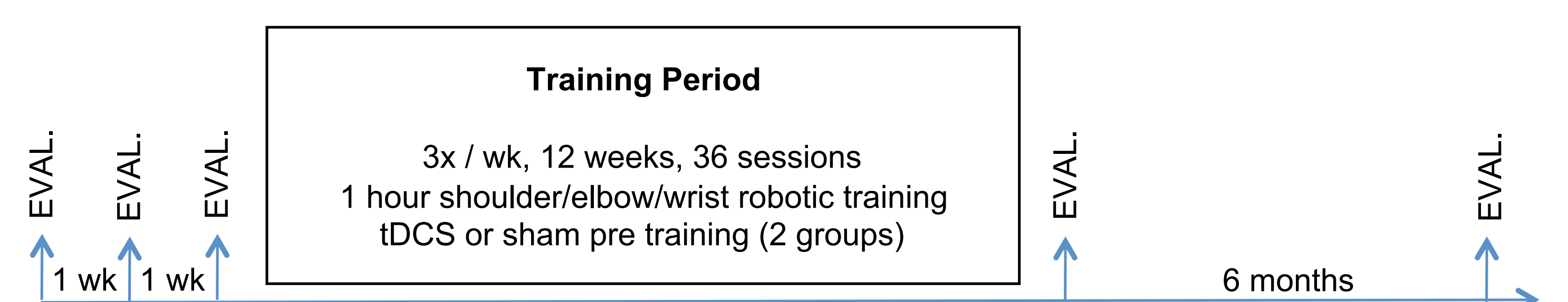
Edwards et al (2009)
Giacobbe et al (2013)

Conditioned / uncond MEP amplitude

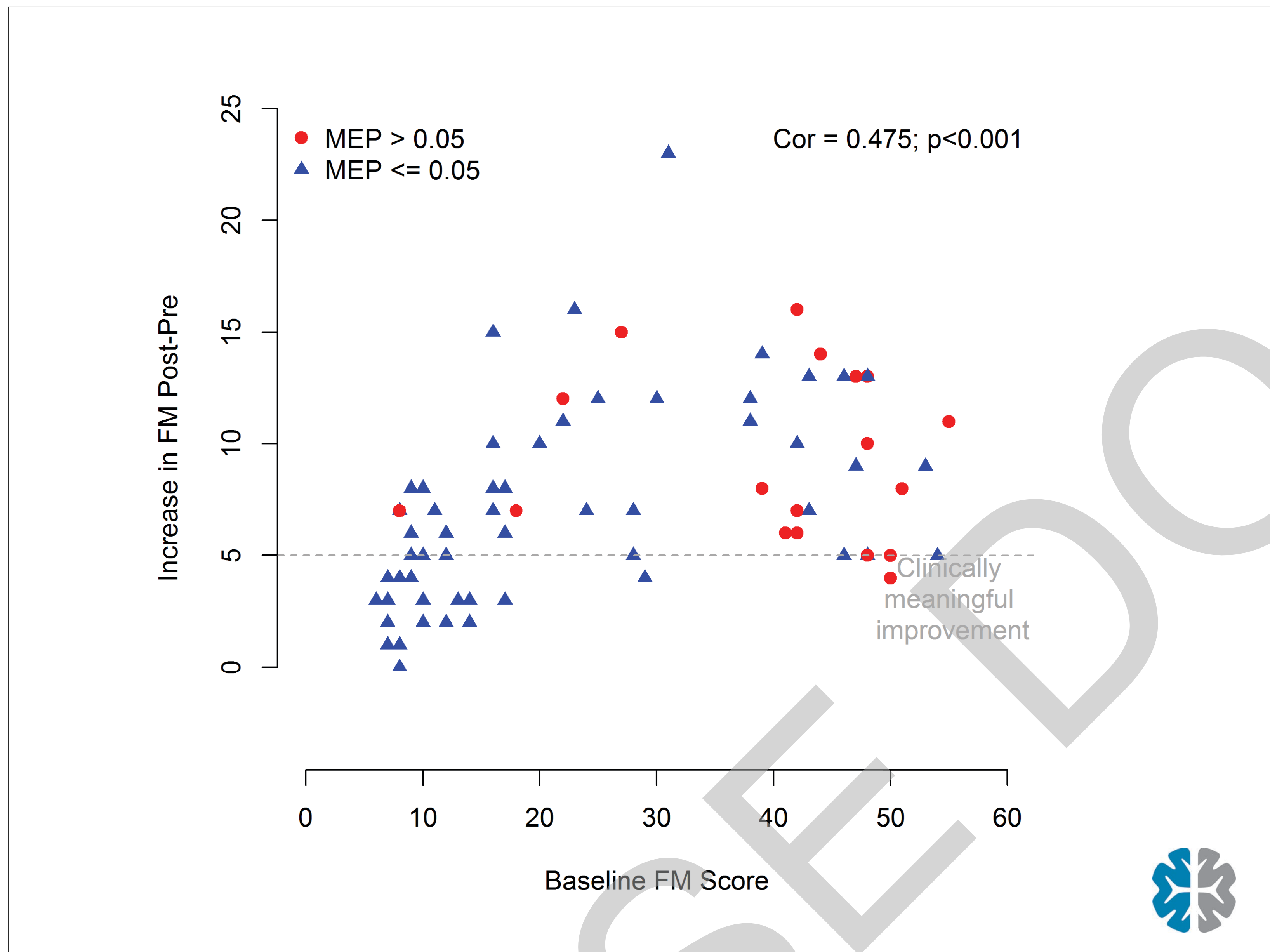
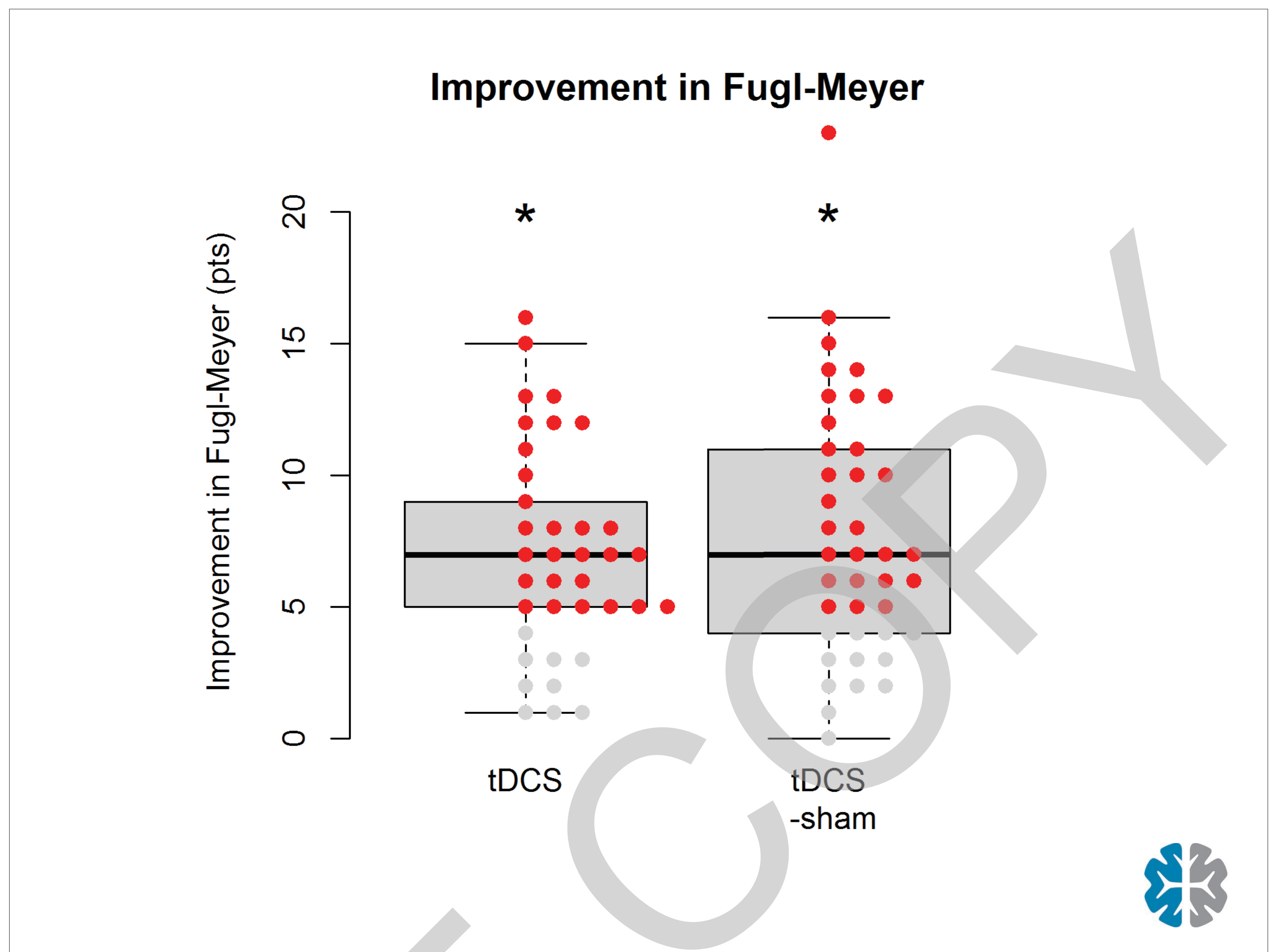
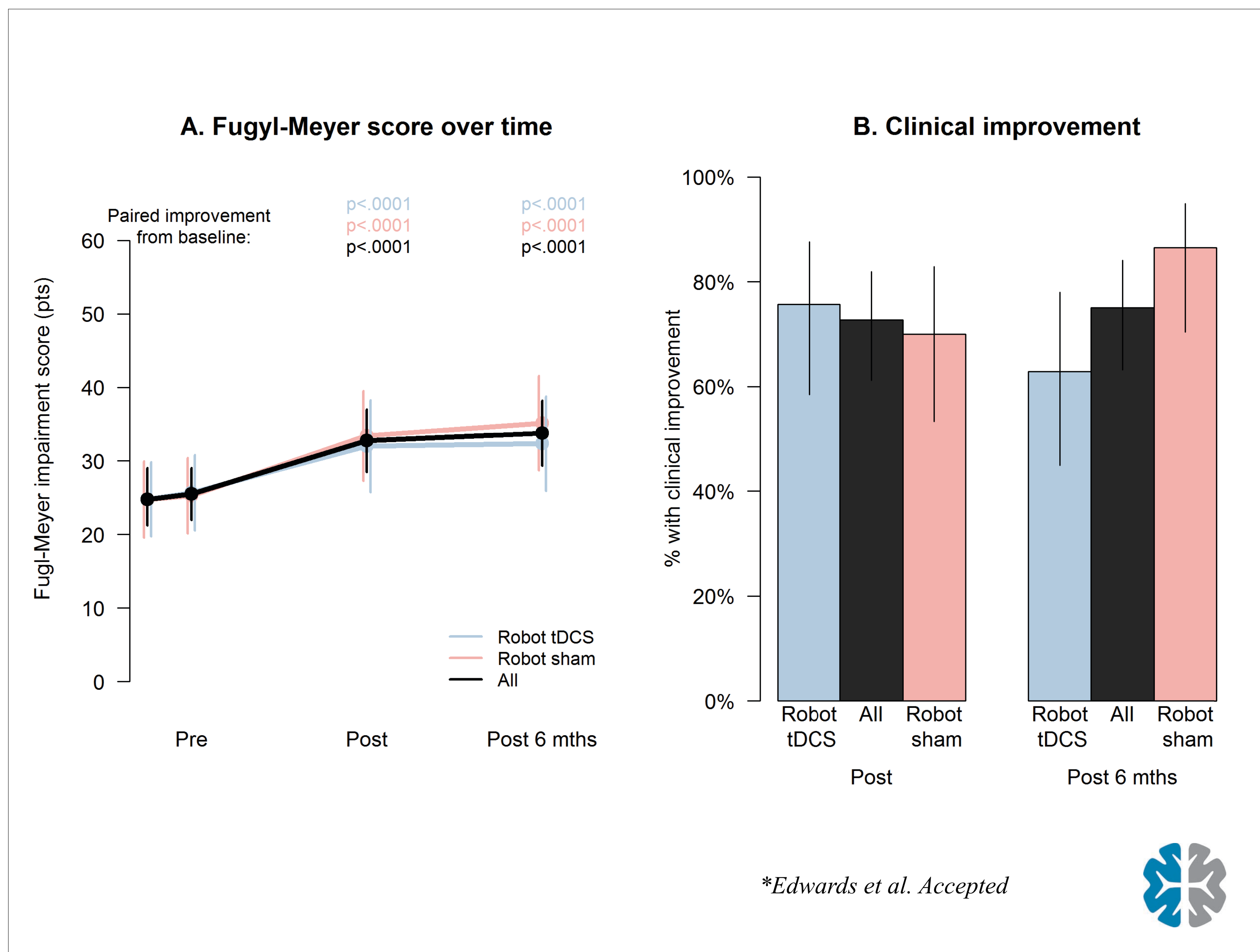
Group SICI Index



H₁: Robot+tDCS > Robot+SHAMtDCS on UEFM improvement



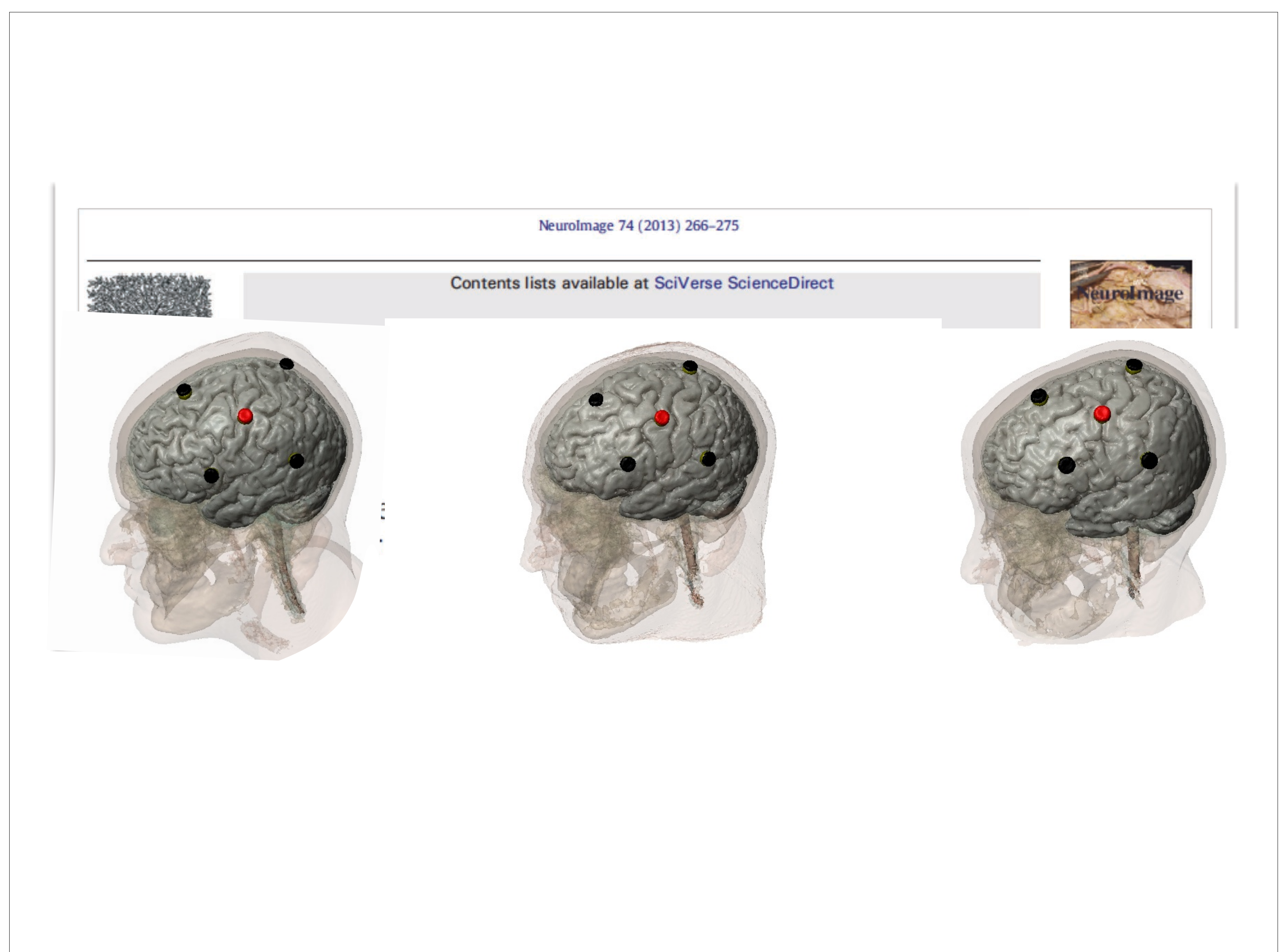
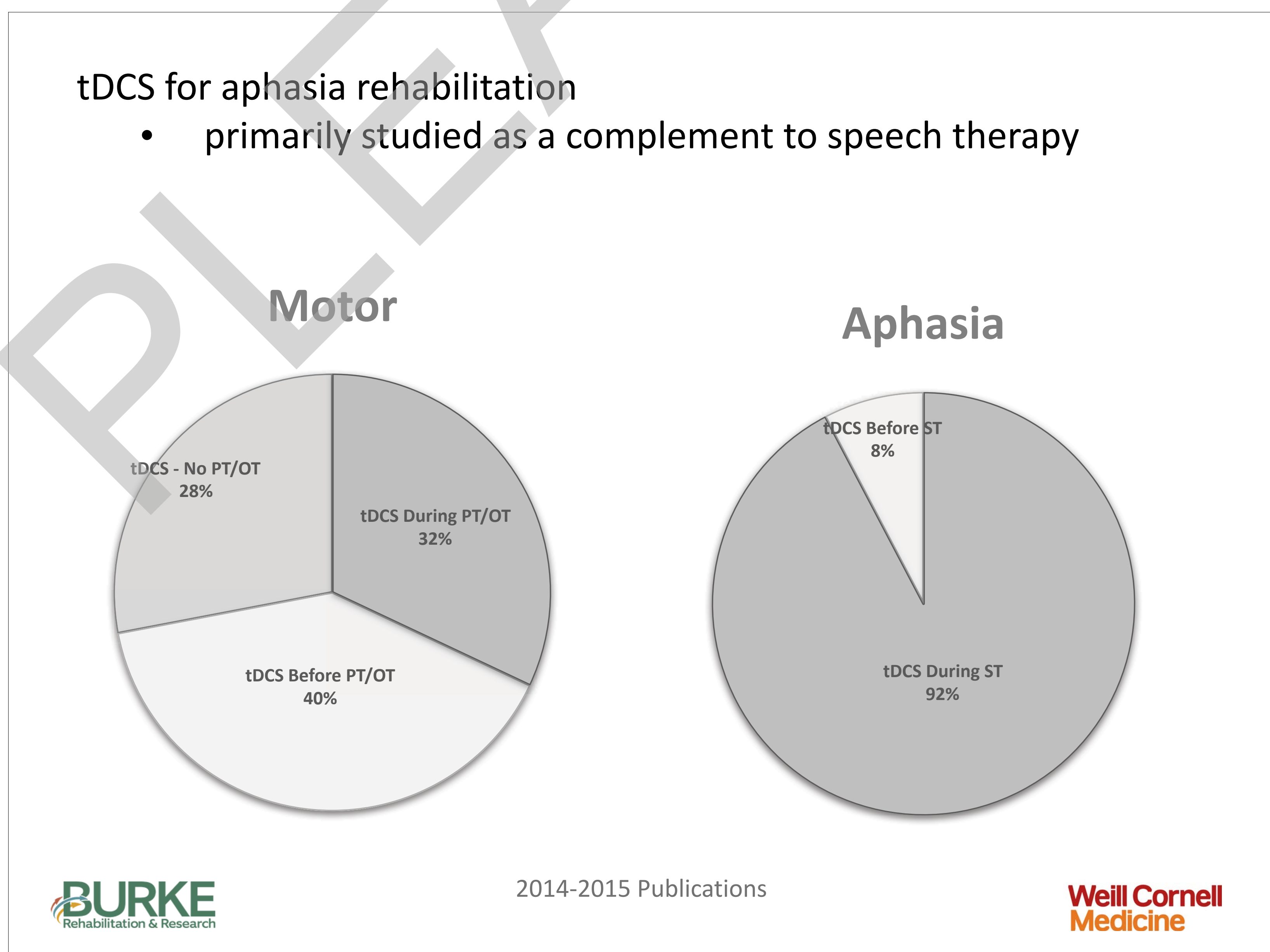
- 82 patients, right hemiparesis
- >6 mnths post first ischemic stroke
- Robotic protocol alternates S/E-wrist robot across sessions
- tDCS 2mA, 35cm². 0.9% NaCl soaked sponges

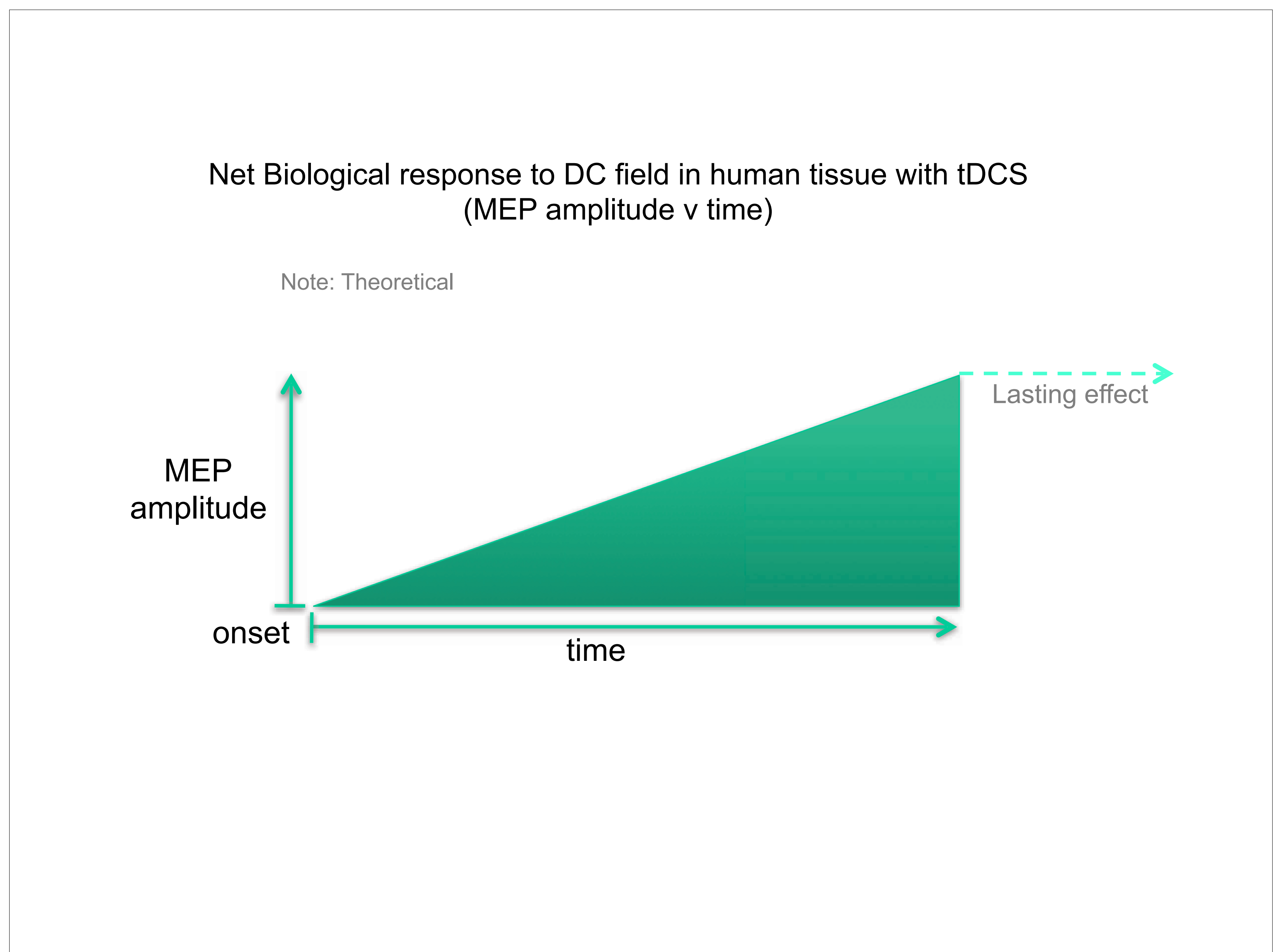
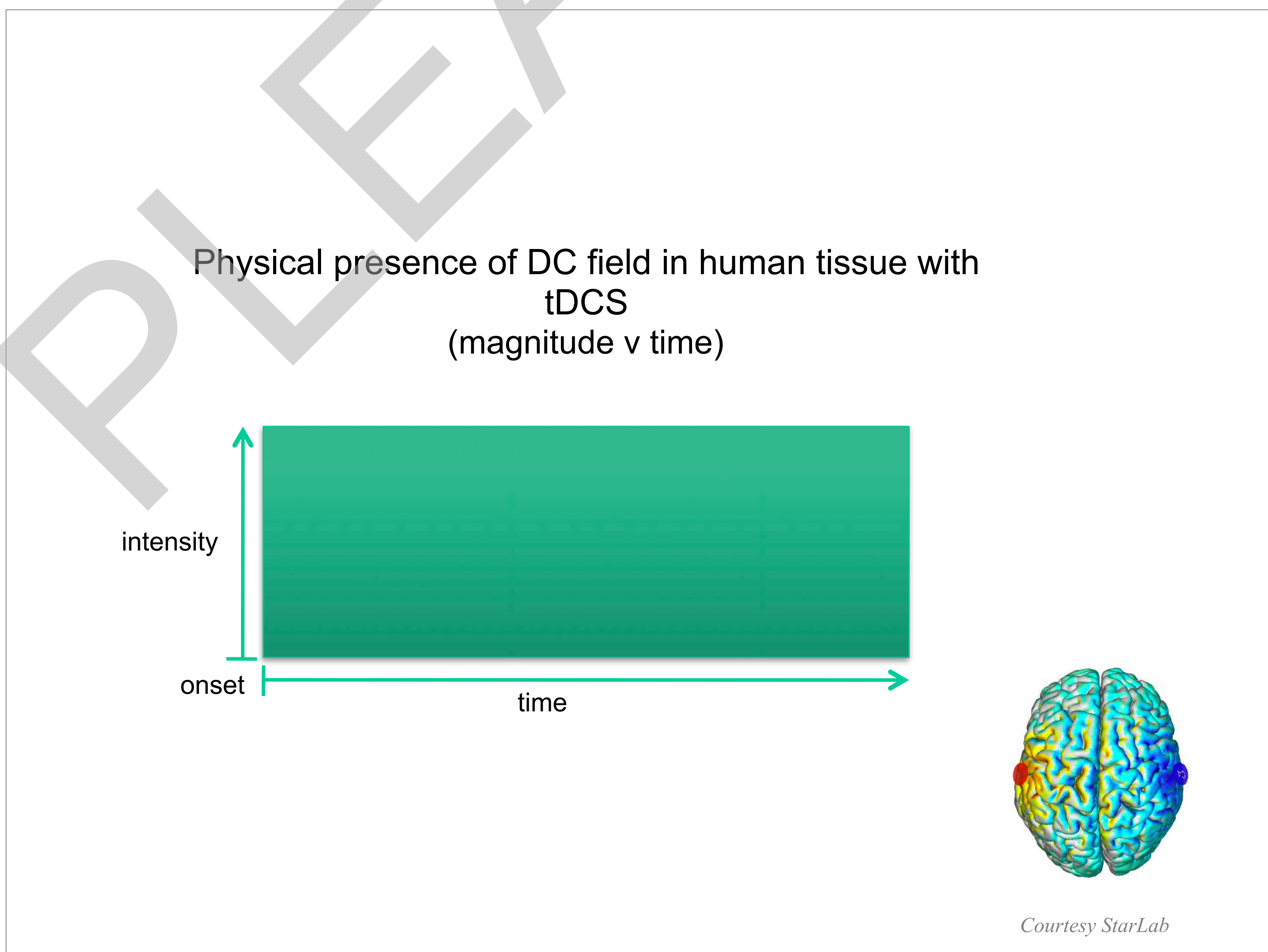
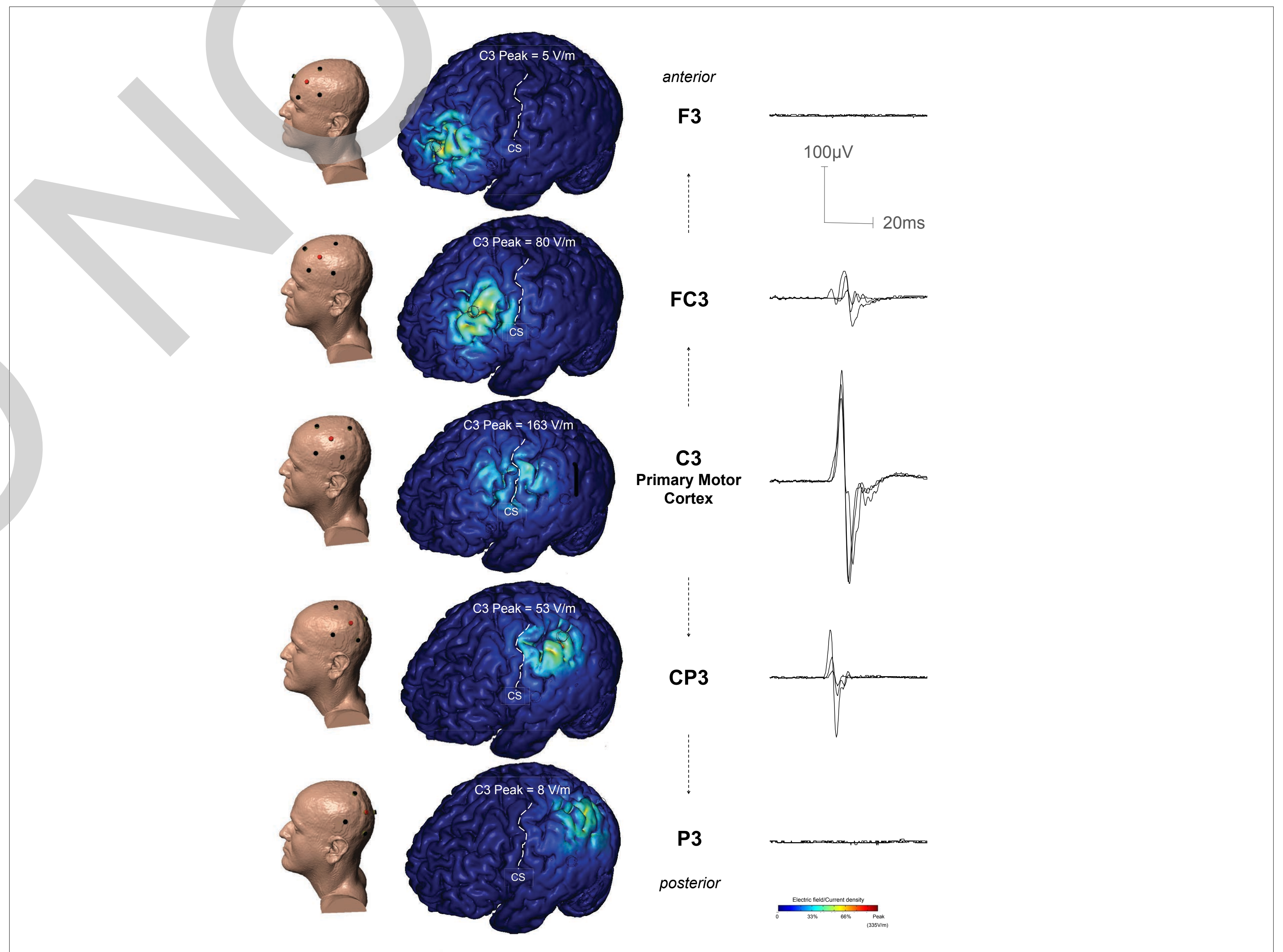
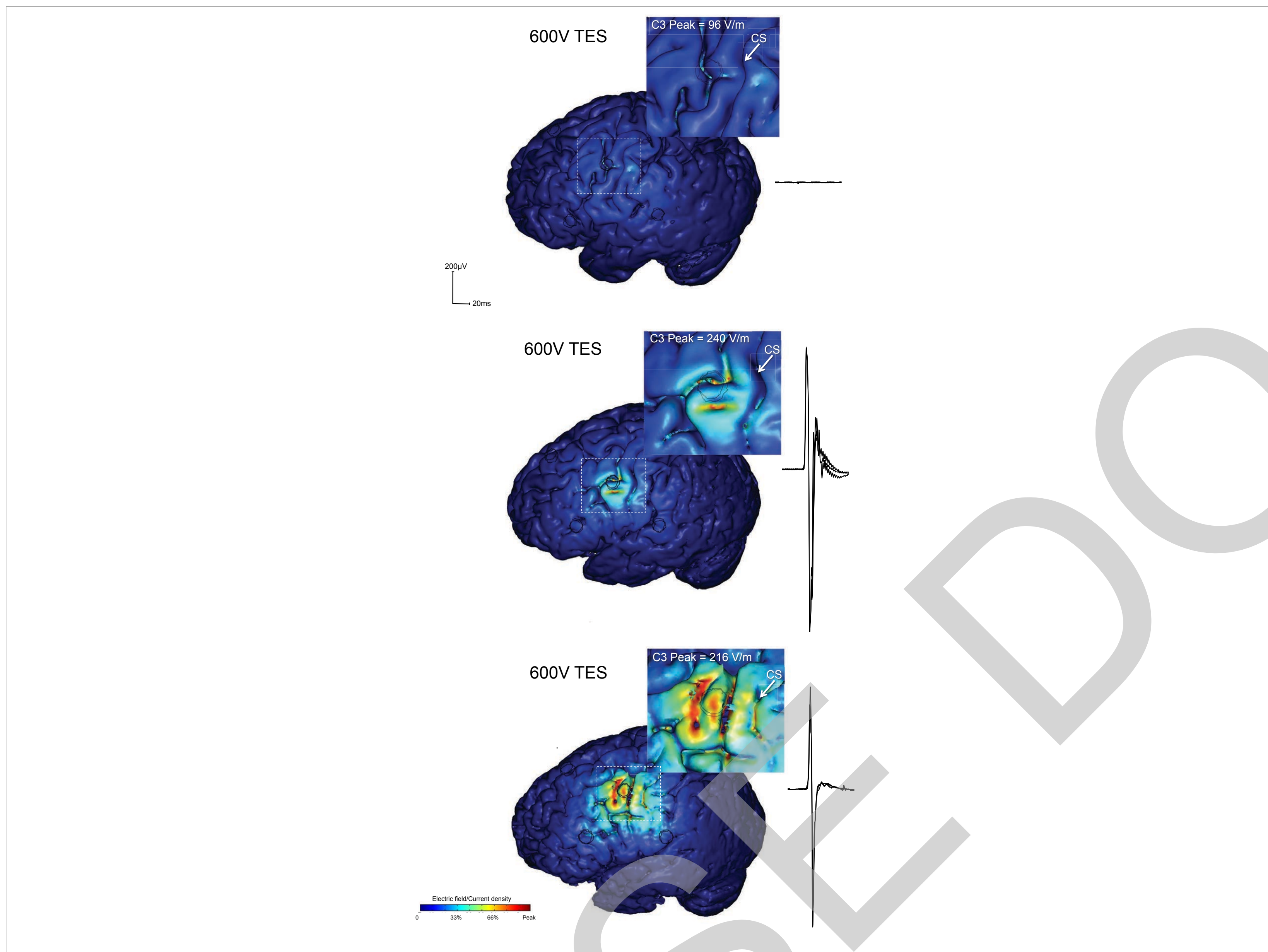
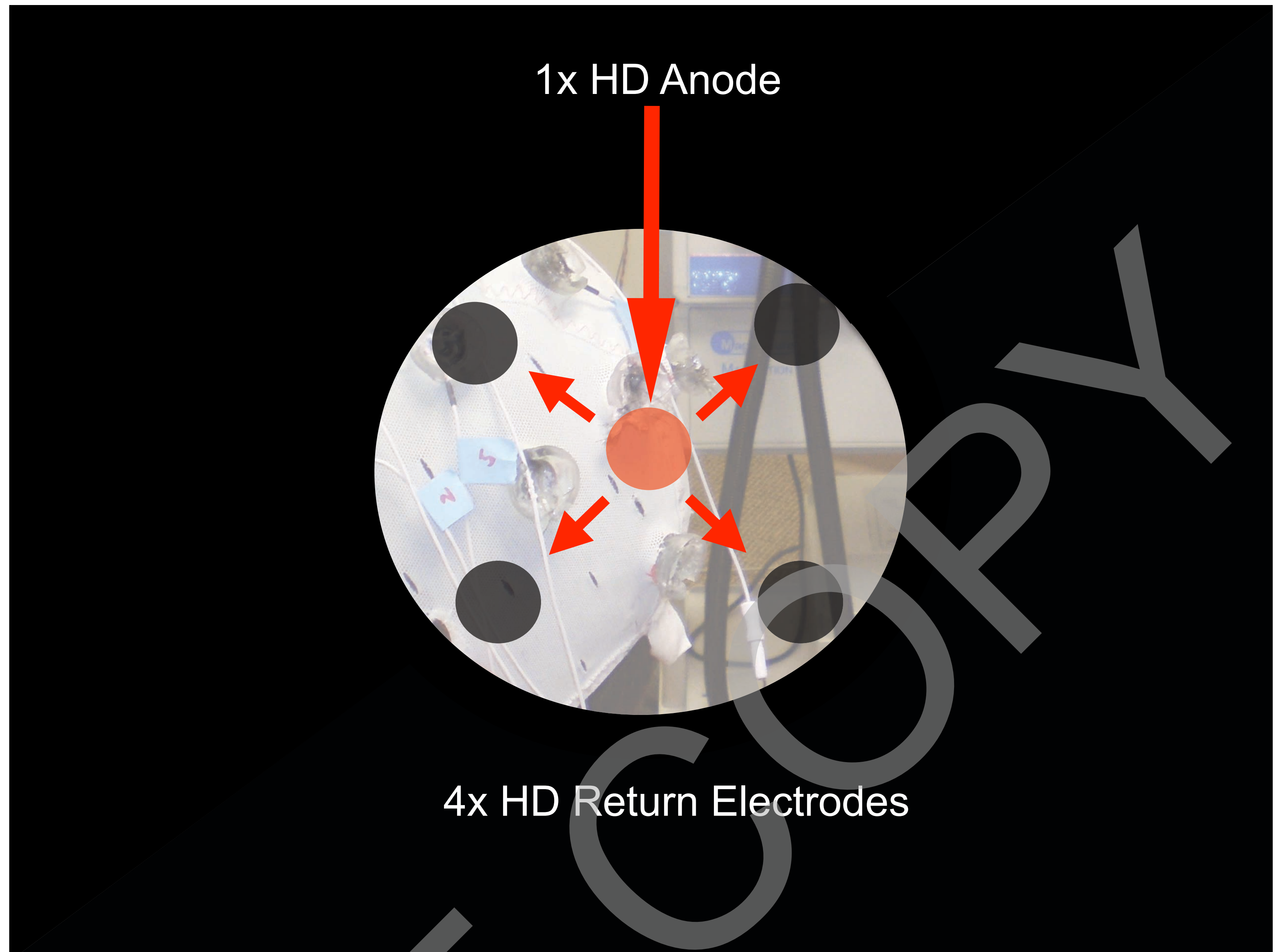
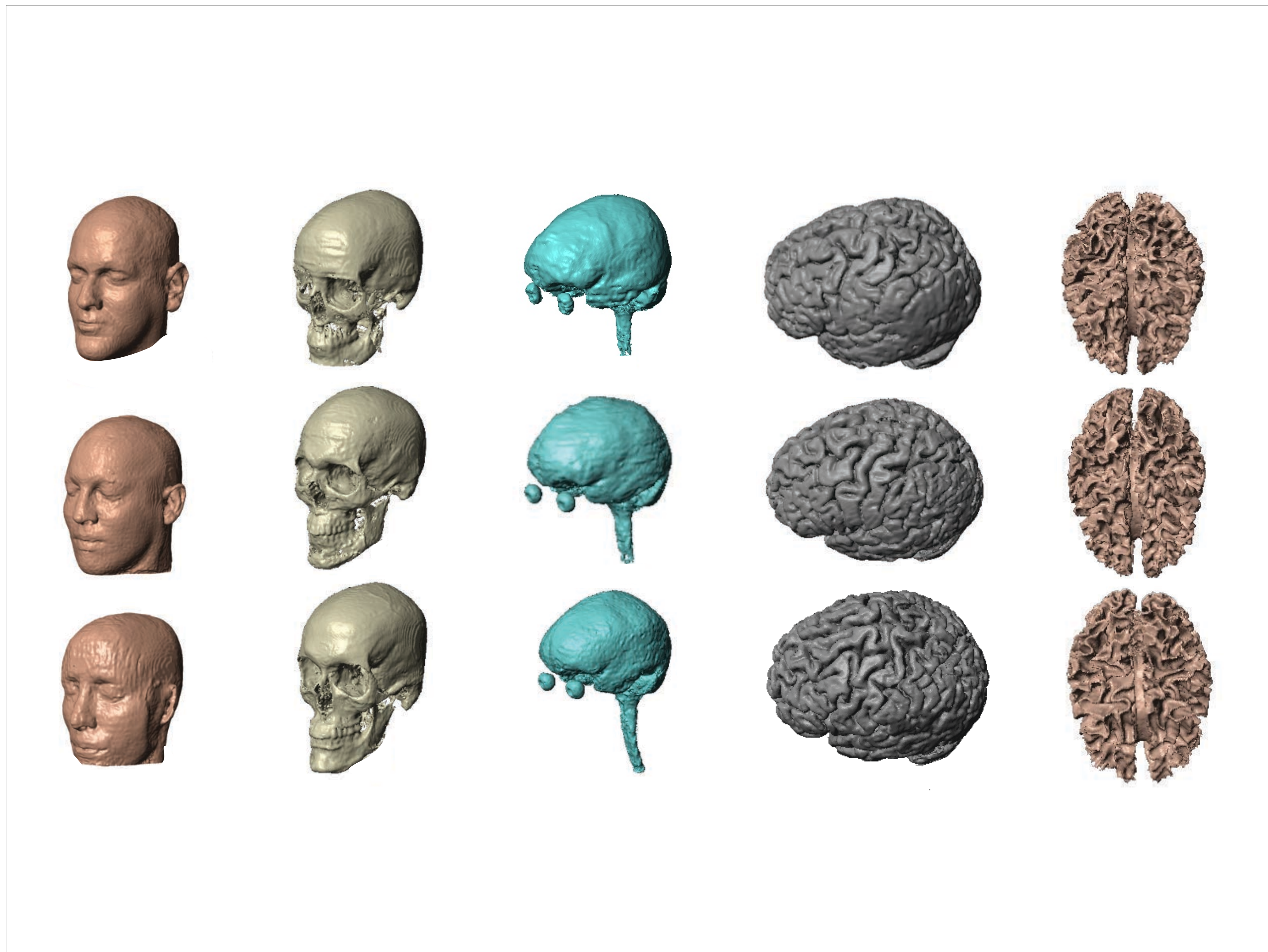


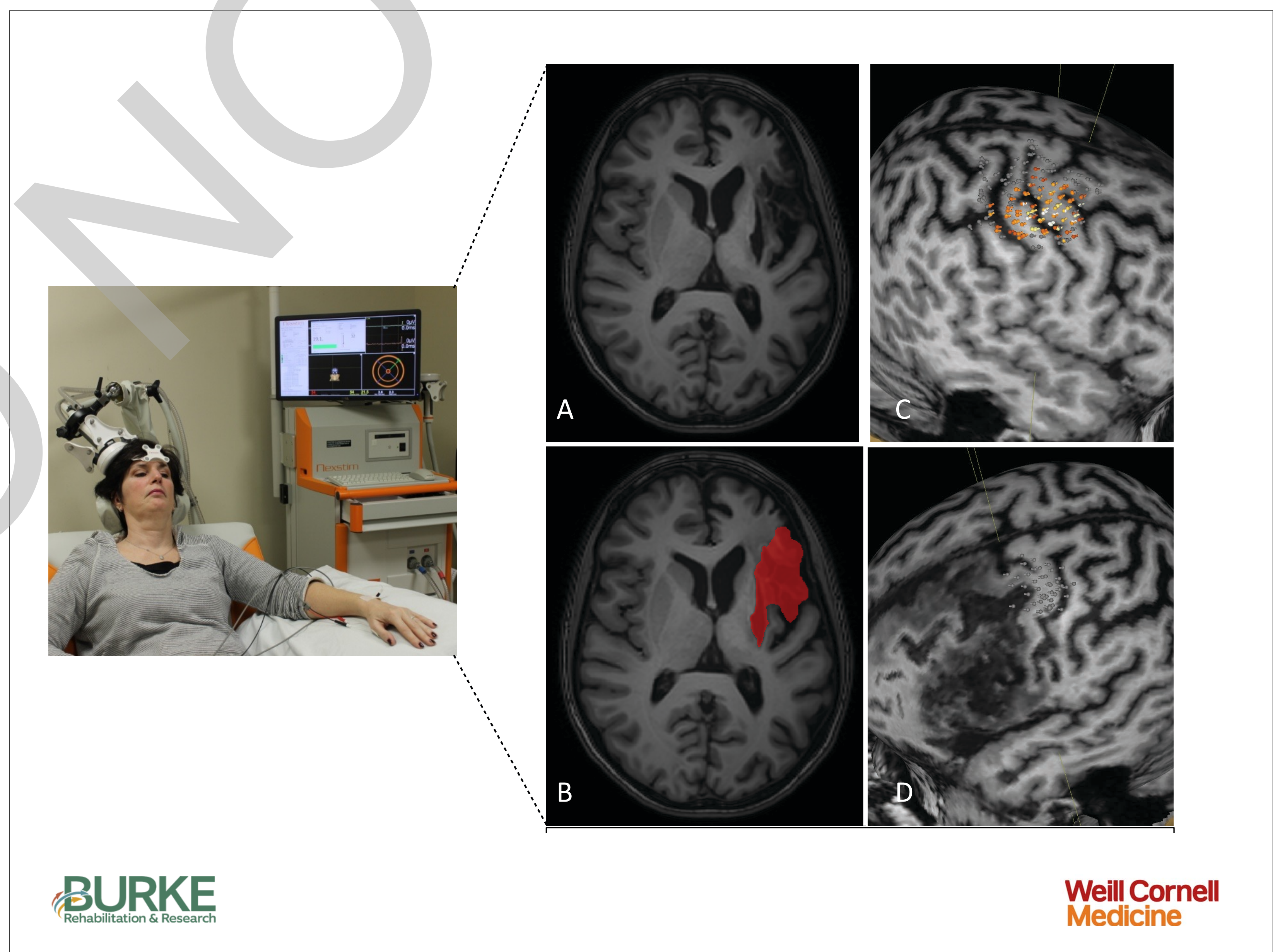
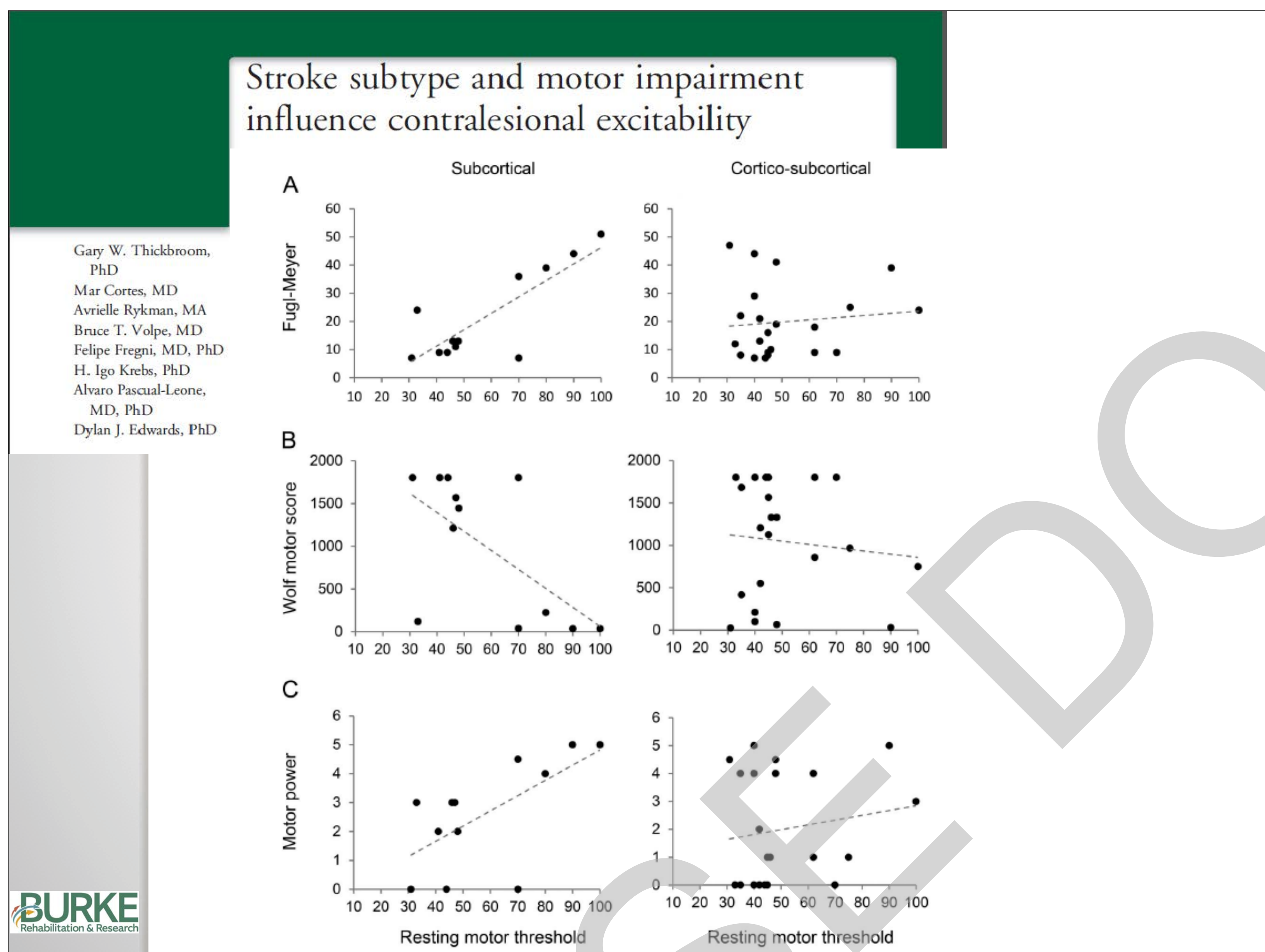
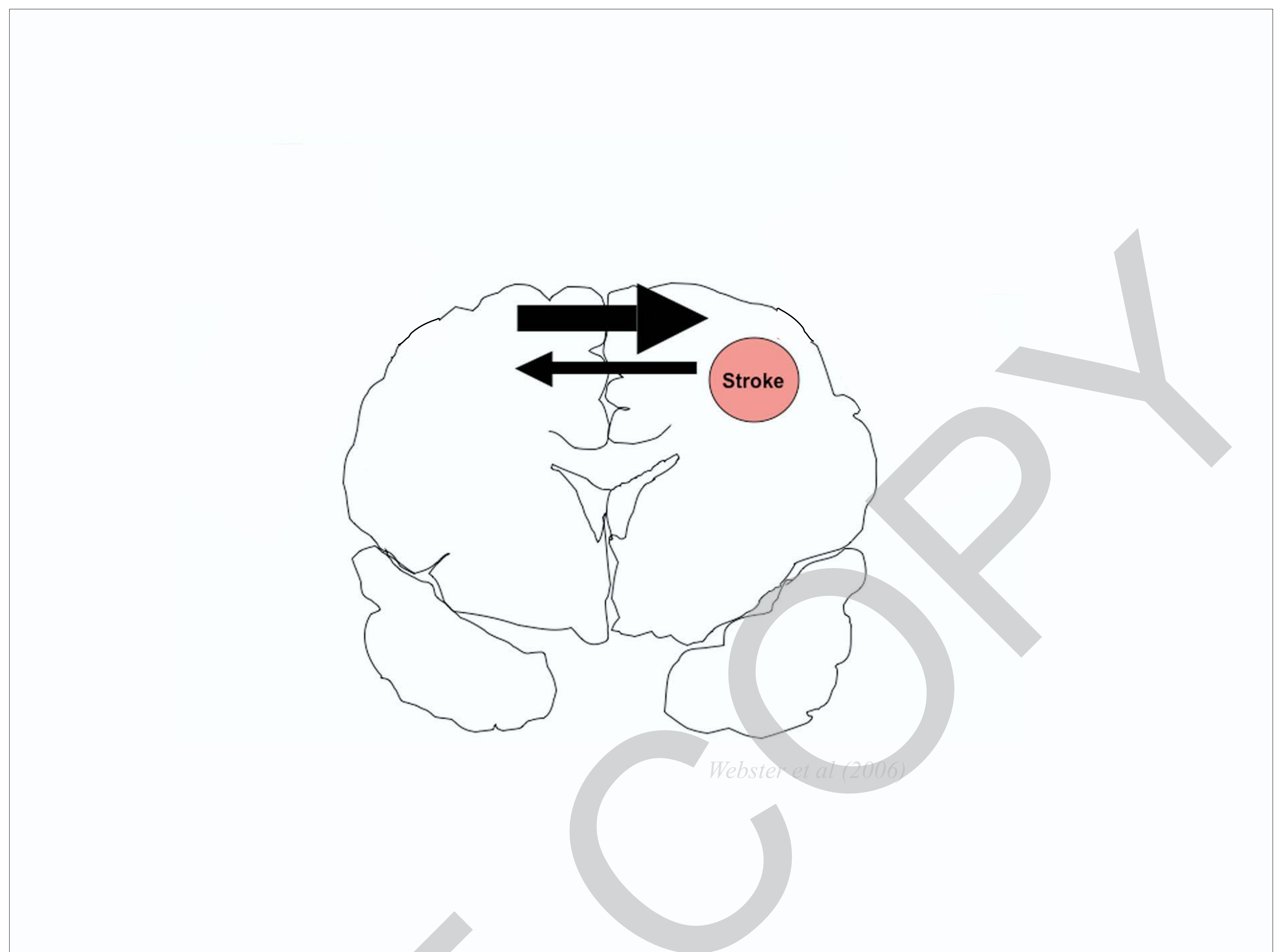
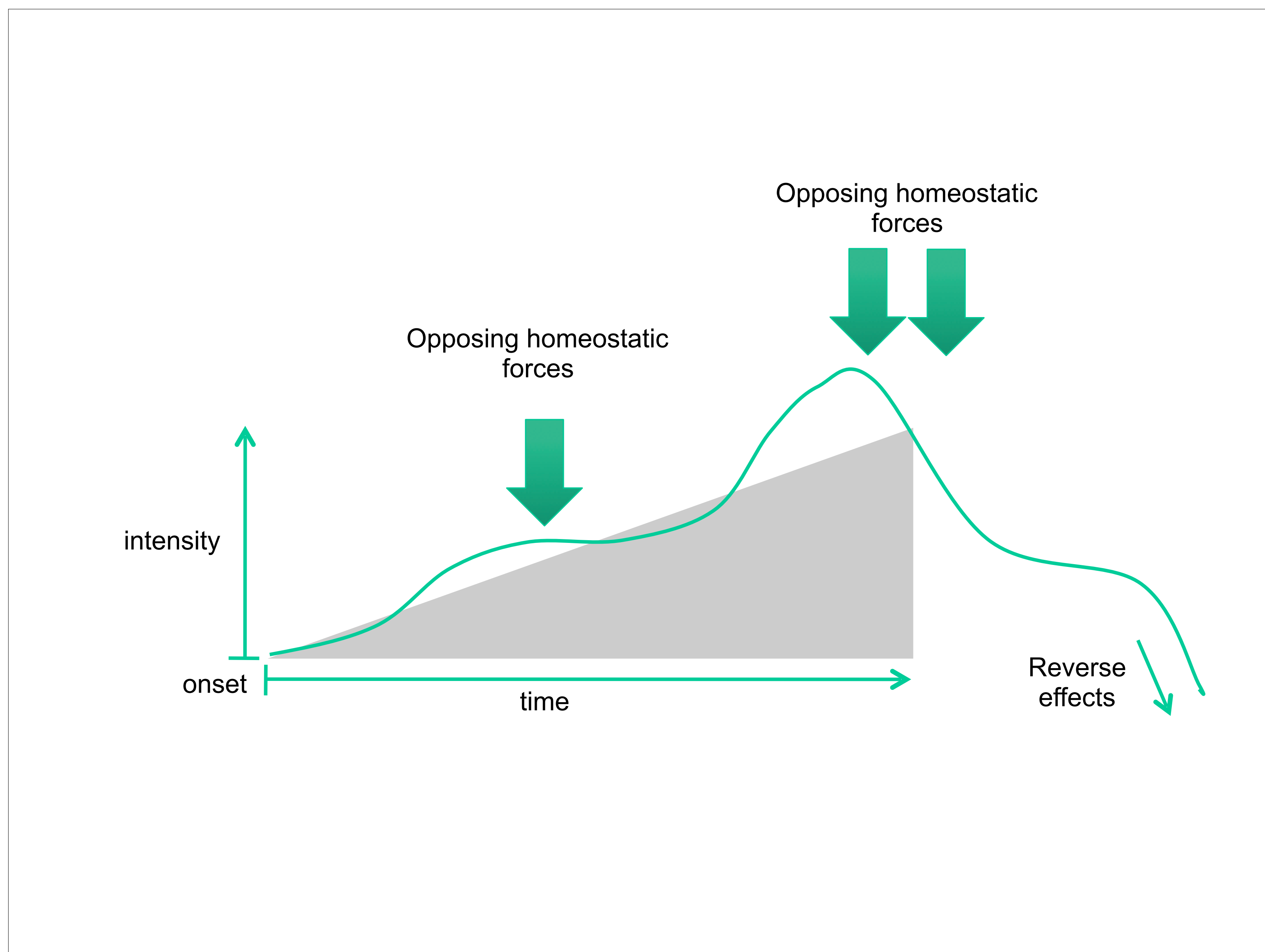
Northstar Trial

Parameters:

- 2 runs of 20 minutes of cathodal stimulation
- Electrode placement
 - Cathode – R pars triangularis
 - Anode – L supraorbital region
- Real – direct current of 2 mA
- Sham – direct current of 0.1 mA
- Worn during speech therapy







NeuroMeasure: a software package for quantification of cortical motor maps using frameless stereotaxic transcranial magnetic stimulation

Michael B. Gerber^{1*}, Alasdair C. McLean¹, Samuel J. Stephen¹, Alex G. Chalco¹, Usman M. Arshad¹, Gary Thickbroom², Josh Silverstein^{3, 4, 5, 2}, Zoe Tsagaris^{3, 2}, Amy Kuceyeski⁶, Kathleen Friel^{3, 4, 5, 2}, Taiza E. Santos⁷, Dylan J. Edwards^{8, 9, 7*}

Two brain surface maps labeled 'a' and 'b' showing motor activation. A color scale on the right ranges from 100 to 600 μV. Map 'a' shows a cluster of blue and green dots, while map 'b' shows a more extensive cluster of blue, green, and yellow dots.

* Gerber et al., 2019

Figure showing the NeuroMeasure software interface. It displays two data sets (Map1.xlsx and Map2.xlsx) and a 'Difference' map. A table of measurements is shown on the right.

Reference Positions:	Post->Ant	Right->Left	Inf->Sup	Distance
Ref->COG1	-6.9336	24.6250	55.3914	61.0137
Ref->COG2	-7.1785	24.2544	55.7750	61.2426
COG1->COG2	0.2448	0.3706	-0.3836	0.5869
Ref->Peak1	-6.1659	23.1581	57.6614	62.4432
Ref->Peak2	-7.2433	22.3590	58.6516	63.1854
Peak1->Peak2	1.0773	0.7991	-0.9902	1.6673

Values:	DataSet1	DataSet2	Difference
COG Value	297.9244	286.4357	11.4887
Peak Value	329.5403	534.8032	205.2629
Surface Area	1.8911e+03	1.7312e+03	159.9204
Volume Integral	1.9107e+05	2.3600e+05	4.4932e+04

Ref->Cursor: [-69.666, -5.505, 43.423]; 82.275
 Cursor Value: 0.000
 RMSE: 78.126

Thank you



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