

Neurophysiological impact of a fronto-temporal transcranial direct current stimulation (tDCS) in healthy subjects: a multimodal imaging approach

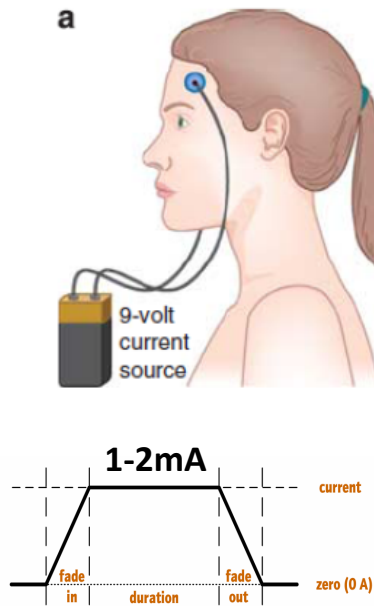
Team: Psychiatric disorders: from Resistance to Response (Ψ R2)

Clara FONTENEAU

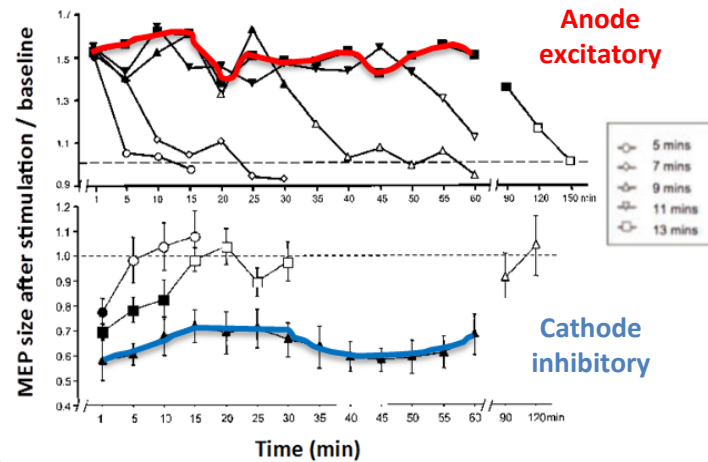


NEUROSTIMULATIONS

transcranial Direct Current Stimulation tDCS



Primary Motor Cortex Stimulation



Nitsche et al., Neurology, 2001

Nitsche et al., Clin Neurophysiol, 2003b

➤ Modulation of **cortical excitability**

Frontal Neurostimulations

DLPFC



An effect in **local** and in **connected regions**

1- Brain activity (ASL)

Antal et al., Human Brain Mapping, 2014
Stagg et al., J Neurosci, 2013

2- Functional connectivity in and between networks, like resting state networks (fMRI)

Keeser et al, Journal of Neuroscience, 2011
Saoite et al, Front Hum Neurosci, 2013
Mondino et al, Schiz Bull, 2015

3- Structural connectivity (DTI)

Peng et al., J Affective Disorders, 2012

4- Subcortical dopaminergic transmission (PET)

Strafella et al., Brain, 2003; Pogarell, J Psy Res, 2006
Cho&Strafella, PLoS One, 2009; Brunelin et al., Schizophr Res, 2011

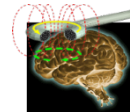
EXAMPLES



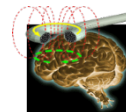
Healthy subjects



Healthy subjects

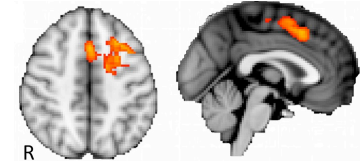


Patients with depression



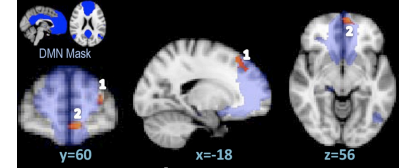
Healthy subjects

Anode vs Cathode



Stagg et al., J Neurosci, 2013

Default Mode Network (RSNc1) (After Real tDCS > Base1) > (After Sham tDCS > Base2)



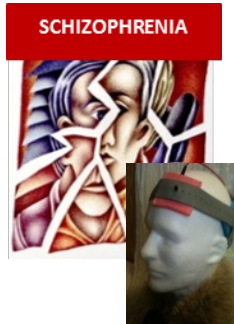
Keeser et al, J Neurosci, 2011

Modification of
anisotropic fraction
after rTMS

Subcortical
Dopamine ↗

**Temporal and functional
organisation ?**

PSYCHIATRY



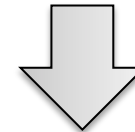
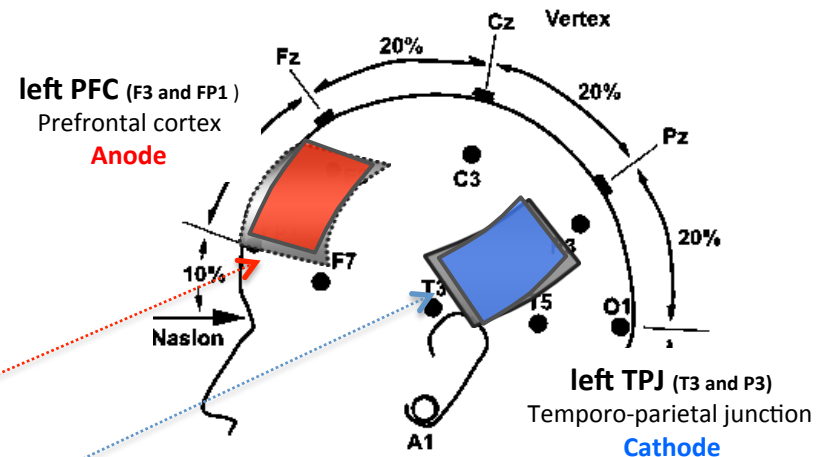
SCHIZOPHRENIA



SCHIZOPHRENIA

Pathophysiology

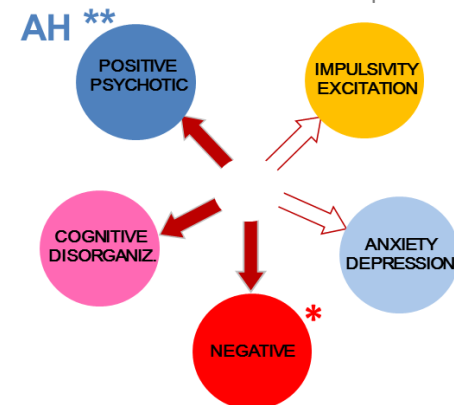
- **Hypoactivity of the left prefrontal cortex**
Lawrie et al, Biol Psychiatry, 2002
Sanfilipo et al, Arch Gen Psychiatry, 2000
- **Hyperactivity of the left temporo-parietal cortex**
Sibersweig et al., Nature, 1995
- **Dopaminergic alterations**
Brunelin et al, Curr Med Chem, 2013



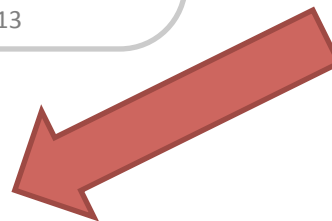
THERAPEUTIC EFFECTS

SCH *

Brunelin et al., Am J Psychiatry. 2012
Koops et al., Front Psychol, 2015



Neurophysiological correlates ?



PROJECT

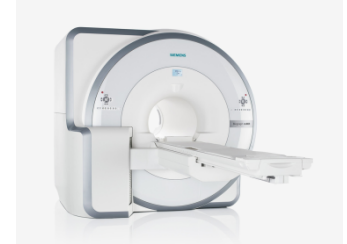
Temporal and functional organisation of the neurophysiological correlates of fronto-temporal tDCS ?

Creating a **coherent ensemble** of the effects of the stimulation combining:

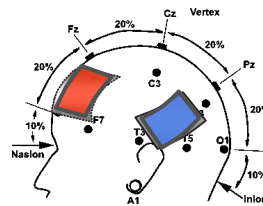
- 1) Structural and functional connectivity
- 2) Brain activity
- 3) Subcortical dopaminergic transmission

➤ Use of a **multimodal imaging system**

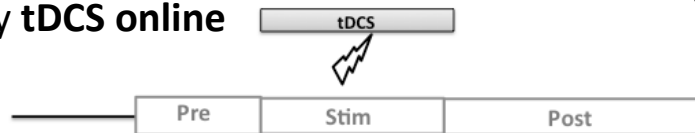
- ✓ Simultaneous imaging MRI & PET (MRI-PET Hybrid machine)
- ✓ Link between connectivity, brain activity and subcortical dopaminergic transmission



➤ Adopt the **fronto-temporal montage**



➤ Apply **tDCS online**

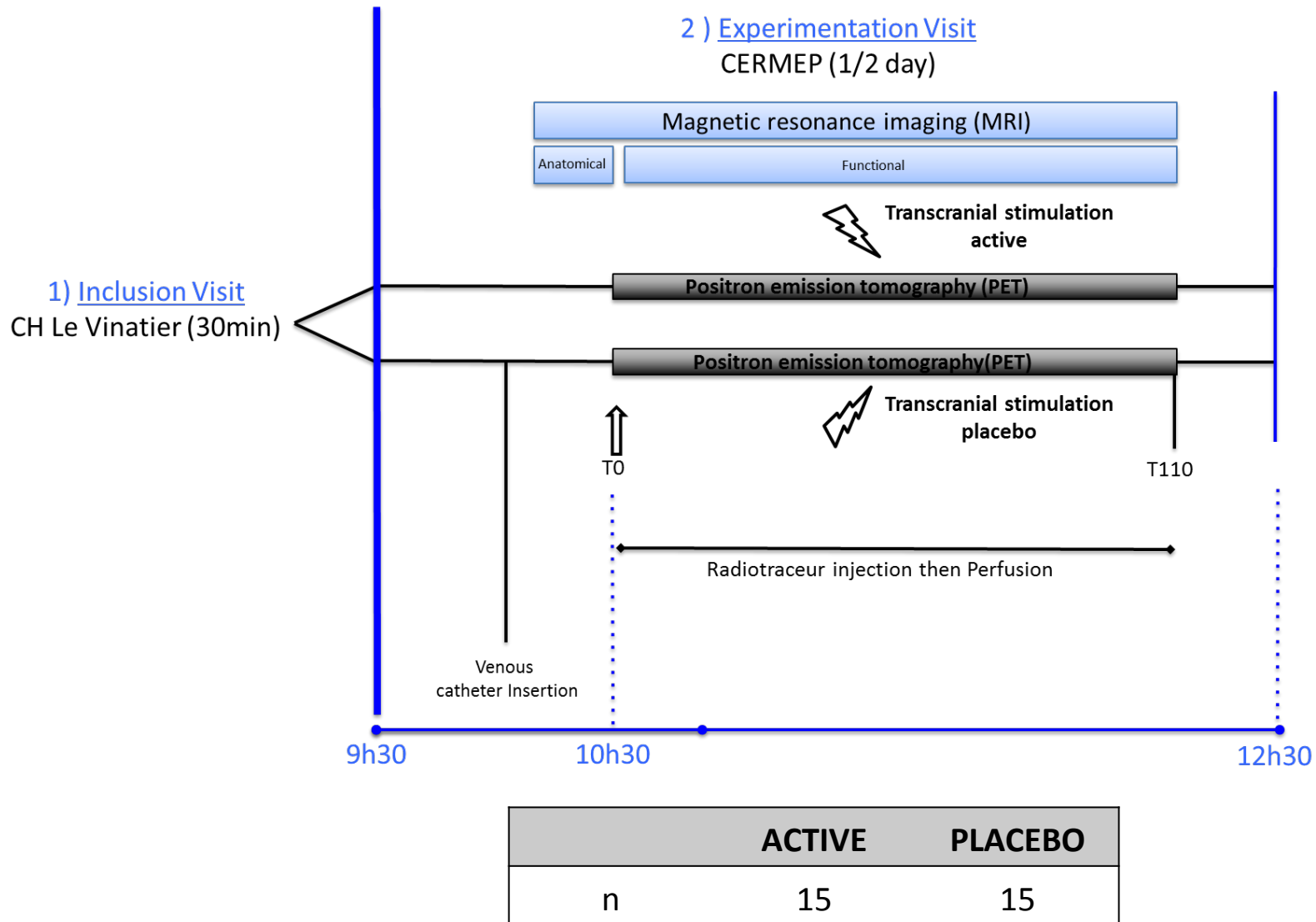


➤ First step in **healthy subjects**

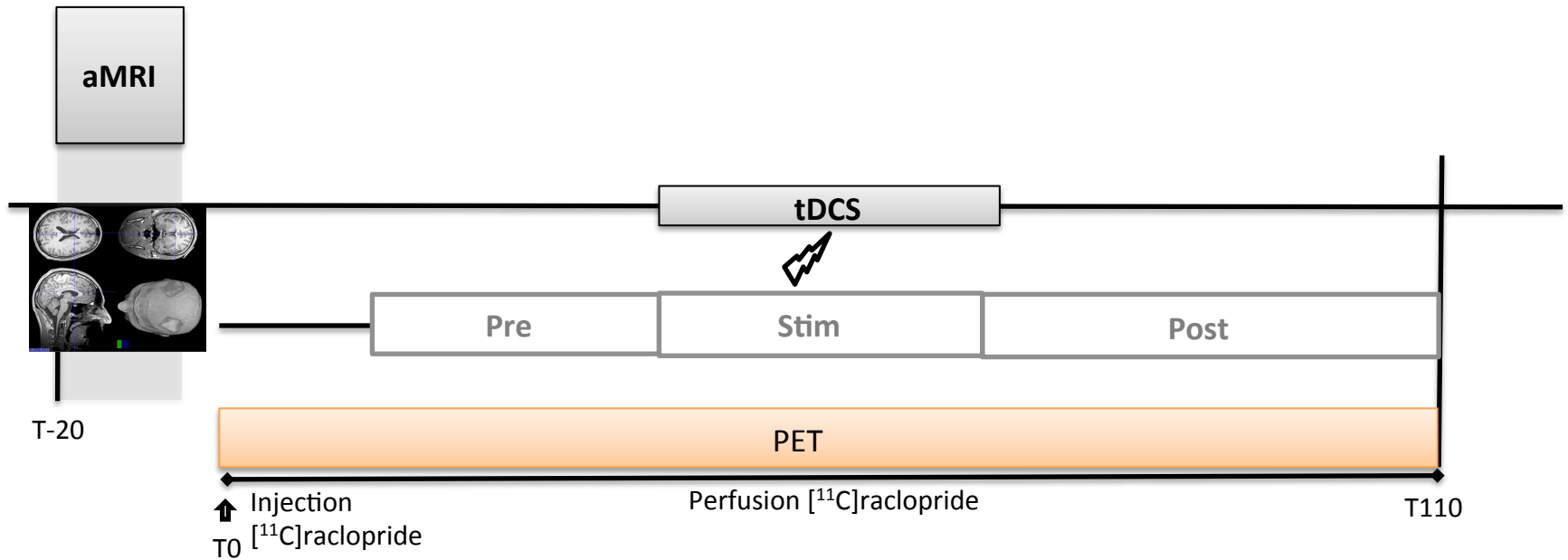


PROJECT

- Randomized, double blind study, 2-arm parallel groups in healthy subjects

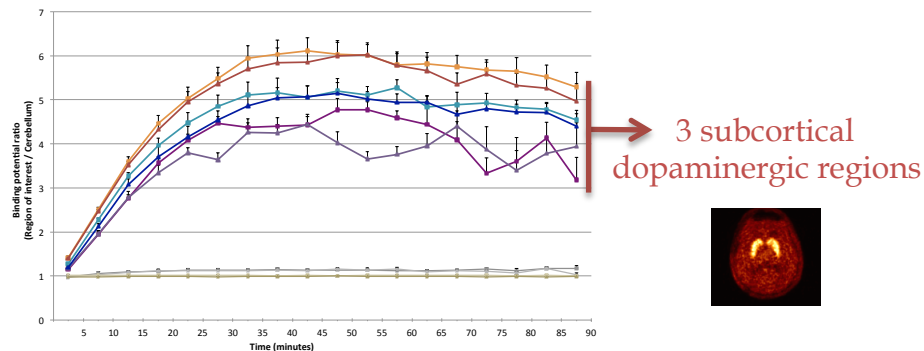


Neurophysiological impact of a fronto-temporal transcranial direct current stimulation in healthy subjects

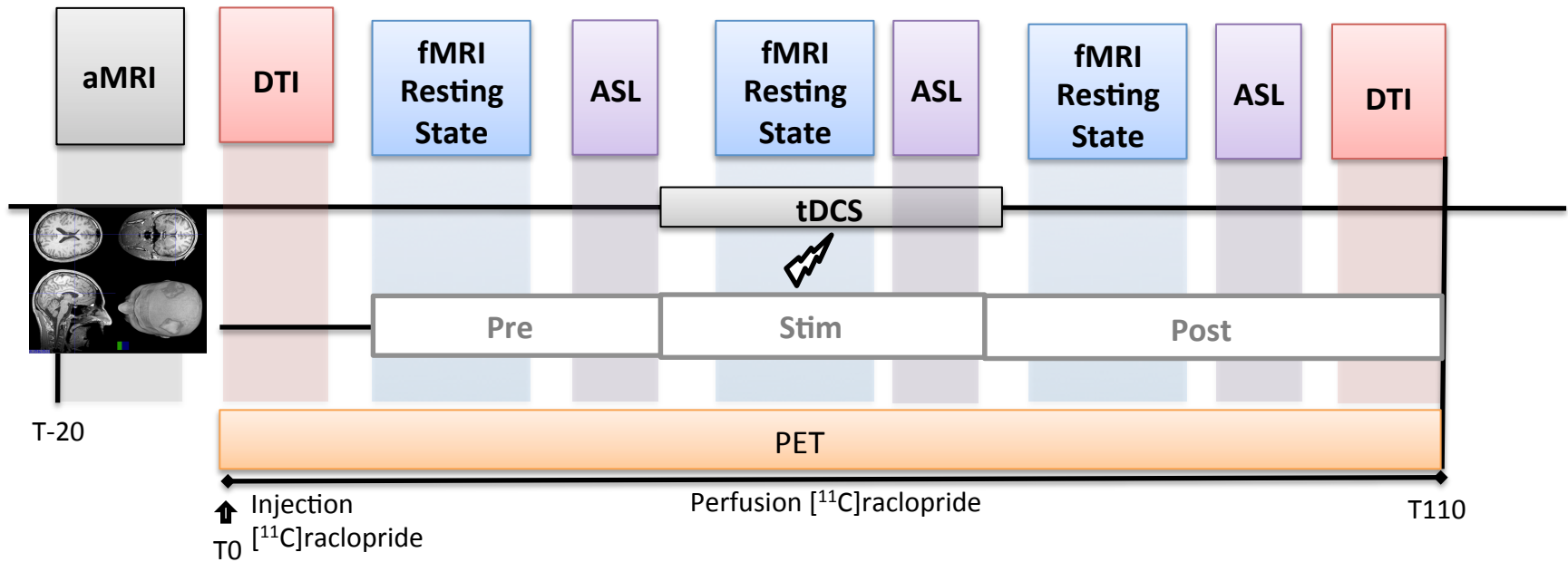


PET – Raclopride:

Dopaminergic transmission in basal ganglia
→ **Static** and **Dynamic** approach



Neurophysiological impact of a fronto-temporal transcranial direct current stimulation in healthy subjects



MRI – Sequences:

Resting state fMRI – functional connectivity

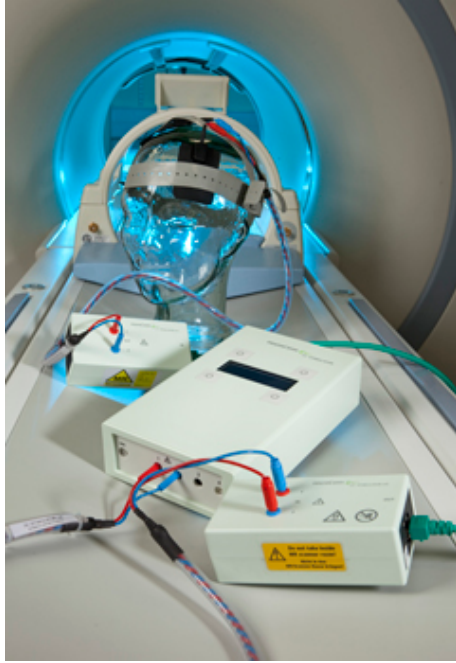
ASL – Brain activity

DTI – Structural connectivity

→ Region of interest analysis

- stimulated regions: **DLPFC & TPJ**
- connected regions: **Resting state networks**

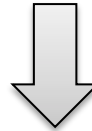
Feasibility



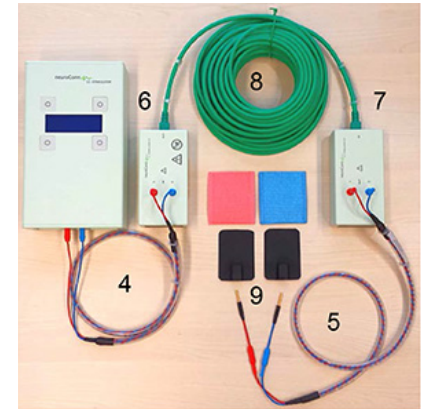
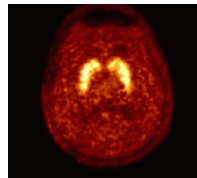
**Stimulator used in
international publications**

Antal, et al., Neurolmage, 2011
Sehm et al., Front Hum Neurosci, 2013
Stagg et al., J Neurosci, 2013
Antal et al., Neurolmage, 2014
Meinzer et al., Journal of visualized experience, 2014

- **Ethics & Radiotracer
Autorisation
(ANSM – CPP)**
- **Neurodis Funding**
- **tDCS online**
MRI compatible, CE
NeuroConn DC stimulator MR



First subjects
N=3



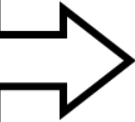
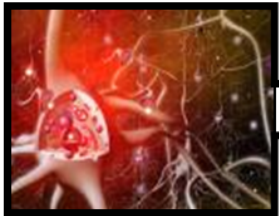
- Collaboration CERMEP

Project impact

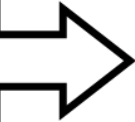


MRI & PET

NEUROBIOLOGY



NEUROPHYSIOLOGY



Coherent ensemble

Dopamine (TEP)



Resting state networks (IRMf)
Perfusion (ASL)
Structural connectivity (DTI)

Imaging biomarker

« Normal brain »



Pathological brain



*Neurophysiological effects of
fronto-temporal tDCS*



Improved psychiatric
symptoms



Multiple applications

Original Article | April 1, 2013

The Sertraline vs Electrical Current Therapy for Treating Depression Clinical Study

Resu

Andre R
Oliveira,
MD, PhD

JAMA P **ELSEVIER**



Contents lists available at bouaziznoomane@gmail.com

Drug and Alcohol Dependence

The World Journal of Biological Psychiatry, 2014; Early Online: 1–15



frontiers in
HUMAN NEUROSCIENCE

REVIEW ARTICLE
published: 14 August 2013
doi: 10.3389/fnhum.2013.00449



Non-invasive brain stimulation can induce paradoxical facilitation. Are these neuroenhancements transferable and meaningful to security services?

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ALVAO¹,