

THE UNIVERSITY of NORTH CAROLINA at CHAPEL HILL



### Mechanism of Transcranial Alternating Current Stimulation (tACS)

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### Conflicts of Interest

- UNC owns IP related with FF as the lead inventor.
- UNC has determined the absence of a conflict of ۲ interest (COI) for the majority of work presented here and has determined a "COI with administrative considerations" for the clinical trials in the Frohlich Lab.
- FF is the founder, chief scientific officer, and majority owner of Pulvinar Neuro LLC. We provide NEUROTECHNOLOGY FOR THE FUTURE solutions for transcranial current stimulation research.
- I speak with many companies and have received industry funding from Tal Medical (travel + research).
- I frequently travel and give presentations. I typically receive reimbursement and a stipend.

### PULVINAR NEURO



### NETWORK NEUROSCIENCE





FLAVIO FRÖHLICH

### Berger 1929

Abb. 4. 40 jähriger Mann. Große linksseitige, von der Stirn bis in die Parietalgegend reichende Knochenlücke. Doppelspulengalvanometer. Kondensation. Nadelelektroden subcutan im Bereich der Knochenlücke, 4,5 cm voneinander entfernt. Oben Schwankungen der epidural abgeleiteten Kurve, unten Zeit in <sup>1</sup>/<sub>10</sub> Sekunden.



# 

Transcranial Alternating Current Stimulation

# VERTICAL INTEGRATION

### Patients



TRACTABILITY

## Model Systems

**Computer Simulations** 



Brain Stimulation, Human Neurophysiology



*In vivo* (Animal) Electrophysiology

In vitro (Animal)

Electrophysiology

**Clinical Trials** 







# RATIONAL DESIGN











Caution: Most tACS literature refers to the peak-to-peak amplitude as amplitude.

# Today's Theory





#### Frohlich and McCormick, 2010





Frohlich and McCormick, 2010





Ali et al. 2013





Ali et al. 2013



Frohlich 2014



#### Frohlich 2014









Kutchko and Frohlich 2013





Kutchko and Frohlich 2013





Schmidt et al. 2015







Schmidt et al. 2015





Alagapan et al. 2016





"Eyes Closed"

Visual Working Memory Task





Alagapan et al. 2016

### 1. Outlasting effects

- 2. Arnold tongue beyond model
- 3. Interacting oscillators
- 4. State-dependence
- 5. Waveform modifiers

Charles Zhou Caroline Lustenberger Sankar Alagapan Yuhui Li Guoshi Li Ehsan Negahbani Juliann Mellin Courtney Lugo Morgan Alexander Philipp Lustenberger Iain Stitt Supritha Dugyala Toheed Khan Quique Toloza Nadia Mishal Mia DeMarco Matt Mattoni Jhana Parikh Hemanth Ambala Florian Schertenleib Carolyn Rapp Alexandra Vossen Franz Hamilton Jessica Page Maadhurya Duvvuri

#### Alumni Lab Members

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> Collaborators ECOG: Dr. Haewon Shin Sleep Spindles: Dr. Bradley Vaughn Modeling ECOG: Dr. Jeremy Lefebvre Electric Field Spatial Targeting: Dr. Angel Peterchev SCZ Clinical Trial: Dr. Fred Jarskog, Dr. John Gilmore Mood Disorders Clinical Trials: Dr. David Rubinow

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