



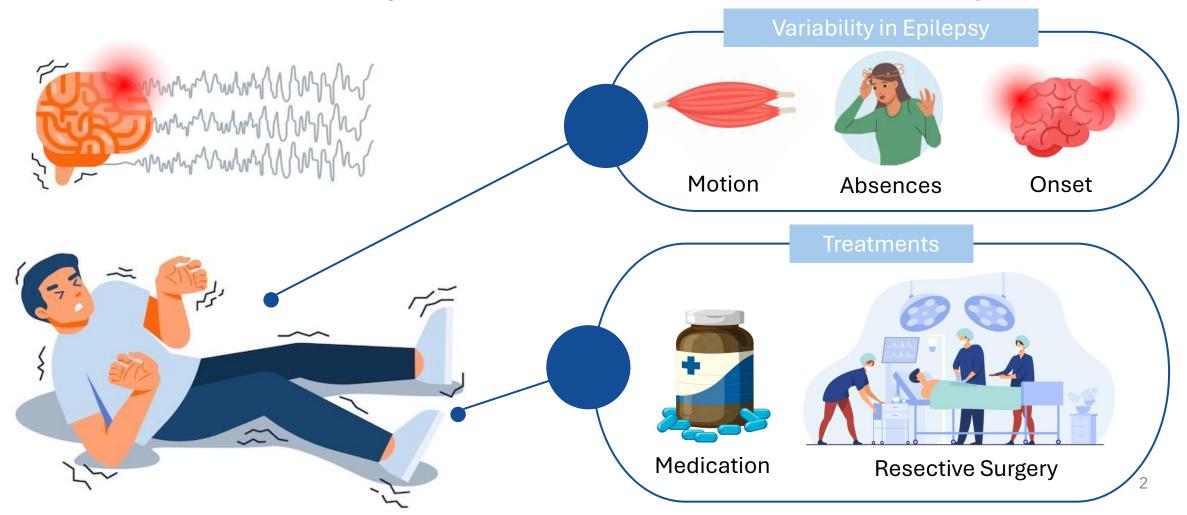
Optimiser et prédire l'efficacité de la stimulation du nerf vague pour combattre l'épilepsie

Marie Dawant



What is epilepsy?

- Epilepsy is a brain disorder that causes recurring, unprovoked seizures
- Seizures are sudden surges of abnormal and excessive electrical activity in the brain



Why do we need response prediction in epilepsy?





1/3 do not respond to medication



Not everyone is eligible for surgery

Vagus Nerve Stimulation



Induces seizure frequency reduction



Only 1/3 respond to therapy

Prediction



No information about the dose

Optimization



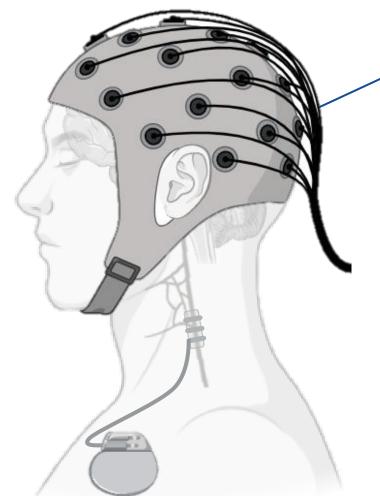
How to investigate VNS impact?







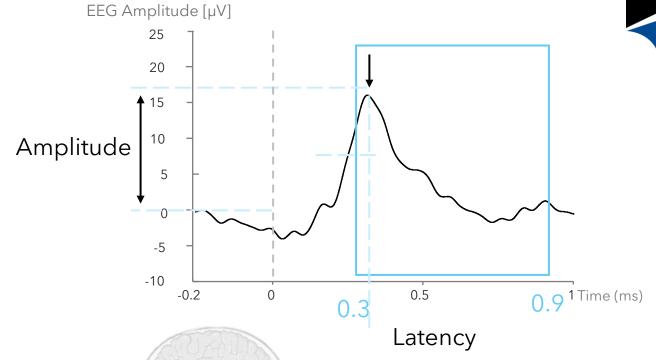
Positive component peaking at 300 ms after the stimuli **P300**

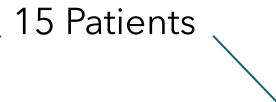


What can we already observe?

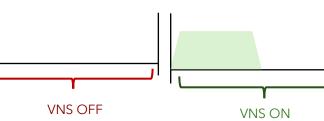
P300 Mean Waveform









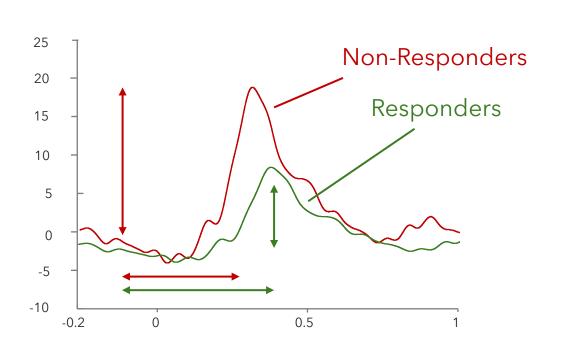


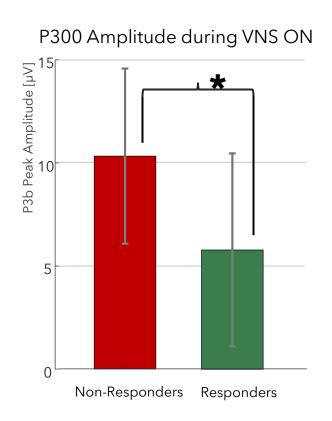
7 Responders

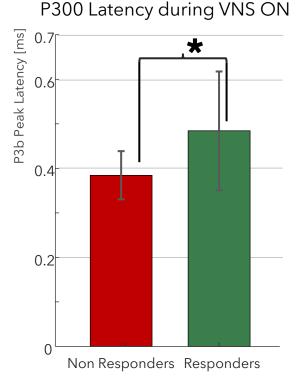
8 Non-Responders

What can we already observe?

- Interaction VNS with Response
- Opposite effects between Responders and Non-Responders in VNS ON







Thank you for your attention

Do you have questions?



What are the mechanisms underlying VNS?

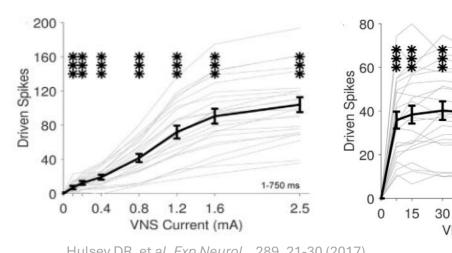
VNS output

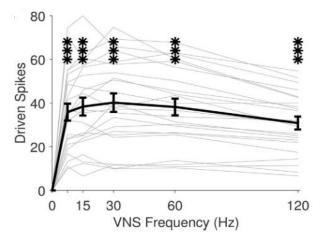
current



Locus Coeruleus

Main source of the Noradrenaline (NE) in the brain

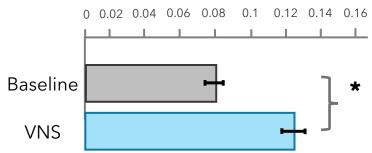






Hulsey DR. et al. Exp Neurol. 289, 21-30 (2017)

NE Concentration (nM)



Higher extracellular NE concentration in hippocampus after VNS

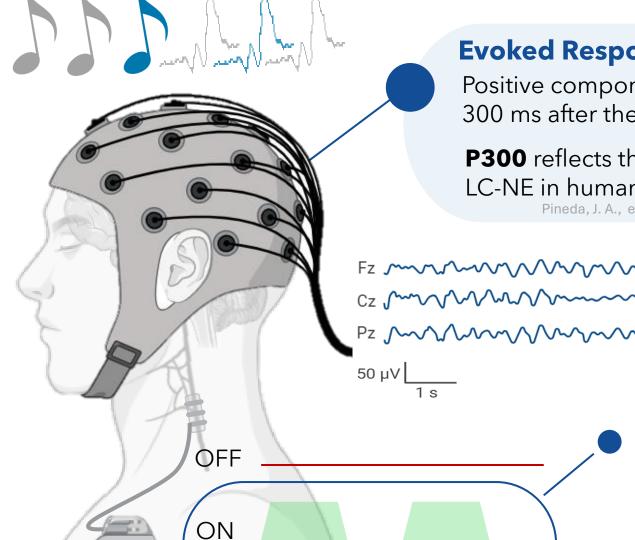
Limitation of seizure development only for an increase higher than 70 %

How to measure LC-NE activity?



P300 Amplitude in VNS Responders





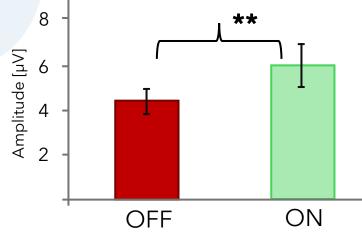
ON part of duty cycle

Evoked Response Potential

Positive component peaking at 300 ms after the stimuli **P300**

P300 reflects the phasic activity of LC-NE in humans

Pineda, J. A., et al. J. Neurosci. 9, 81-93 (1989)

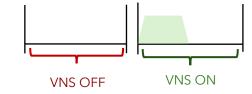


De Taeye, L. et al. Neurotherapeutics 11, 612-622 (2014)

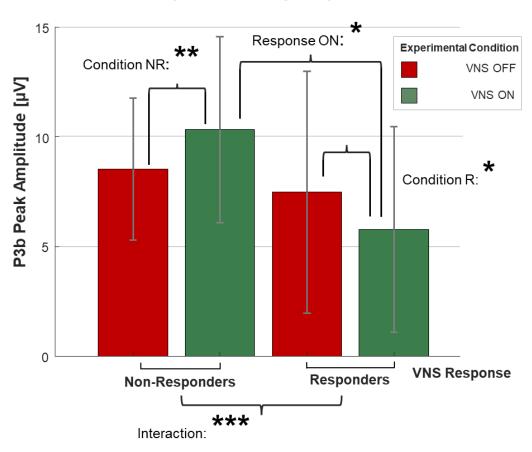
Can we measure the LC-NE activity underlying VNS during the stimulation pulses in humans?

Proof-of-Concept of a reliable Goal online VNS detection for P300

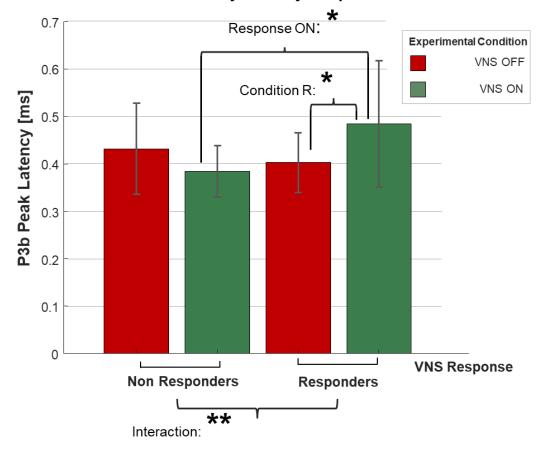
Summary of the electrophysiological results



Bar Plot of P3b Amplitude at Pz by Response and Condition



Bar Plot of P3b Latency at Pz by Response and Condition



Summary of the electrophysiological results viscoff VNS HIGH ON **VNS LOW ON**

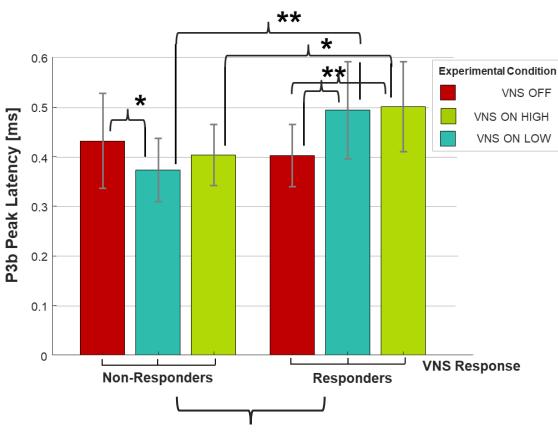
Bar Plot of P3b Amplitude at Pz by Response and Condition

Experimental Condition 16 VNS OFF ** * P3b Peak Amplitude [µV] VNS ON LOW VNS ON HIGH ** **VNS Response** Non-Responders Responders

Interaction: Response x (OFF x ON LOW)

Interaction: Response x (OFF x ON HIGH) **

Bar Plot of P3b Latency at Pz by Response and Condition



Interaction: Response x (OFF x ON LOW)

Interaction: Response x (OFF x ON HIGH) **

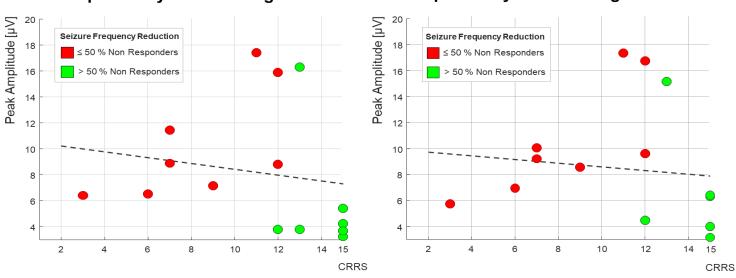
Summary of the electrophysiological results viscoff











P3b Amplitude by Score during VNS OFF

P3b Amplitude by Score during VNS ON LOW

