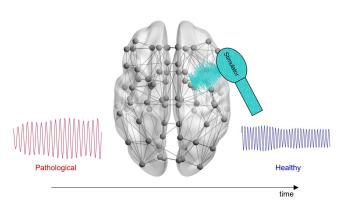


## New computational tool could help optimize treatment of Alzheimer's disease

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Reversion of pathological electroencephalographic activity in Alzheimer's disease with minimal energy deposition over the tissue can be achieved through delivering a computationally tuned brain stimulation that considers individual neuroimaging data. Credit: Lazaro Sanchez-Rodriguez, partially using BrainNet Viewer

Scientists have developed a novel computational approach that incorporates individual patients' brain activity to calculate optimal, personalized brain stimulation treatment for Alzheimer's disease. Lazaro Sanchez-Rodriguez of the University of Calgary, Canada, and colleagues present their new framework in *PLOS Computational Biology*.

Electrical stimulation of certain parts of the <a href="brain">brain</a>
could help promote healthy activity in neural
circuits impaired by Alzheimer's disease, a
neurodegenerative condition. This experimental
treatment has shown some promise in <a href="clinical trials">clinical trials</a>
. However, all patients currently receive identical
treatment protocols, potentially leading to different
outcomes according to individual variations in brain
signaling.

To investigate the possibility of personalized brain stimulation, Sanchez-Rodriguez and colleagues took a theoretical approach. They built a computational tool that incorporates patients' MRI scans and physiological brain signaling measurements to calculate optimal brain stimulation signals, with the goal of delivering efficient, effective personalized treatment.

The new approach is based on a computational strategy known as the state-dependent Riccati equation control (SDRE), which has been applied in other fields—such as aerospace engineering—to optimize input signals that control dynamic, nonlinear systems like the human brain. This strategy enabled the new tool to identify specific brain regions that would not benefit from brain stimulation.

The researchers also used their new framework to show that certain parts of the brain, the limbic system and basal ganglia structures, could serve as particularly powerful targets for brain stimulation in Alzheimer's disease. Moreover, they found that patients whose neural structures are highly integrated in the brain network may be the most suitable candidates for stimulation.

"With our new framework, we are getting closer to erasing the knowledge gap between theory and application in <u>brain stimulation</u>," Sanchez-Rodriguez says. "I think we will soon see a boom in the application of our framework and similar tools to study other diseases involving impaired <u>brain activity</u>, such as epilepsy and Parkinson's."

Next, the researchers plan to refine their tool so that it accounts for additional variation in brain activity between patients. The approach will need to be tested in animals before it enters clinical trials.

More information: Sanchez-Rodriguez LM, Iturria-Medina Y, Baines EA, Mallo SC, Dousty M, Sotero RC, et al. (2018) Design of optimal nonlinear network controllers for Alzheimer's disease. *PLoS Comput Biol* 14(5): e1006136. doi.org/10.1371/journal.pcbi.1006136



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