

Virtual Brain Twins in EBRAINS

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EBRAINS

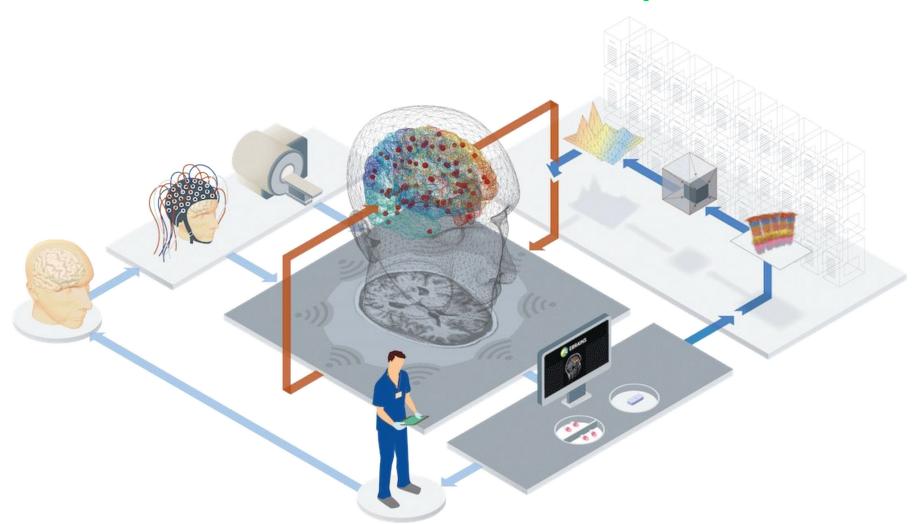
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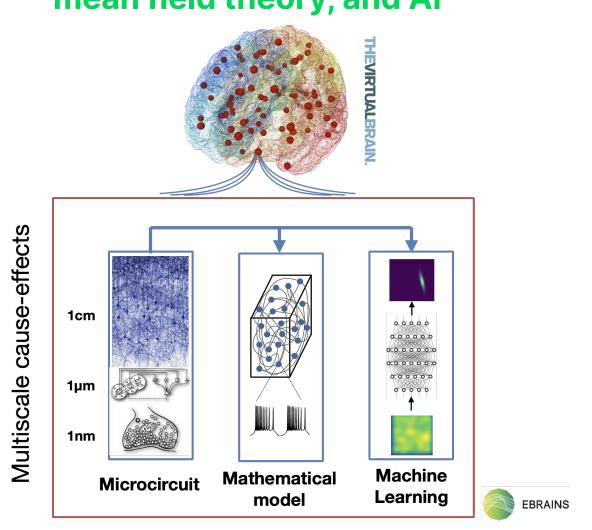
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Virtual Brain Twin workflow: from computer to bedside



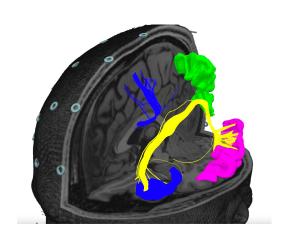
Patient-specific recordings constrain the personalized virtual brain twin. We use machine learning methods to further parametrize the patient's twin for specific conditions, with the goal of aiding clinical diagnosis and predicting best interventions, such as medication, surgery, and stimulation. In particular, one workflow (right) is the computationally demanding multiscale cause-effect simulations with the second workflow integrating patient specific data into a iteratively improved personalized virtual twin towards a system that can support clinicians with guidance in decision making for individual patients on which medication to take, how to adjust the dosage, when to change medications, when to suggest other factors in life such as physical activity.

Multiscale estimation of cause-effect using high-performance computation, mean field theory, and Al



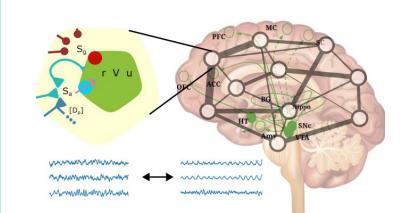
Three ways to Identify how mechanisms traverse across scales and express themselves mathematically in The Virtual Brain (TVB)

Personalized Virtual Brain Twin



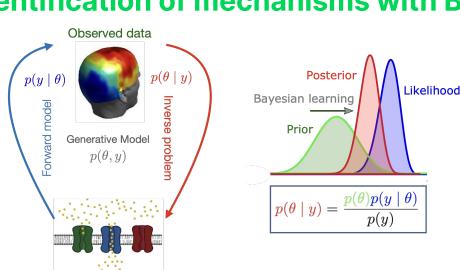
The virtual brain twin (VBT) is a personalization a general model produced on HPC using a subject's anatomical data T1including diffusion weighted Magnetic Resonance Imaging (MRI).

Neuromodulators drive the brain dynamics & entry points for pharmacological interventions



Besides the white matter connectivity each brain node is equipped with a model of activity neural that the effect captures neuromodulators (such dopamine).

Identification of mechanisms with Bayesian statistics



Hidden causes

Identification of the most likely model parameters (linked multiscale mechanisms) for the biomarkers

Virtual brain twin vs real surgery

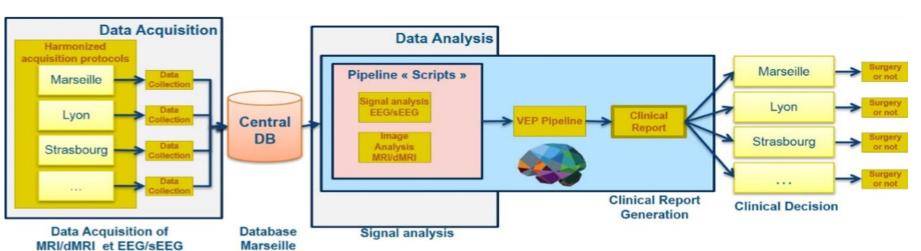
0.3 0.4 0.5 0.6 0.7 0.8 0.9 1.0

Correspondance with virtual brain prediction

Not-seizure-free (n=13)

Seizure-free (n=12)

Virtual Brain Twins in clinical trial: improving surgical outcome for drug resistant epilepsy



EPINOV Clinical Trial (2019-2023): randomized parallel-group study trial Sponsor : Assistance publique - Hôpitaux de Marseille Coordinator: Fabrice Bartolomei

Objective: evaluate the role of personalized Virtual Epileptic Patient (VEP) brain models for surgery

13 French clinical centers

Scientific Director: Viktor. Jirsa

356 prospective patients

310 patients randomized (VEP and 154 control)

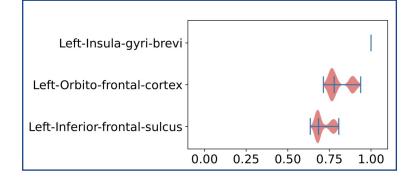
160 VEP reports sent (133 VEP and 27 control) 178 surgeries performed (93 VEP and 85 control) **X** epinov

Blue: Clinical Hypothesis for Epileptogenic Zone Red: Additional epileptogenic zone 29% of all cases fall in the red epileptogenic zone DASSAULT SYSTEMES areas and are not accessible via SEEG

SEEG time series during seizure

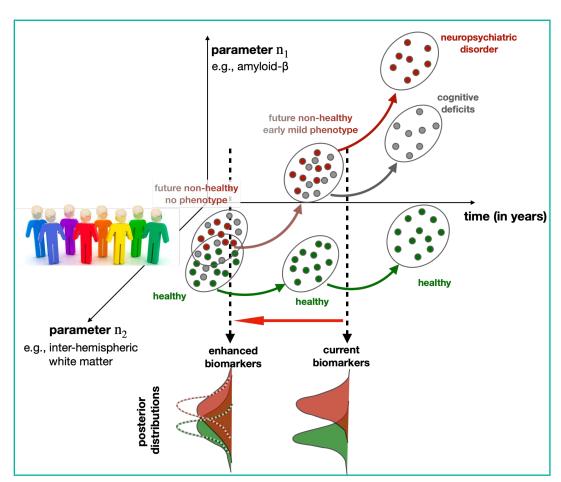
Time in seconds

Most surgical failures when TVB finds the clinical hypothesis to be incomplete



Estimated epileptogenic zone network with high excitability values

Virtual Brain Twins and inference predict individual aging, cognitive performance and brain health



accelerates The use of VBT diagnostics separating indistinguishable parameter distributions making pathophysiological causes identifiable.

Significant benefits of diagnostic acceleration by earlier access to pharmacological non-pharmacological treatments.



Healthy aging example Fitted fluidity Interhemispheric empirical simulated virtually aged connectivity decline (proxy for age fluidity related decline) $0.\overline{00}$ $0.\overline{25}$ $0.\overline{50}$ Interhemispheric α

Website

To learn more about the Virtual Brain Twin personalized treatment Psychiatric Disorders project, please visit website the virtualbraintwin.eu.



References

[1] Wang, H. E., Woodman, M., Triebkorn, P., Lemarechal, J.-D., Jha, J., Dollomaja, B., Vattikonda, A. N., Sip, V., Medina Villalon, S., Hashemi, M., Guye, M., Makhalova, J., Bartolomei, F., & Jirsa, V. (2023). Delineating epileptogenic networks using brain imaging data and personalized modeling in drug-resistant epilepsy. Science Translational Medicine, 15(680).

[2] Jirsa, V., Wang, H., Triebkorn, P., Hashemi, M., Jha, J., Gonzalez-Martinez, J., Guye, M., Makhalova, J., & Bartolomei, F. (2023). Personalised virtual brain models in epilepsy. The Lancet Neurology, 22(5), 443–454.

[5] Livingston G, Huntley J, Sommerlad A, et al. Dementia prevention, intervention, and care: 2020 report of the Lancet Commission Lancet. 2023 Sep 30;402(10408):1132.

