THE ATTRACTOR STATES OF THE FUNCTIONAL BRAIN CONNECTOME

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Abstract

Abstract: todo

Keywords

Key Points:

- We propose a high-level computational model of "activity flow" across brain regions
- The model considers the funcional brain connectome as an already-trained Hopfield neural network
- It defines an energy level for any arbitrary brain activation patterns
- and a trajectory towards one of the finite number of stable patterns (attractor states) that minimize
 this energy
- The model reproduces and explains the dynamic repertoir of the brain's spontanous activity at rest
- It conceptualizes both task-induced and pathological changes in brain activity as a shift on the "attractor landscape"
- We validate our findings on healthy and clinical samples (~2000 participants)

1 Introduction

Brain function is accompanied by the activation and deactivation of anatomically distributed neuronal populations. While changes in the activity of a single brain area is often associated with various tasks or conditions, in reality, regional activation never occurs in isolation (ref). Regardless of the presence or absence of explicit stimuli, brain regions seem to work in concert, resulting in a rich and complex spatiotemporal fluctuation over time. This fluctuation shows quasi-periodic properties (Thompson et al. [2014]), with a limited number of recurring states known as "brain states" (Gutierrez-Barragan et al. [2019], Vidaurre et al. [2017]). These states are often interpreted as sporadic intervals during which information can be efficiently exchanged between a characteristic subset of brain regions (Hutchison et al. [2013], Liu and Duyn [2013], Zalesky et al. [2014]). Brain state dynamics can be assessed with multiple techniques, including independent component analysis (ref), co-activation patterns (Liu and Duyn [2013]) and hidden markov models (Vidaurre et al. [2017]).

While such efforts, by their nature, do not shed light on the driving forces of the complex spatiotemporal dance of brain acctity, they provide accumulating evidence for the neurobiological relevance of these dynamics, with promising perspectives for facilitating the clinical translation of functional neuroimaging techniques (Lee et al. [2021]).

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Why does such interregional communication manifest in co-activation? Which activity configurations does the brain visit and which not? How do these relate to each other? How does this dyanmic repertoir of activation patterns result in task-related activity maps, as obtained with functional magnetic resonance imaging? What is the meaning of activity and connectivity differences across individuals or in various clinical conditions?

Traditional tools of computational neuroscience try to address these and similar questions whith methods of varying complexity from those based on the Fokker–Planck equation, to neural mass and neural field models. Commons amongst these models is the assumption that spatiotemporal patterns of neural dynamics arise from interactions between functionally specialized cell populations connected by a topologically complex array of short- and long-range axonal connections (Bullmore and Sporns [2009], Yuste [2015]), the latter ofen being estimated at macroscopic scales by diffusion magnetic resonance imaging (dMRI).

These models have found broad success in modeling seizures11, encephalopathies12,13, sleep14, anesthesia15, resting-state brain networks16,17 and the human alpha rhythm18,19, and as a tool for multimodal data fusion20. Technical advances in model inversion (estimating the likelihood and parameters of a model from empirical data) place mean field models within reach of widespread application to cognitive neuroscience21.

Nevertheless, there is no broadly accepted mathematical theory for the collective activity of neuronal populations and such models have shown limited success to bridge levels of explanations from singel neurons to complex behavior, mainly due to the grand challenges of estimating all free parameters.

Todo: shorten and focus on the fact that most comp models aim to solve the task all the way: to construct a "biophysical model" that accounts for empirical brain data and behavior.

Thus, the penetration of dynamic models of large-scale brain activity into mainstream neuroscience has been slow, and they may be unknown to many neuroscientists.

An alternative approach: Neuroconnectomist approach (Doerig et al. [2023])

A further scenario rests on the role of ghost attractors [109](https://www.nature.com/articles/nn.4497#ref-CR109 "Deco, G. & Jirsa, V.K. Ongoing cortical activity at rest: criticality, multistability, and ghost attractors. J. Neurosci. 32, 3366–3375 (2012).", a dynamic landscape of remnant attractors each of which has an incomplete basin, hence allowing the system to 'wander' through large swathes of the phase space under the influence of weak noise 110.

In this work, we aim for:

- simpliest, highest-level genrative computational model: a multistable dynamic system with **maximal empirical validity**
- bypass the challenges of estimating parameters, by building on the activity. flow
- no mechanistic model, not aiming to explain biophisical background
- links to Neuroconnectomism

multistable dynamical systems theory

Our framework - with minimal and reasonable assumptions about the "activity flow" (ref: Cole-papers) between two, functionally connected regions - considers the stationary functional brain network as an already-trained artificial neural network. In the proposed framework, the topology of the stationary brain connectome defines a cost (energy) for any arbitrary brain activation patterns and a trajectory towards one of the finite number of stable patterns that minimize this cost (so-called attractor states).

• noise

Here we propose these attractors states as robust and neurobiologically relevant characteristics of the functional brain connectome, with a wide variety of potential applications.

We demonstrate that the proposed attractor states highly resemble to the dynamic brain states commonly observed by dynamic functional connectivity methods (e.g. CAP-analyses (ref)) and provide a proof-of-concept for the biomedical validity of our framework, by showing that the average brain activations corresponding to the attractor states during resting sate display manifold significant associations to cognition.

Due to the known noise-tolerance of the applied eANN-s, the proposed approach can be expected to be highly robust/reliable/replicable, which we demonstrate with independent datasets (total n=xxx).

List all the aims: hierarchy, generalizability etc

2 Results

- introduce the idea in more detail (fig 1)
 - attractor states and added noise
 - construcct validity
- attractor states (fig 1)
- face validity
- clinical validity

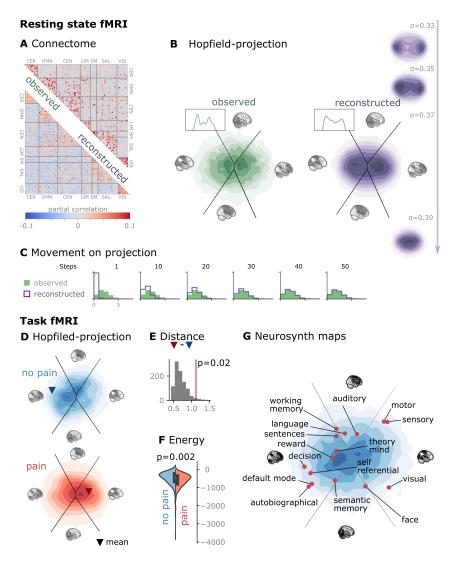


Figure 1: Empirical Hopfield-networks reconstruct real brain activity.

Here I refer to Figure 1.

3 Discussion

significance

4 Methods

Todo

Todo

References

- Garth John Thompson, Wen-Ju Pan, Matthew Evan Magnuson, Dieter Jaeger, and Shella Dawn Keilholz. Quasi-periodic patterns (QPP): Large-scale dynamics in resting state fMRI that correlate with local infraslow electrical activity. NeuroImage, 84:1018–1031, jan 2014. doi:10.1016/j.neuroimage.2013.09.029. URL https://doi.org/10.1016%2Fj.neuroimage.2013.09.029.
- Daniel Gutierrez-Barragan, M. Albert Basson, Stefano Panzeri, and Alessandro Gozzi. Infraslow state fluctuations govern spontaneous fMRI network dynamics. *Current Biology*, 29(14):2295–2306.e5, jul 2019. doi:10.1016/j.cub.2019.06.017. URL https://doi.org/10.1016%2Fj.cub.2019.06.017.
- Diego Vidaurre, Stephen M. Smith, and Mark W. Woolrich. Brain network dynamics are hierarchically organized in time. *Proceedings of the National Academy of Sciences*, 114(48):12827–12832, oct 2017. doi:10.1073/pnas.1705120114. URL https://doi.org/10.1073%2Fpnas.1705120114.
- R. Matthew Hutchison, Thilo Womelsdorf, Elena A. Allen, Peter A. Bandettini, Vince D. Calhoun, Maurizio Corbetta, Stefania Della Penna, Jeff H. Duyn, Gary H. Glover, Javier Gonzalez-Castillo, Daniel A. Handwerker, Shella Keilholz, Vesa Kiviniemi, David A. Leopold, Francesco de Pasquale, Olaf Sporns, Martin Walter, and Catie Chang. Dynamic functional connectivity: Promise, issues, and interpretations. *NeuroImage*, 80:360–378, oct 2013. doi:10.1016/j.neuroimage.2013.05.079. URL https://doi.org/10.1016%2Fj.neuroimage.2013.05.079.
- Xiao Liu and Jeff H. Duyn. Time-varying functional network information extracted from brief instances of spontaneous brain activity. *Proceedings of the National Academy of Sciences*, 110(11):4392–4397, feb 2013. doi:10.1073/pnas.1216856110. URL https://doi.org/10.1073%2Fpnas.1216856110.
- Andrew Zalesky, Alex Fornito, Luca Cocchi, Leonardo L. Gollo, and Michael Breakspear. Time-resolved resting-state brain networks. *Proceedings of the National Academy of Sciences*, 111(28):10341–10346, jun 2014. doi:10.1073/pnas.1400181111. URL https://doi.org/10.1073%2Fpnas.1400181111.
- Jae-Joong Lee, Hong Ji Kim, Marta Čeko, Bo yong Park, Soo Ahn Lee, Hyunjin Park, Mathieu Roy, Seong-Gi Kim, Tor D. Wager, and Choong-Wan Woo. A neuroimaging biomarker for sustained experimental and clinical pain. *Nature Medicine*, 27(1):174–182, jan 2021. doi:10.1038/s41591-020-1142-7. URL https://doi.org/10.1038%2Fs41591-020-1142-7.
- Ed Bullmore and Olaf Sporns. Complex brain networks: graph theoretical analysis of structural and functional systems. *Nature Reviews Neuroscience*, 10(3):186–198, feb 2009. doi:10.1038/nrn2575. URL https://doi.org/10.1038%2Fnrn2575.
- Rafael Yuste. From the neuron doctrine to neural networks. *Nature Reviews Neuroscience*, 16(8):487–497, jul 2015. doi:10.1038/nrn3962. URL https://doi.org/10.1038%2Fnrn3962.
- Adrien Doerig, Rowan P. Sommers, Katja Seeliger, Blake Richards, Jenann Ismael, Grace W. Lindsay, Konrad P. Kording, Talia Konkle, Marcel A. J. van Gerven, Nikolaus Kriegeskorte, and Tim C. Kietzmann. The neuroconnectionist research programme. *Nature Reviews Neuroscience*, 24(7):431–450, may 2023. doi:10.1038/s41583-023-00705-w. URL https://doi.org/10.1038%2Fs41583-023-00705-w.