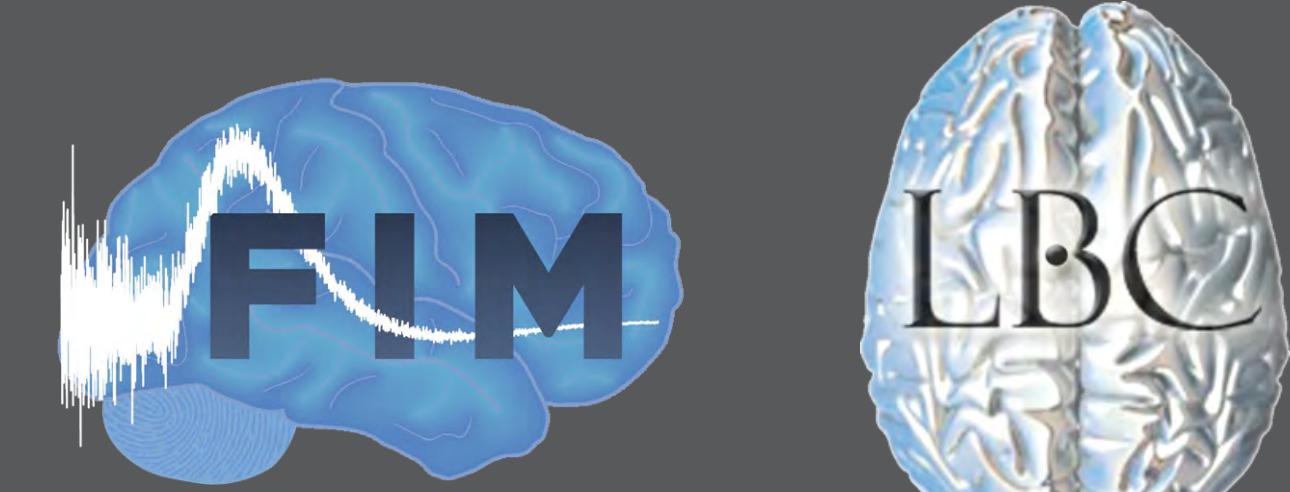


# On the static and dynamic features of edge time series

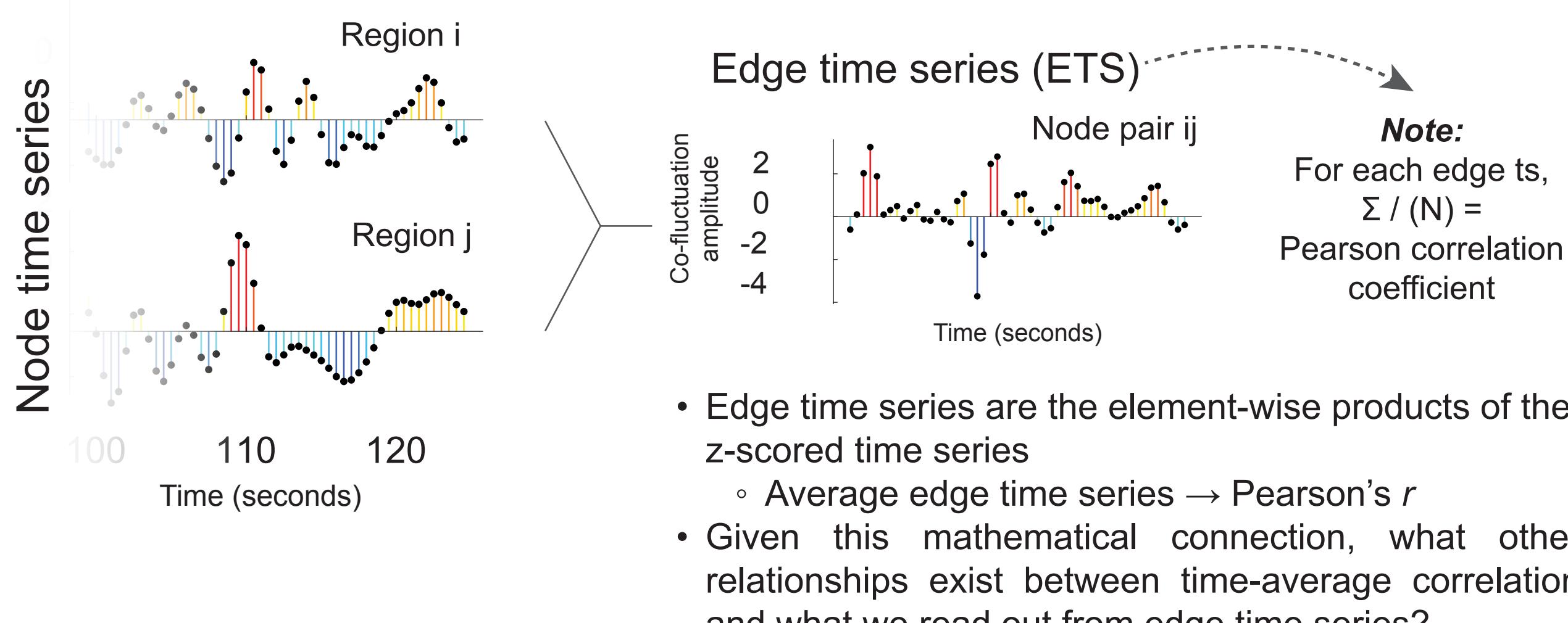
Joshua Faskowitz<sup>1</sup>, Tyler Morgan<sup>1</sup>, Daniel A. Handwerker<sup>1</sup>,  
Javier Gonzalez-Castillo<sup>1</sup>, Peter A. Bandettini<sup>1,2</sup>

<sup>1</sup> Section on Functional Imaging Methods,  
<sup>2</sup> Functional Magnetic Resonance Imaging Core Facility,  
National Institute of Mental Health, Bethesda, USA



## Introduction

- Functional magnetic resonance imaging (fMRI) has repeatedly shown that BOLD signals will fluctuate across the cortex during task-free conditions
  - Fluctuating regions are often compared using **Pearson correlation**
  - Taking correlation between all region pairs forms a correlation, i.e., *functional connectivity*, matrix.
- Measuring the **dynamic nature of correlations** is increasingly popular; can reveal connectivity states<sup>5</sup> and transients<sup>11</sup>
- Edge time series**<sup>3,16</sup> render connectivity dynamics at the temporal resolution of the input time series
  - Recently, it has been shown that features of edge time series can be partially explained with features of static correlation<sup>9,12,13</sup>.
- Here we further explore the relationship between static versus dynamic edge time series features.

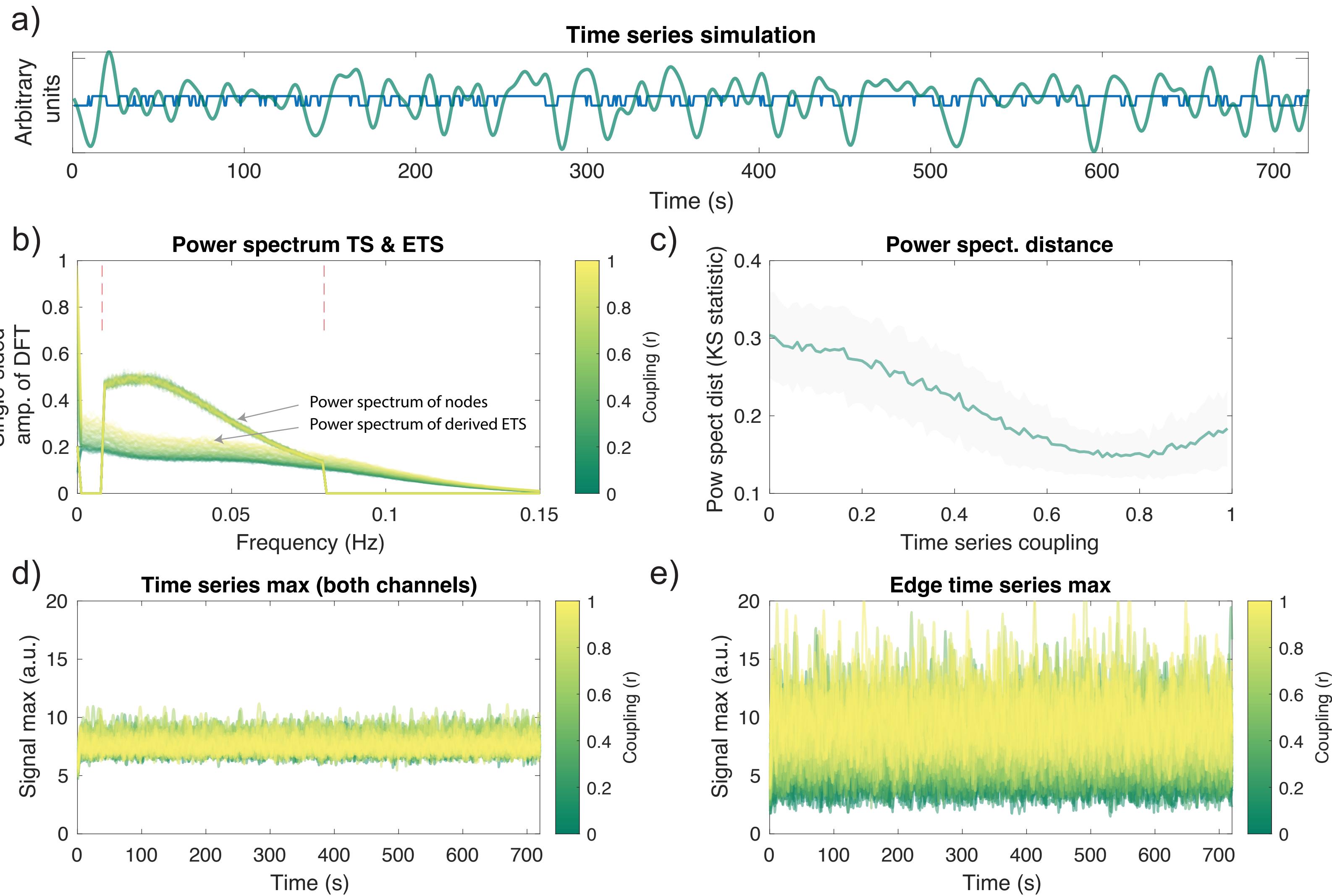


- Edge time series are the element-wise products of the z-scored time series
  - Average edge time series → Pearson's  $r$
- Given this mathematical connection, what other relationships exist between time-average correlation and what we read out from edge time series?

## Methods

- Simulated data**
  - Time series generated with a 2-state Markov chain, convolved with a canonical HRF for five-second stimulus and bandpass filtered (0.008–0.08 Hz)
  - Target covariance patterns were enforced on randomly generated and uncorrelated channels by projecting the data onto the axes of the eigen-decomposed covariance<sup>10</sup>
    - Using this method, we can control coupling (i.e., correlation) between two channels
  - Randomly generated target covariance using the stochastic block model<sup>1</sup> — a generative network model that allows for planted community structure with parameterized weight and edge existence
- Real data**
  - Human Connectome Project resting-state data (0.72 TR, 1200 TRs, ~15 mins)
  - Minimally pre-processed<sup>4</sup>: motion, distortion, bandpass, first/last 50 TRs discarded
  - Nuisance regressed using aCompCor<sup>2</sup> components (5 WM, 5 CSF) and 24 motion parameters. Time series constructed by averaging vertex data within 200 node of Schaefer parcellation<sup>15</sup>, at each time point, using Connectome Workbench
- Edge time series**
  - Computed via the element-wise product of two z-scored time series; sum of which is correlation
  - RSS: root of the sum of squares of all edge time series magnitudes, taken at each time point

## Results

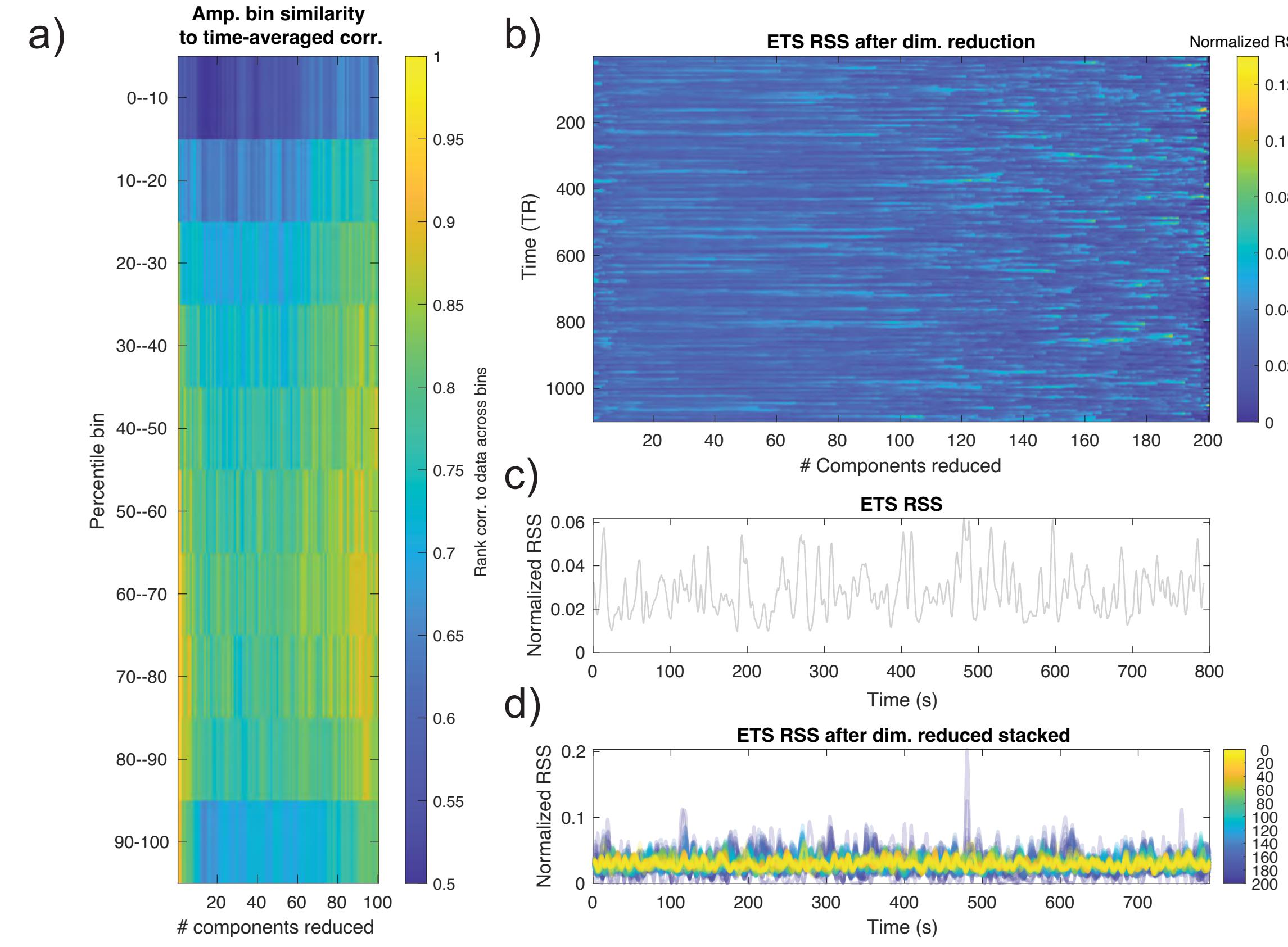
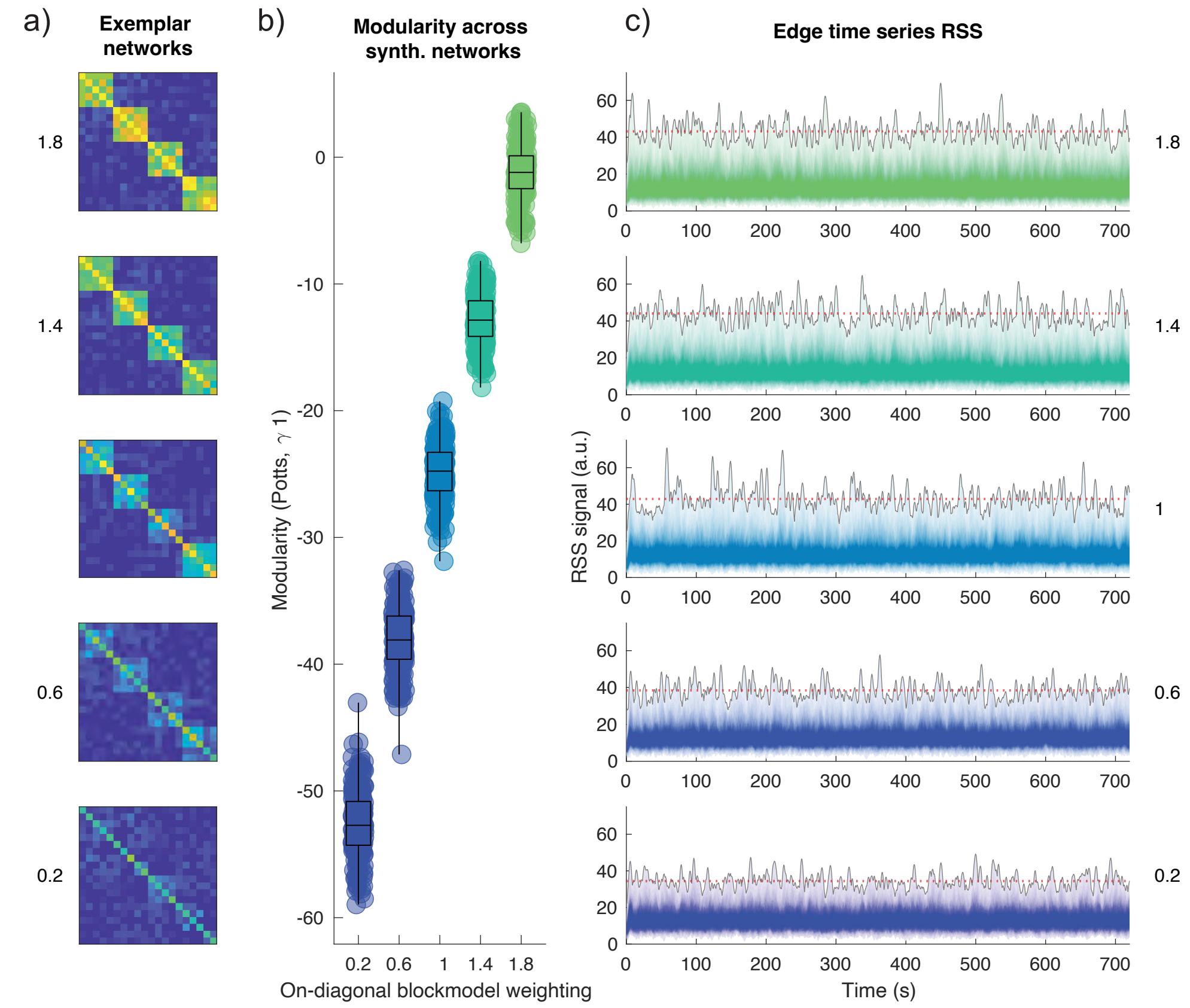


**Figure 1. Using simulated data with parameterized coupling, we observe relationship between the similarity of node time series and the amplitude of edge time series.** a) Example time series simulation for one channel using two-state Markov chain (blue) that is subsequently convolved with the HRF and bandpass filtered (green); 1000 time points generated b) Individual channels ( $n=200$ ) adhere to bandpass filter (vertical red lines), whereas the ETS generation induces higher/lower amplitudes c) The distance (Kolmogorov-Smirnov) between node and ETS power spectra is systematically modulated by coupling of two channels, with min. around 0.8 d) Maximum values of time series within a narrow range compared to (e), the maximum values of ETS; coupling values derived from 200 simulations at each of 100 coupling values (0–99, 0.01 steps), denoted by green-yellow color map

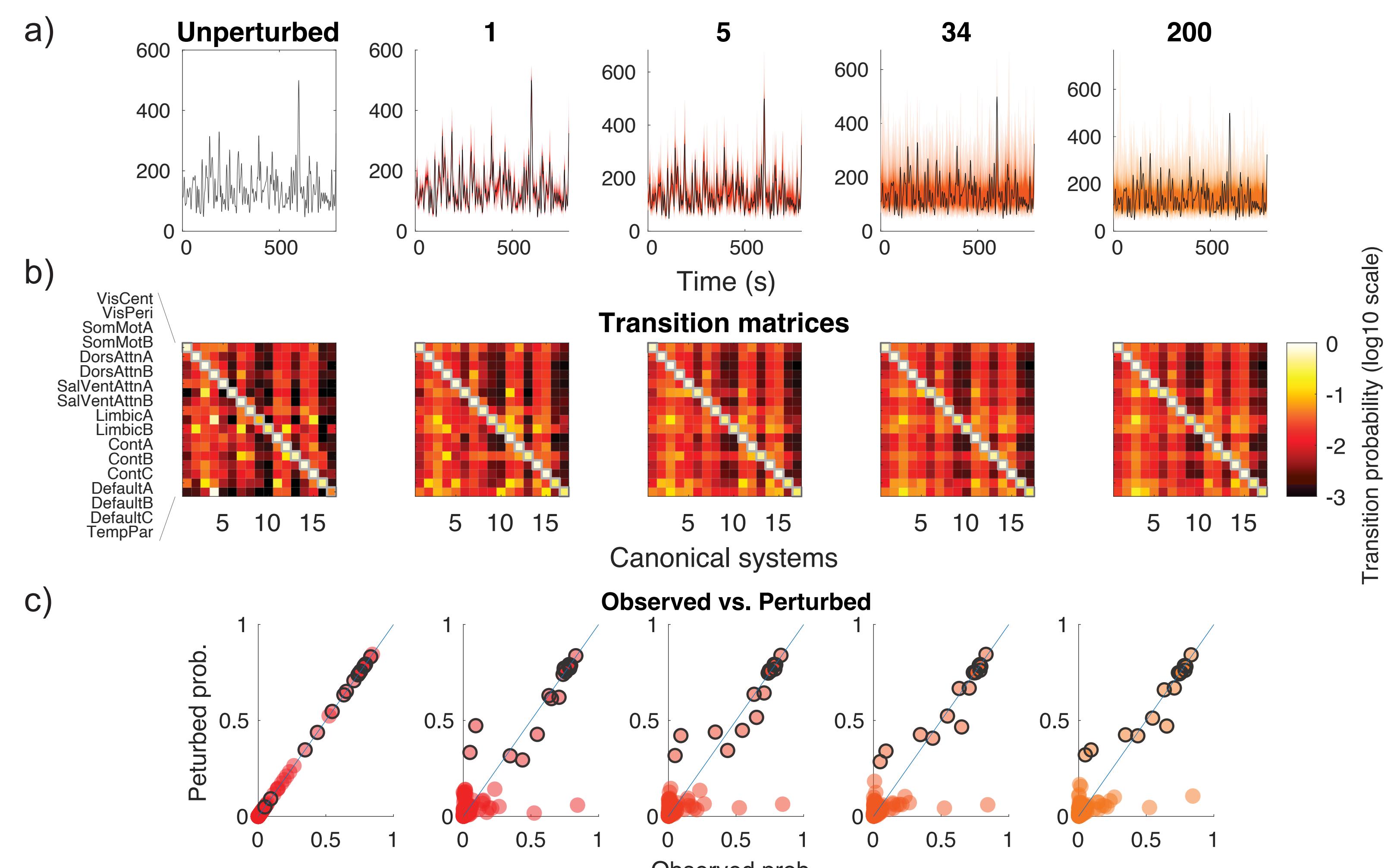
## References

- Aicher, C., Jacobs, A. Z., & Clauset, A. (2015). Learning latent block structure in weighted networks. *Journal of Complex Networks*
- Behzadi, Y. (2017). A component-based noise correction method (CompCor) and perfusion based fMRI. *Neuroimage*
- Faskowitz, J. (2020). Edge-centric functional network representations of human cerebral cortex reveal overlapping system-level architecture. *Nat. Neuro.*
- Glasser, M. F. (2015). The minimal preprocessing pipeline for the Human Connectome Project. *Neuroimage*
- Gonzalez-Castillo, J. (2015). Tracking ongoing cognition in individuals using brief, whole-brain functional connectivity patterns. *PNAS*
- Gonzalez-Castillo, J. (2021). Human-scale network organization links to variation in hormone concentrations over the menstrual cycle. *Net. Neuro.*
- Gutierrez-Barraquer, D. (2022). Unique spatiotemporal fMRI dynamics in the awake mouse brain. *Current biology*
- Han, S. (2012). Global changes in fMRI connectivity. *Neuroimage*
- Ladwig, Z. (2022). BOLD fluctuation ‘events’ are predicted from static functional connectivity. *Neuroimage*
- Laumann, T. O. (2017). On the stability of BOLD fMRI correlations. *Cerebral cortex*
- Liu, X. (2013). Time-varying functional network information extracted from brief instances of spontaneous brain activity. *PNAS*
- Matsu, T. (2022). On co-activation pattern analysis and non-stationarity of resting brain activity. *Neuroimage*
- Novelli, L. (2022). A mathematical perspective on edge-centric brain functional connectivity. *Nature communications*
- Pope, M. (2021). Modular origins of high-amplitude cofluctuations in fine-scale functional connectivity dynamics. *PNAS*
- Schaefer, A. (2018). Local-global parcellation of the human cerebral cortex from intrinsic functional connectivity MRI. *Cerebral cortex*
- Zamani Esfahanian, F. (2020). High-amplitude cofluctuations in cortical activity drive functional connectivity. *PNAS*

**Figure 2. By generating a toy system of 20 node time series with a planted covariance structure, we observe that modularity affects the maximum edge time series amplitude.** a) Example matrices produced by the stochastic blockmodel, at different concentrations, which result in different modularity values b) based on Potts model with a gamma of 1. c) After generating 200 synthetic matrices, time series, and subsequently edge time series for each network, the room-sum-square (RSS) of the edge time series are taken; more extreme RSS produced, as indicated by the red line, by more modular structures, as predicted and expected based on previous studies<sup>13,14</sup> which discussed dependency of “events” on modularity or a relatively large leading eigenvector of the covariance matrix; max RSS: grey line, mean of max RSS: red dotted line



**Figure 3. After increasing amount of principle components removed from time series data, binning patterns and RSS ETS change.** a) RSS % bins were used demarcate data to construct 10 matrices & each matrix was compared to time-averaged connectivity, at each level of components removed b) ETS RSS as a matrix, showing how pattern of high amplitude events changes as function of variance removed c) Unperturbed ETS RSS d) Normalized ETS RSS at varying levels of perturbation, distinguished by color map; because generating ETS involves z-scoring, data with many components removed (>100) produce similar RSS magnitude as less perturbed, yet more noisy.



**Figure 4. Applying randomization that preserves static covariance can still change edge time series phenomena.** a) ETS RSS time series from an HCP resting-state scan, with varying levels of randomization applied (100 surrogate time series, as colored lines), according to the number of probabilistically sampled frequencies that were phase-shifted<sup>8</sup>; shift applied to all channels equally, maintaining original covariance pattern b) Transition probability matrices<sup>7</sup> derived from ETS and surrogate data; based on t and t+1 transitions for each node, where affiliation is dictated by max. ETS magnitude at each time point c) Original vs. surrogate data transition probability values compared, demonstrating that even a slight perturbation to data, with time-averaged covariance held constant, causing between-system changes; suggests target for future ETS applications, not tethered to time-averaged covariance.

## Discussion

- Here we probe time series and edge time series characteristics, to obtain a better understanding of how their features are related. We illustrate:
  - ETS do not respect bandpass filter of original data (Fig1b), and the similarity of time series modulates the magnitude of ETS (Fig1c-e); implicates role of correlation for high amplitude ETS
  - Higher modularity of a simulated system results in higher amplitude ETS RSS (Fig. 2)
  - Similarity of ETS RSS binned data to time-averaged data holds for intact data (Fig3), and relationship is diminished after removing variance via incomplete PCA reconstructions
  - Small perturbations to time series (Fig 4a) results in large dynamics changes (Fig4b,c), even if time-averaged covariance is kept constant by the randomization method
- Indeed, some features of the time series are sufficient to estimate ETS properties<sup>13</sup>—notably those ETS properties that collapse across time, such as edge community similarity<sup>3</sup> or the distribution of high-amplitude events into specific bins<sup>9</sup>.
  - Suggests that future work on edge time series should explore dynamics using the fine-grained precision that ETS can offer; opportunity for state-based analyses<sup>6,7</sup>
  - Additional line of research will focus on comprehensively characterizing ETS channel properties (power/frequency, duty cycle, burstiness) to potentially distinguish if correlation appears different, for different functional relationships; i.e., investigating different fluctuation regimes across brain.