

# Psilocybin Drives Idiosyncratic Brain Responses to Movies

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## Background and Questions

- Ingestion of psilocybin (“magic mushrooms”) causes changes to perception<sup>1</sup>, emotion<sup>2</sup>, memory<sup>3</sup>, reasoning<sup>4</sup>, and more<sup>5</sup>
- To what extent are such changes similar or idiosyncratic across subjects?
- Where in the visual processing pathway are perceptual changes associated with altered neural representations?
  - As early as V1, or only in higher-order regions?
- How does psilocybin alter encoding and memory of narrative information?
- How does psilocybin alter large-scale brain activity during naturalistic perception, and how does this differ from alterations during task and task-free conditions?

## Methods

- 13 participants (of a planned 30) received 10mg/70kg psilocybin or placebo on separate visits
- Beginning ~60 minutes post-dose, participants watched one hour of short audiovisual movies during fMRI
- Movies were segmented into events by human raters
- Intersubject brain similarity during movie watching was measured via:
  - Intersubject pattern correlation (pISC), i.e. ISC of parcel-wise multivoxel patterns averaged over movie events<sup>6</sup> (Fig. 1)
  - Temporal intersubject correlation (tISC), i.e. ISC of each voxel’s timeseries<sup>7</sup> (Fig. 2)
  - Inter-regional brain coupling was assessed via intersubject functional connectivity analysis (ISFC), i.e. tISC between brain regions<sup>8</sup> (Fig. 3)
- Coupling of visual brain activity to stimulus was assessed by fitting a model of low-level visual features to predict voxel timeseries

## Bonus Materials and Contact

Figure captions  
Movie stimuli



References

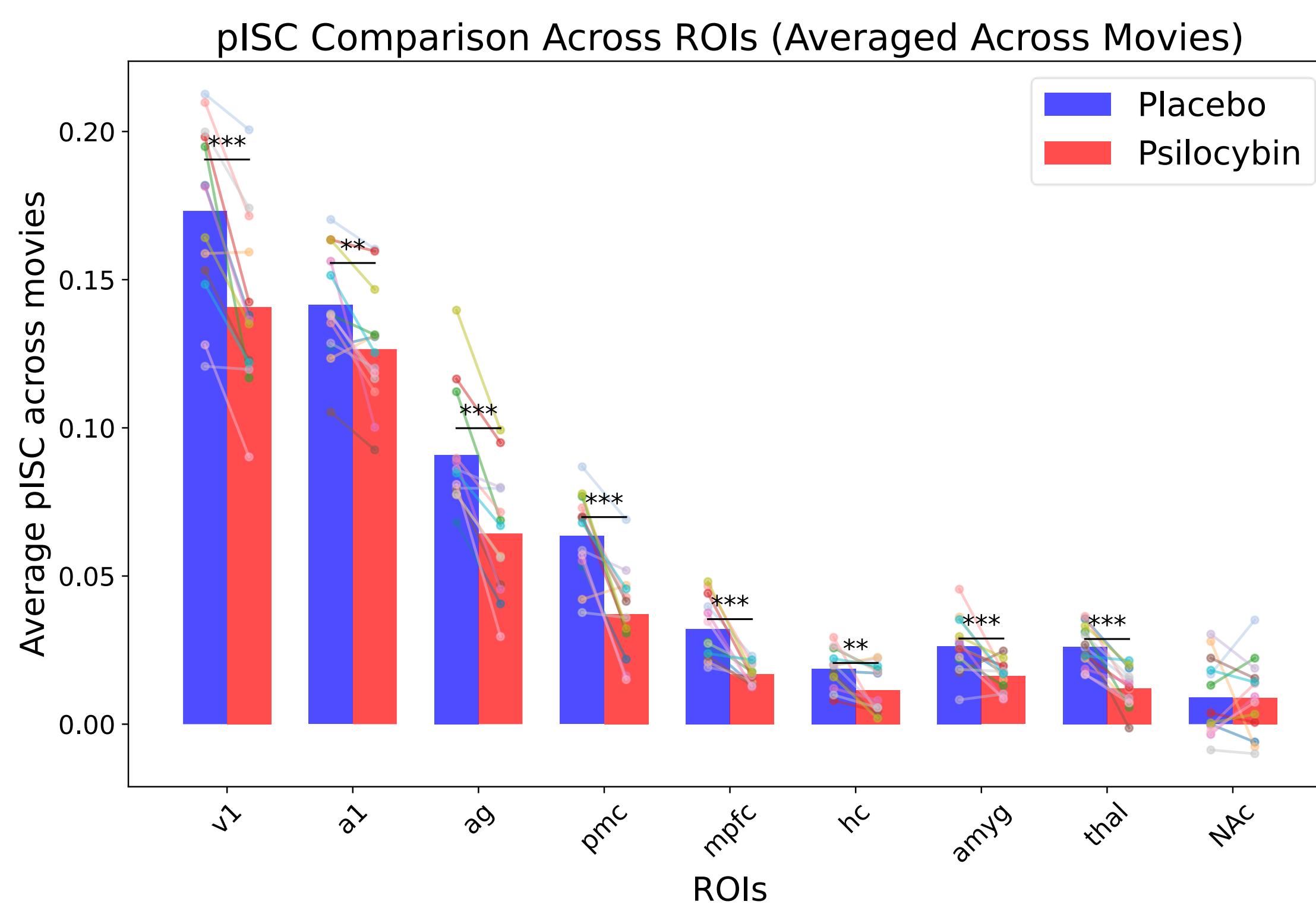
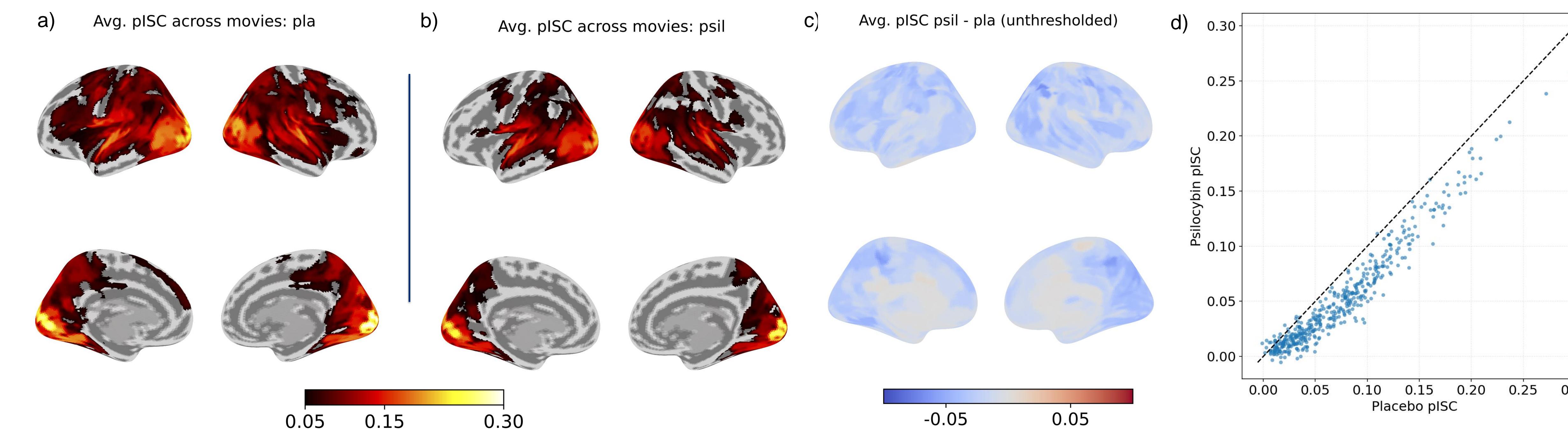


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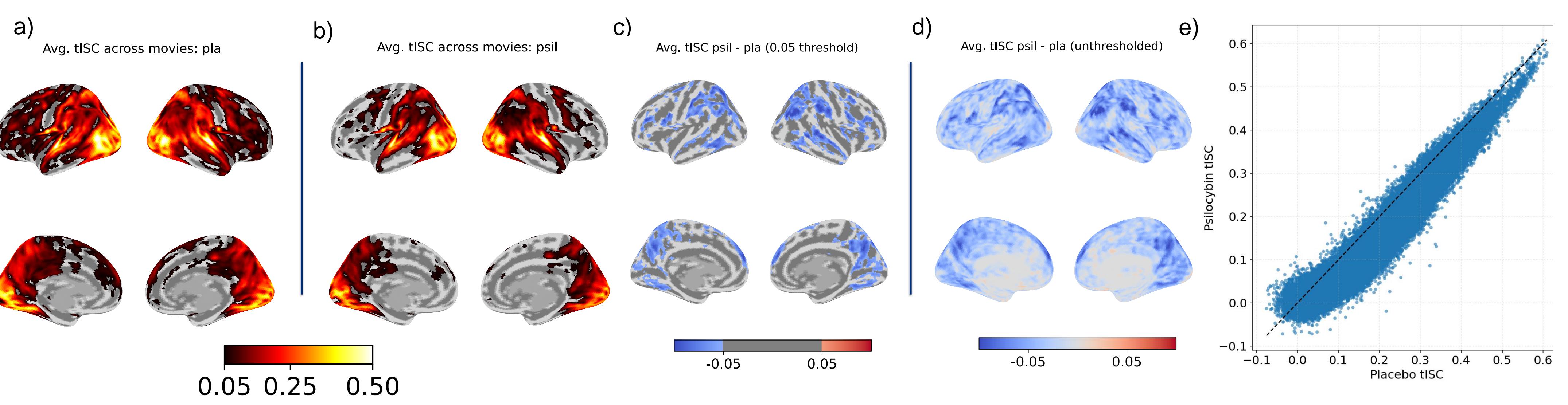


## Results

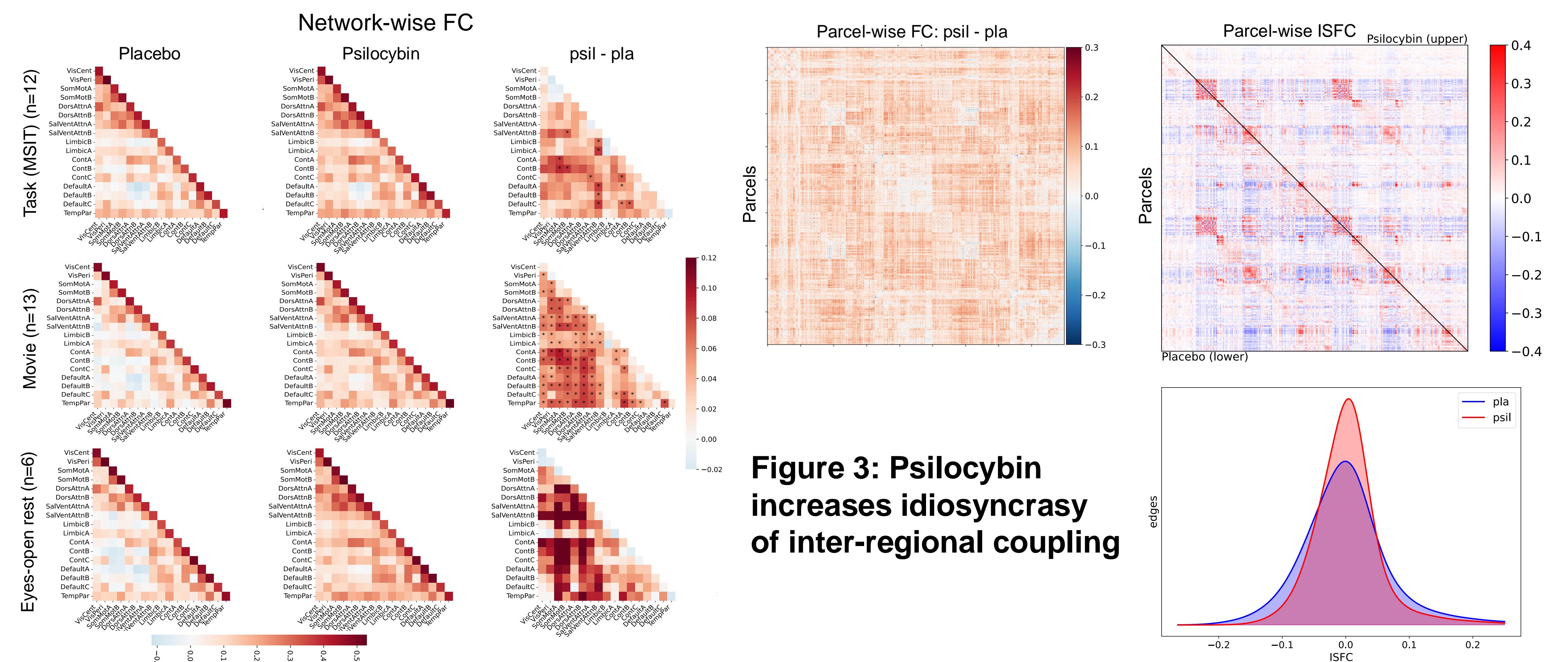
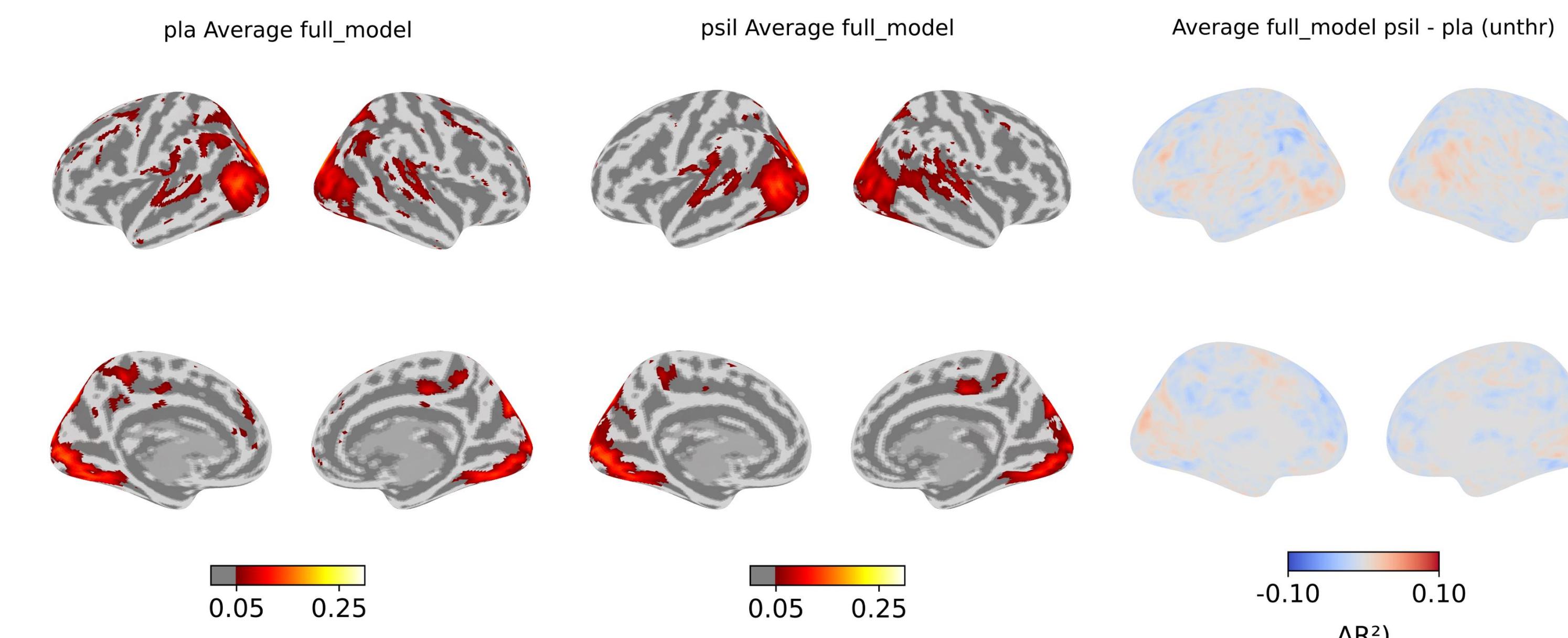
**Figure 1— Psilocybin reduces intersubject pattern correlations (pISC) across cortex**



**Figure 2: Psilocybin reduces temporal intersubject correlations (tISC) across cortex**



**Figure 4: Low-level visual stimulus features predict visual responses equally across drugs**



**Figure 3: Psilocybin increases idiosyncrasy of inter-regional coupling**

## Conclusions

- Psilocybin reduced temporal, spatial, and interregional ISC across the whole brain, likely reflecting idiosyncratic changes to perceptual and cognitive representations of audiovisual narratives
- Despite reduced ISC in V1, a model of low-level visual features predicted visual responses equally across drug conditions, raising questions about top-down influence on sensory processing in the production of illusions and hallucinations
- Psilocybin increased FC between most brain regions and networks but caused ISFC to move towards 0 (uncorrelated). This may suggest psilocybin causes spatially widespread effects that are consistent across subjects but unrelated to the stimulus