



Mutual Information measures as a connectivity indicator

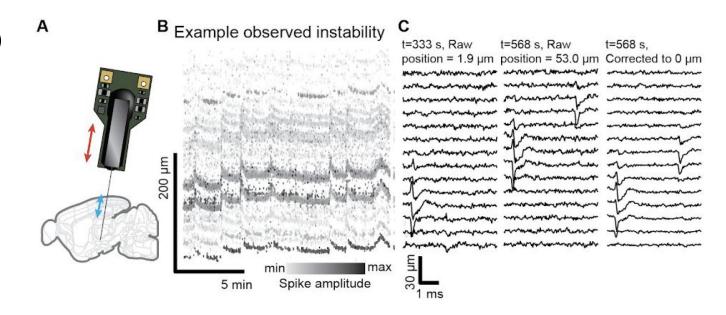
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Introduction. Dataset

- Which data?
 - Mice with 2 genotypes, wild type (eu) and trisomic (ts)
- How?
 - **Neuropixel** high density probes
- Where? Prefrontal cortex, with neurons from:
 - Supplementary Motor Cortex (M2)
 - Anterior Cingulate Cortex (AC)
 - PreLimbic Cortex (PrL)
 - InfraLimbic Cortex (IL)

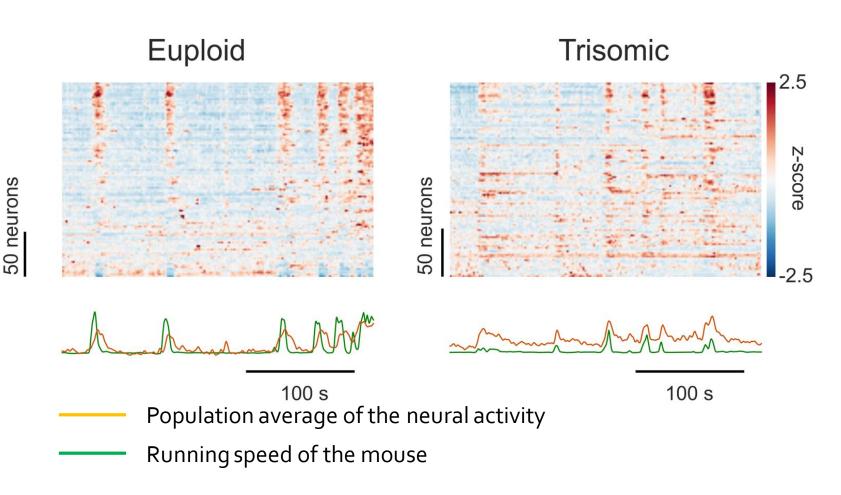


Extracted from Figure 3 of *Nicholas A. Steinmetz et al.*, Neuropixels 2.o: A miniaturized high-density probe for stable, long-term brain recordings. Science, 372, eabf4588(2021). DOI:10.1126/science.abf4588

Introduction. Problem

Control

From the Neuropixels,
we can extract the
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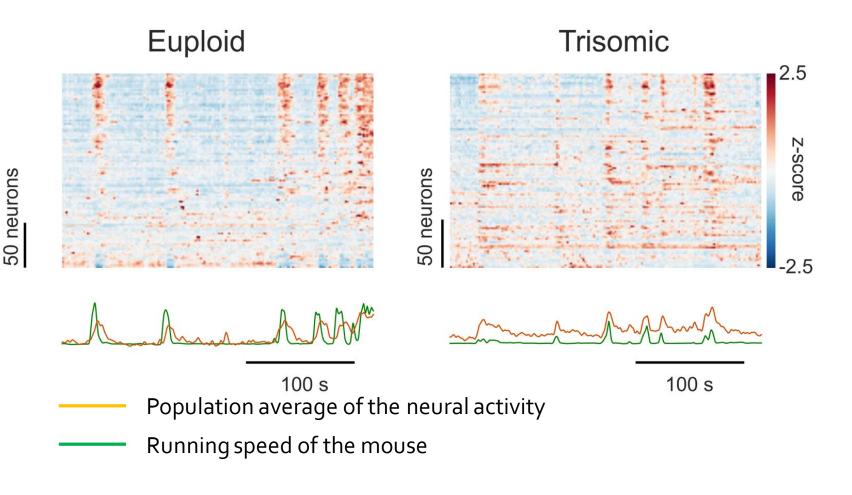


Introduction. Problem

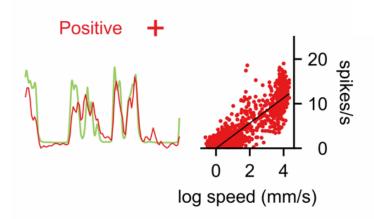
Control

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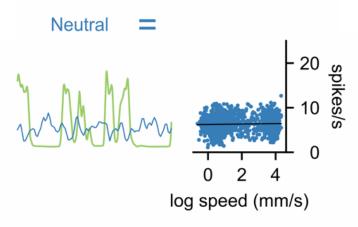
Why do the neural activity and locomotion decorrelate in the trisomic mice?

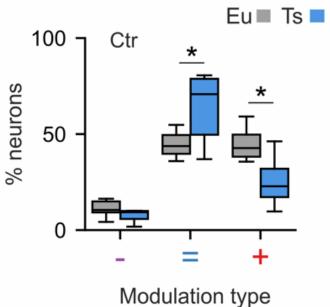


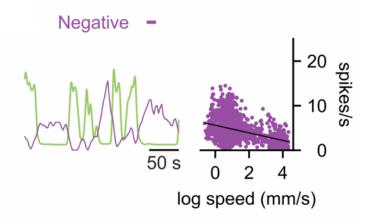
Preliminar results



Attending to this classification, we see the following...







... but can we learn something in this dataset from the network point of view?



Entropy & Mutual Information

Shannon Entropy

$$H(X) = -\sum p(x) \log_2 p(x) dx$$

$$p(x) \equiv \text{probability distribution of } yariable X$$

- Measures the uncertainty in the probability distribution, the more random (uniform) the higher the entropy
- If measured in bits, it also correspond to the number of yes/no questions we need to characterize the distribution

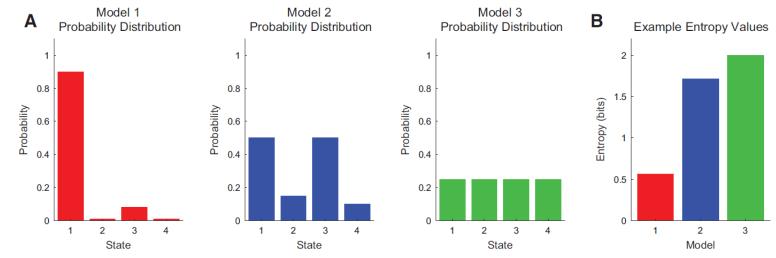


Figure 3. Example entropy calculations. **A**, Example probability distributions for three models (red, blue, and green); **B**, their associated entropy values. Model 1 was most likely to be in state 1, so it had low entropy. Model 3 was equally likely to be in all four states, so it had maximum entropy. Uniform count binning (see *Data Binning*) will produce equally likely states and maximize entropy, similar to Model 3.

Nicholas M. Timme, Christopher Lapish, *A Tutorial for Information Theory in Neuroscience*, eNeuro 29 June 2018, 5 (3) ENEURO.0052-18.2018; DOI: 10.1523/ENEURO.0052-18.2018

Entropy & Mutual Information

Shannon Entropy

$$H(X) = -\sum p(x)\log_2 p(x)dx$$

 $p(x) \equiv probability distribution of variable X$

Mutual Information

I(X,Y) = H(X) + H(Y) - H(X,Y) $H(X,Y) \equiv \text{ joint entropy (shared)}$

 MI tells us the reduction in uncertainty that we get in one variable by knowing the other

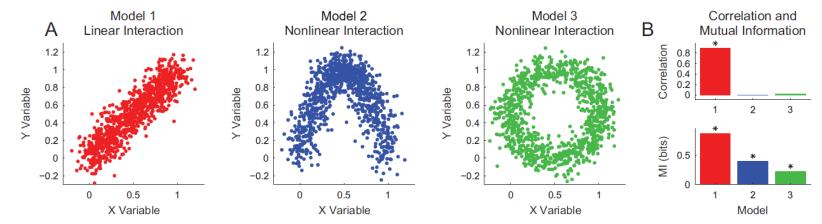
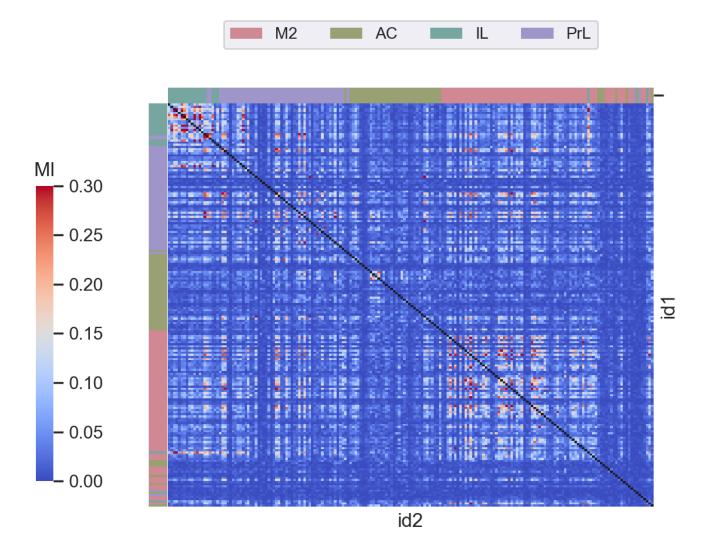


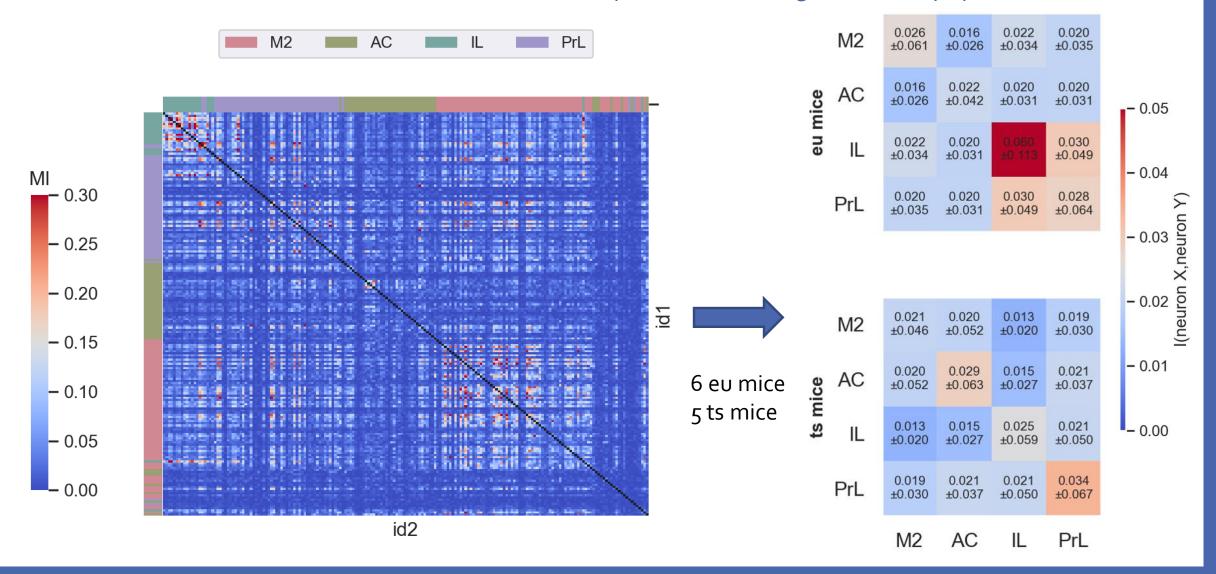
Figure 5. Example of linear versus nonlinear analysis methods. **A**, Example data for three models (red, blue, and green) with linear (red) and nonlinear (blue and green) interactions; **B**, the associated correlation coefficient and mutual information (MI) values for all three models (star: p < 0.05, correlation coefficient and p-value calculated via MATLAB, mutual information and p-value calculated via the Neuroscience Information Theory Toolbox; see *Data Binning and Significance Testing*, 4 bins and 1000 null data sets).

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Neuropixels \rightarrow Spike trains \rightarrow Continuous time series of spiking rates with time bin $\Delta t \rightarrow$ Mutual Information measure between neurons pairs

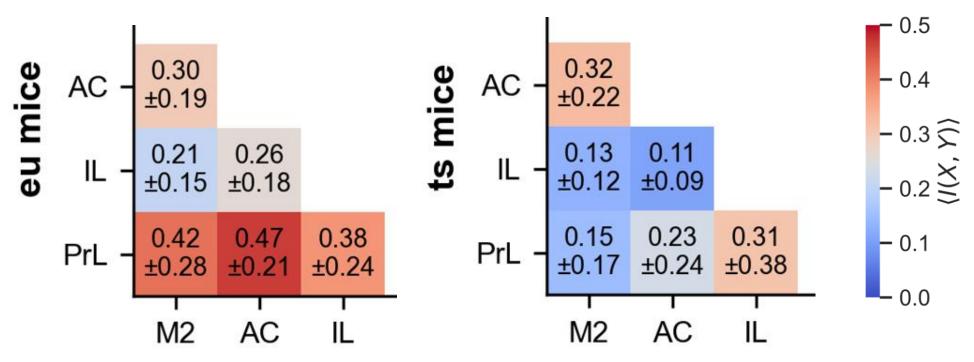


Neuropixels \rightarrow Spike trains \rightarrow Continuous time series of spiking rates with time bin $\Delta t \rightarrow$ Mutual Information measure between neurons pairs \rightarrow Average between populations and mice



Neuropixels → Spike trains → Spiking rates → Population averages by cortical region → Mutual Information measure between population pairs

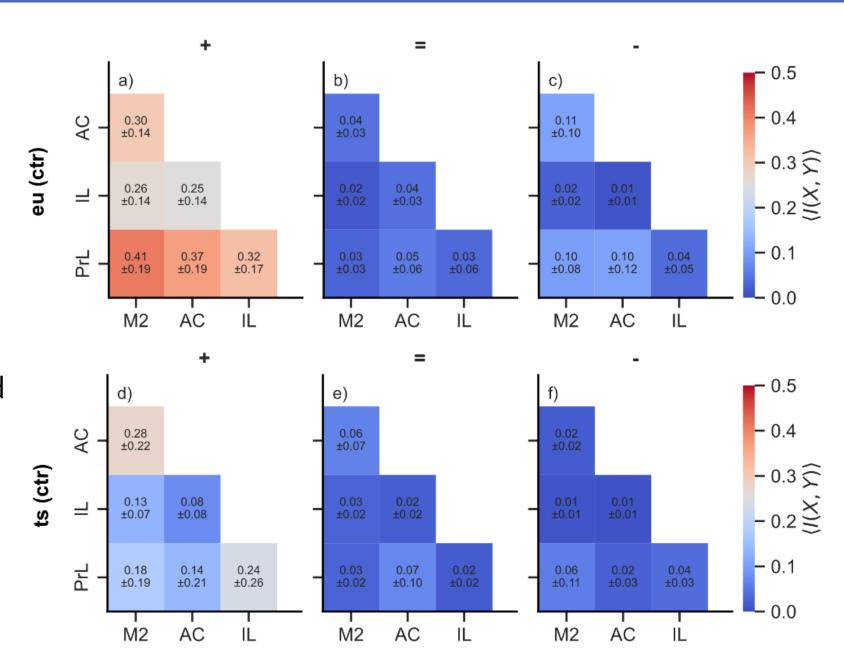
What if we now consider the MI between each population rate instead of single neurons?



Again, each cell shows the average also across mice (n = 6 for eu, n = 5 for ts)... but still errors are very big, why?

What if we also consider the classification attending to locomotion?

We see that the = and – guys are barely contributing to the encoding, while the + ones encode less



Take home ideas



- Mutual information makes no assumption about the data and can help us to reveal non-linear correlations and intricate encoding patterns
- In our case, it reveals specific connectivity patterns present only at network level and finding loss of coherence in the trisomic genotype

Thanks for your attention! Questions?

Special thanks to,

Alessandro Torcini



Alberto Bacci



Javier Zorrilla

