The Gatsby Data Center Distributing and Applying Advanced Statistical Tools to Understand How Neural Circuits Generate Behavior

Joaquín Rapela

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My Academic History



PhD in Electrical Engineering (Signal Processing) – University of Southern California

Postdoc on statistical analysis of EEG recordings – University California San Diego



Postdoc on statistical analysis of recordings from patients with epilepsy- Brown University



Postdoc (and hopefully research engineer soon) building the Gatsby Data Center – Gatsby Computational Neuroscience Unit

Gatsby Computational Neuroscience Unit

Theoretical Neuroscience

Neural data modeling Statistics of sensory stimuli Perception Signal processing Neural networks Learning

Machine Learning

Kernel methods Comparison of distributions Graphical models Networks Bayesian non-parametrics Symmetries

- Faculty: 4
- Postdocs: 9
- PhD students: 19



Joaquín Rapela (Gatsby)

The Gatsby Data Center

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- How does the brain make decisions and select actions?
- How does the brain learn and remember?

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Joaquín Rapela (Gatsby)

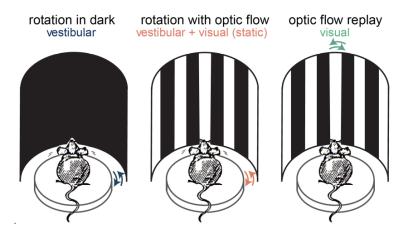
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Example experiment (1/3): behavioral and electrophysiological recordings related to social behavior



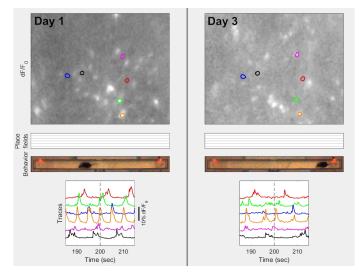
Cristina Mazuski (SWC)

Example experiment (2/3): electrophysiological recordings related to the integration of visual and vestibular information



Sepiedeh Keshavarzi (SWC)

Example experiment (3/3): optical recordings related to spatial navigation



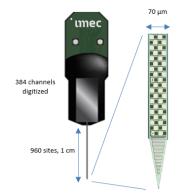
Sheintuch et al., 2017.

Joaquín Rapela (Gatsby)

Partial Summary

The SWC is doing sophisticated experiments to understand the relation between neural circuits and behavior.

Neuropixels Probes

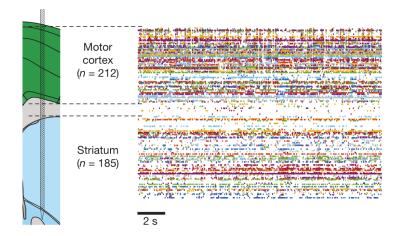


Neuropixels Waveforms

а b 0.56 1.09 V1B V2L -----0.93 1.05 0.37 1 mm

10 ms

Neuropixels Spikes



Partial Summary

The SWC is recording (electrically and optically) the activity of large populations of single neurons with state of the technology.

Mission:

- global: distribute high-quality implementations of advanced statistical methods devised at the unit, and allow neuroscientists without training in statistics to perform sophisticated analysis of their neural recordings,
- local: assist neuroscientists at the SWC in the use of advanced statistical methods to model their recordings.

Overview

 Collaboration: discovering the syllables of behavior in mouse social interactions

- 2 Distribution of the Sparse Variational Gaussian Processes Factor Analysis Python package
- 3 Collaboration: characterizing neural populations related to the integration of visual and vestibular information



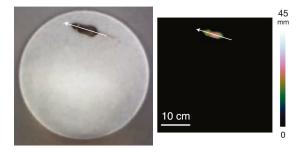
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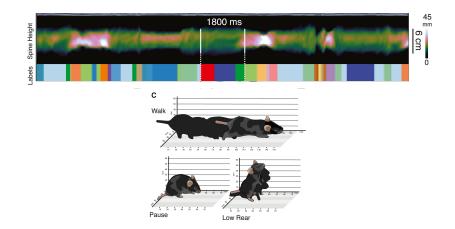
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Conclusions

Behavioral syllables for isolated mice have been reported $\left(1/2 \right)$



Behavioral syllables for isolated mice have been reported $\left(2/2\right)$



Discovering behavioral syllables in social interactions



- approach
- head-to-tail
- broadly-social
- following
- no-contact
- o conspecific-contact
- head-to-head
- head-to-flank

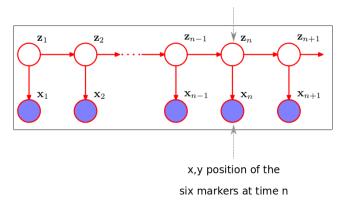
Cristina Mazuski (SWC)

Hidden Markov Model for finding behavioral syllables in social interactions

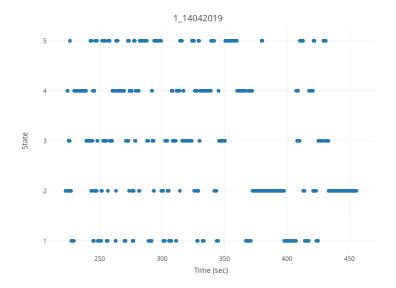
social behavioral state at time n

(e.g., following, approaching, head-

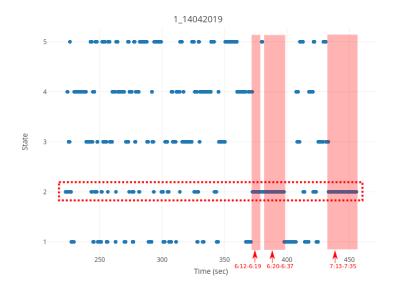
to-head, head-to-tail, etc)



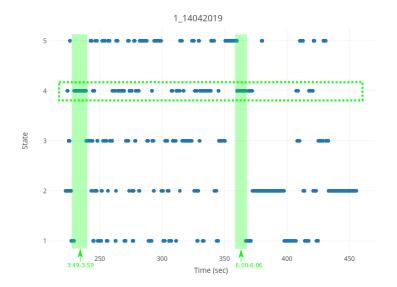
Behavioral syllables for social interactions



Non-Social state



Male-Head Female-Tail state



Joaquín Rapela (Gatsby)

Partial Summary

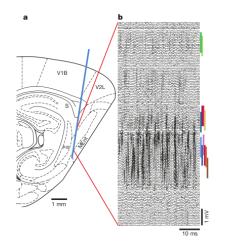
We gave the first steps in the quantitative characterization of social behavior in mice.

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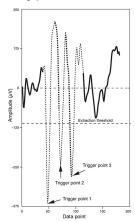
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Raw data

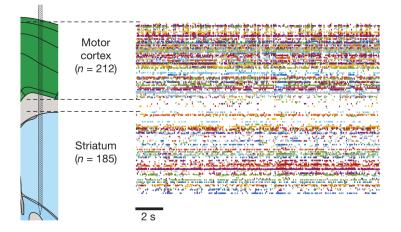


Extraction of spikes



A High-pass filtered data

Spikes data

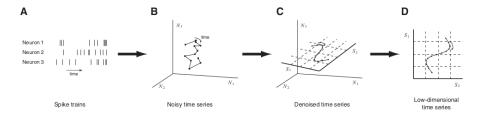


Our task now is to extract meaning from the spikes of populations of neurons.

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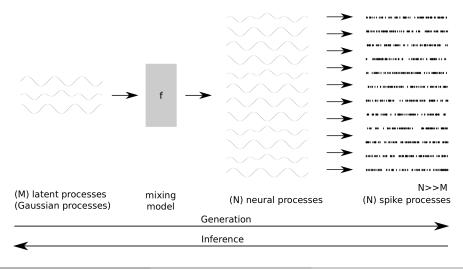
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Modeling neural populations spike recordings with Gaussian Processes Factor Analysis (intuition)



Yu et al., 2009

Modeling neural populations spike recordings with Gaussian Processes Factor Analysis (model sketch)



The Gatsby Data Center

Modeling neural populations spike recordings with Gaussian Processes Factor Analysis (mathematical expressions)

$$x_k(t) \sim GP(\mu_k(t), k_k(t, t')), \ k = 1, \dots, K$$

$$y_n(t) = \sum_{k=1}^{K} C[n, k] x_k(t) + d[n] + v_n(t), \ v_n(t) \sim N(0, \sigma_n), \ n = 1, \dots N$$

Letting
$$X(t) = [x_1(t), ..., x_K(t)], Y(t) = [y_1(t), ..., y_N(t)],$$

 $d = [d[1], ..., d[n]]$ and $R = diag(\sigma_1, ..., \sigma_N)$ the model is:

$$Y(t) \sim N(CX(t) + d, R)$$

Parameters: $\theta = [C, d, \{\mu_k, \text{hyperparameters of } \{k_k(t, t')\}_{k=1}^K\}]$ Yu et al., 2009 Modeling neural populations spike recordings with Gaussian Processes Factor Analysis (estimation)

Estimation using an approximate variational method with inducing points.

Variational method: approximates the full posterior $p(\theta|y_1, \ldots, y_n)$. Inducing points: allow to overcome the $O(N^3)$ complexity in Gaussian Process inference.

Duncker and Sahani, 2018

Goals of the distribution

- open source
- orrect
- fast running time
- easy to modify/extend
- appealing to users
 - ► GUI
 - documentation
 - support/help infrastructure

Tools for the distribution

- open source Github
- o correct Travis Cl
- fast running time GPyTorch
- easy to modify/extend OO Design: class and interaction diagrams
- appealing to users
 - ► GUI <mark>Q</mark>t
 - documentation Sphinx/Read the Docs
 - support/help infrastructure Github Issues, mailing list

Partial Summary

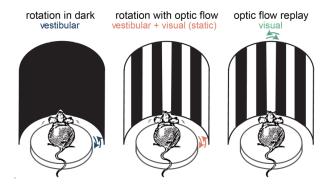
We made good progress in the distribution of the Python package Sparse Variational Gaussian Process Factor Analysis to model spiking activity in populations of neurons.

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Experiment



Sepiedeh Keshavarzi (SWC)

- Recordings from the same cell in the three conditions (visual, vestibular and visual+vestibular stimulation).
- Recordings from multiple brain regions, cortical layers and cell types (inhibitory and excitatory).
- Recordings from large populations of neurons (Neuropixels probe).

Joaquín Rapela (Gatsby)

The Gatsby Data Center

Question: In what cortical areas, layers, and neuron types there is integration of visual and vestibular information?

Hypothesis: The response of neurons that integrate visual and vestibular information should be more correlated to stimuli speed in conditions with visual+vestibular stimulation than in conditions with only visual- or vestibular-only stimulation.

Methodological approach

Fit a state-space model to predict spiking activity of neurons using stimuli speed inputs. Find brain regions, layers and cell types for which predictions of models in the visual+vestibular stimulation condition is significantly larger than predictions of models in the visual- or vestibular-only condition.

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- Istate-space model that can take inputs to generate predictions: Kalman filter estimated with the EM algorithm.
- 2 method to compare model predictions across cortical regions, cortical layers and cell types: analysis of variance (ANOVA).

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Examples of Kalman filter results

The Kalman filter yields excellent predictions. We found neurons that integrate visual and vestibular information.

Partial Summary

The Kalman filter appears to be a good model for large population recordings from the visual and vestibular integration experiment. In combination with the ANOVA, it may help us understand how multimodal information is integrated in the brain.

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- Its members are highly skilled in neural data analysis and machine learning methods, and they are excellent software engineers.
- The Gatsby+Data Center+SWC research demonstrates that the open-science approach produces excellent results.

Thanks



Cristina Mazuski



Sepiedeh Keshavarzi



Maneesh Sahani