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Application to the BBR24 funding opportunity

Enabling the Next Generation of Naturalistic Long-Duration Neuroscience Experimentation with Advanced Machine Learning

May 2, 2024

1 Summary

Word limit: 550

In plain English, provide a summary we can use to identify the most suitable experts to assess your application.

Clearly describe your proposed work in terms of:

- context
- the research the infrastructure, facility or resource will enable
- aims and objectives
- potential user communities, applications and benefits

2 Core team

List the key members of your team and assign them roles from the following:

- project lead (PL)
- project co-lead (UK) (PcL)
- researcher co-lead (RcL)
- specialist
- professional enabling staff
- research and innovation associate
- technician
- visiting researcher

Only list one individual as project lead.

A research technical professional or research software engineer can be listed as a project lead or project co-lead, provided that:

- their appointment is resourced from the central funds of their research organisation at the time of application
- their level of responsibilities and duties is appropriate to a person with substantial research experience
- their contract extends beyond the duration of the project

The researcher co-lead role has replaced the research co-investigator role previously used in Je-S grant applications. They will be an individual who merits appropriate recognition for making a substantial contribution to the formulation and development of the application and will be closely involved with the project.

project lead (PL) Dr. Joaquin Rapela

project co-lead (UK) (PcL) Prof. Tiago Branco, Prof. Maneesh Sahani, Prof. Thomas Mrcic-Flogel

researcher co-lead (RcL) Dr. Dario Campagner

specialist Dr. Nicholas Guilbeault

3 Application questions

3.1 Vision

Word limit: 1,700

What are you hoping to achieve with your proposed work?

What the assessors are looking for in your response

Explain how your proposed work:

- is of excellent quality and importance within or beyond the field(s) or area(s)
- has the potential to advance current understanding, or generate new knowledge, thinking or discovery within or beyond the field or area
- is timely given current trends, context, and needs
- impacts world-leading research, society, the economy, or the environment

Include the following in your statement:

- the uniqueness and expected added value of the proposed resource to the UK bioscience research community and infrastructure landscape
- how the resource relates to past and current resources in the subject area in both the UK and abroad
- full details of the resource and an overview of the associated objectives. Details on how these objectives are delivered should be included in the Approach section
- a description of the types of research that will be enabled by the resource
- consideration of the potential impact on the scientific community and other possibly dependent resources if the resource did not exist
- only if applicable, relevance of the proposed work to the plant health spotlight

In your vision, you should also clearly identify which of the following categories your proposed resource falls under, and expand on the relevant points raised below:

- establishment of a new and innovative resource that will be beneficial to a broader BBSRC user base. Explain why a new resource is needed and what unique and important features it will offer
- maturation and subsequent maintenance of a project-based resource into a community-based one. Briefly explain the background to the resource, current usage, proposed changes and the benefits this will lead to for the research community

- further development or essential maintenance of an existing community resource, with well-established access mechanisms. Explain current usage and how this project will increase its relevance, quality and utility, for example:
 - by enabling the resource to support FAIR (findable, accessible, interoperable, reusable) principles
 - by enabling new uses, for example metadata enrichment for machine learning and AI approaches
- association, or integration, of distinct resources. Explain current usage and how the proposed plans will create an upgraded resource with a greater value than the sum of the parts

References may be included within this section.

You may demonstrate elements of your responses in visual form if relevant. Further details are provided in the Funding Service.

Conventional versus modern neuroscience experimentation

Conventional system neuroscience experiments heavily constrain the behaviour of their subjects in order to simplify behavioural and neural analysis. However, important aspects of brain function may not be expressed in these constrained behaviours and these aspects may only manifest in naturalistic experimental conditions.

Recent technological advances now allow to build complex naturalistic experiments, while measuring a large number of experimental variables and recording from a large number of neurons simultaneously. These advances allow unrestrained subjects behaviour with precise monitoring of what happens in the subjects' environment and brain.

Important examples of these experiments are naturalistic foraging experiments, where freely moving subjects search for food in unrestrained environments. Foraging studies are currently underway at major research centres around the world (e.g., the Allen Institute for Brain Research, Janelia Farm, the Max Plank Institute of Animal Behaviour).

A complication of naturalistic neuroscience experiments is extracting meaning from the plethora of data that they generate (Juavinett, 2022). Machine learning methods are specially helpful to address this problem. For example, they can be used to extract subjects' internal states from behavioural measurements, to extract low-dimensional latent variables from high-dimensional neural recordings, and to relate the inferred subjects' internal state to the neural latent variables.

The data generated by conventional system neuroscience experiments can be understood by just plotting a few of its variables. However, interpreting the output of naturalistic experiments is more difficult and machine learning methods become essential for this.

Long-duration and naturalistic experiments at the SWC

We have developed a new platform that allows housing of mice for several weeks to months in a large enclosure (>2m diameter), while manipulating and monitoring their behaviour at high spatiotemporal resolution. The arena floor is tessellated to form a honeycomb of modular hexagonal tiles (Figure 1a), each of which can be equipped with a newly designed underground feeder (Figure 1b). Pellets are dispensed onto a foraging wheel once the mouse has spun it for a pre-defined programmable distance threshold using its forepaws (fictive digging). By modulating the digging cost/reward ratio at multiple arena location, this feeder allows the simulation of different foraging scenarios and probe mice behavioural strategies. The arena contains up to six scale-equipped nesting modules that allows housing of mice in the arena and weight monitoring. Mice can live in the arena for prolonged periods of time weeks to months and express their full natural behavioural repertoire.

Behavioural monitoring is achieved by an array of high-speed cameras (up to 15), by which mouse location, mouse identity and body parts can be track in real time.

Our platforms also enables continuous long term monitoring of neural activity using Neuropixels probes, the state of the art recording electrode, able to measure the activity of thousands of neurons simultaneously spanning the entire brain depth. Using this platform we have collected several week long datasets both with single mouse and multiple mice. This dataset capture a rich behavioural repertoire including a foraging behaviour, social learning task, defensive and nesting behaviours.

These long-duration experiments are allowing us to address foraging questions that cannot be studied in shorter experiments. For example, scientifically these experiments allow us to ask how do mice foraging patterns change across days, and what are the neural mechanisms underlying these changes. Statistically, the large amount of data recorded in the new experiments allows us to estimate parameters of much more complex models than those that can be fitted to data from conventional system neuroscience experiments.

Proposed resource

The initial SWC foraging project was supported by core funds from the SWC. The data it generated has only been analyzed by SWC and Gatsby Unit scientists. Due to its uniqueness, we aim to distribute the generated data widely to benefit the broad scientific community. However the complexity and size of these data create unprecented challenges for their distribution and analysis. Here we propose to mature this foraging project and create a resource to (1) share data from our foraging experiments to provide scientists across the world access to unprecedented experimental measurements, (2) share machine learning software implementations of methods to simulate and analyse these data to enable scientists around the world to generated insights from these recordings, (3) to promote the development of novel tools to simulate and analyze this data

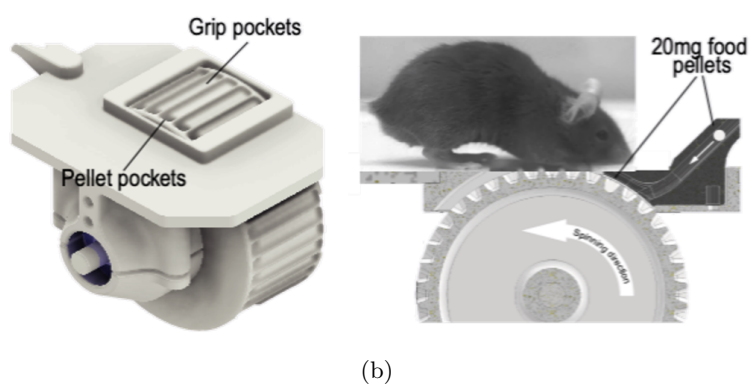
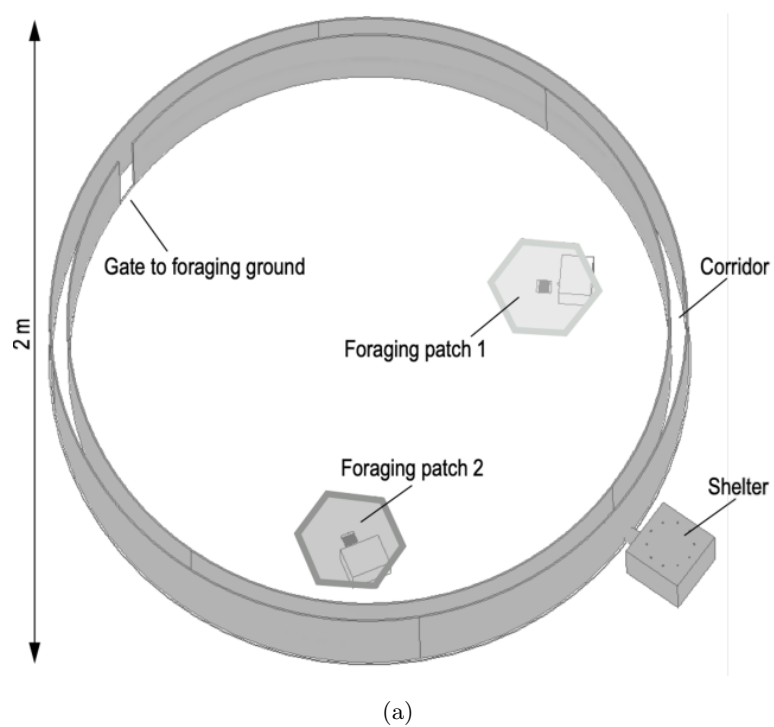


Figure 1: Foraging arena (a) and patch (b).

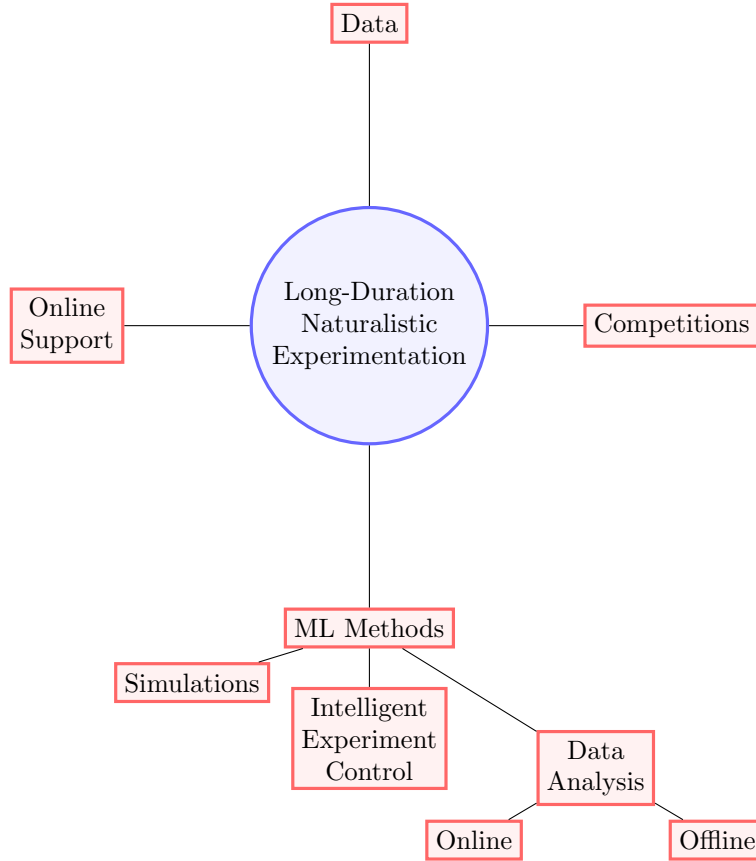


Figure 2: Resource theme (blue) and deliverables (red).

by organising foraging data simulations and analysis competitions and (4) provide online support to users of the resource to facilitate their use of the data and methods provided in this resource (Figure 2).

Data generated by our long-duration and naturalistic foraging experiments will be made publicly available, following FAIR standards, in the [DANDI](#) archive.

Whenever possible we will distribute offline and online implementations of all data analysis methods. Funded by a BBR award¹ we at the SWC, GCNU and NeuroGEARS are adding machine learning methods to Bonsai. This project will use will use these methods, add a few more methods to Bonsai, and use them to intelligently control our long-duration naturalistic foraging experiments with Bonsai.

We will initially distribute methods that we have already used for our foraging experiments. For several functionalities we have used more than one method,

¹[award details](#)

and we will distribute all of them. In this way experimental neuroscientists using our resource should be able to compare different distributed methods and choose the one more suitable to their needs.

The capability to simulate realistic foraging behaviour is important for our experiments. First, we often have to decide between several options to configure our foraging arenas (e.g., should we use two or three foraging patches? how much distance should mice travel to obtain a pellet? where should patches be located?). It would be helpful to evaluate the different options on simulated experiments, rather than evaluating them on expensive real ones. Second, we are often interested in understanding how much the behaviour of mice in our experiments differ from optimal behaviour. To address this issue we will simulate foraging agents and compare their behaviours with those of real mice.

We will encourage contributions from machine learning method developers interested in applying their methods to data generated by our foraging experiments. To motivate them to contribute their methods we will organise data competitions where participants will be given a data problem to solve (e.g., to simulate a given environment or to analyse a given dataset), they will provide their solutions, and we will select the winning one.

Intelligent experimental control with Bonsai

Bonsai is a reactive visual programming environment developed by NeuroGEARS Ltd that is widely used in Neuroscience for controlling sophisticated neuroscience experiments. Reactive programs are very different from procedural ones (like C or Python). The main entity of a reactive program is stream and a reactive program is a sequence of stream transformations that get an input stream, transform it, and generate an output one. Reactive programs are ideal to process online data, like those generated in animal neuroscience experiments.

Not all algorithms that we propose to distribute in this resource admit and online implementation, however most of them do. Whenever possible, we will distribute online versions implemented in Bonsai of the methods in this resource.

Online machine learning methods are relevant to long-duration naturalistic experimentation for at least two reasons. First the extremely large size of recorded dataset may forbid storing all raw data and online machine learning algorithms can help decide what data to store. For instance, we want to record the behaviour of multiple animals in large arenas with high resolution. This requires using multiple high-definition cameras to record videos of different parts of the arena. It is not feasible to store the videos by all cameras in long duration experiments. To overcome this problem, we can use probabilistic machine learning methods to track online the positions of the mice in the arena. When the tracking confidence of this methods is high, we would only save the high-resolution videos of the cameras filming mice, but when their confidence is low we would save the videos of all cameras.

Second, we want to make neural interventions informed by online inferences from machine learning methods. For example, in a foraging experiment we could find that a neural latent variable peaks before the instant when a mouse start

accelerating to leave the current patch (the latent variable could be estimated using a Poisson linear dynamical system and the acceleration using a linear dynamical system²). We could hypothesise that the neural population associated with the latent variable is responsible for the decision of leaving a patch. We could then test this hypothesis by optogenetically silencing this population while an animal is on a patch and checking if it leaves the patch or not.

Resource background

In July 2020 the SWC began building hardware and software infrastructure to record behaviour and neural activity of freely moving mice foraging in large arenas. Our initial focus was on monitoring behaviour for extended periods of time. We have been able to record behavioral videos, monitor the movement of the foraging wheels, register pellet deliveries and record mice weight for up to two weeks in experiments with two mice. More recently we have been able to record behaviour (as described above) and neural activity (Neuropixel array, four shanks, 384 electrodes per shank) continuously for up to 27 hours. Our next target is to record behavioral and electrophysiology data continuously for one month.

We expect that a position paper describing our foraging experiments will be published by the end of 2024.

Why is the proposed resource unique and timely

Long-duration and naturalistic experimentation is the future of experimental neuroscience. Animal foraging is a central neuroscience problem today and multiple groups around the world are working on it (e.g., Allen Institute for Neural Dynamics, Janelia Farm, University of Konstanz). Yet, none of this groups is focused on the unique long-duration and naturalistic experiments that we are developing at the SWC. The foraging data and the machine learning methods that we propose to distribute are key elements for world-class foraging research. If this resource is not created the emergence of naturalistic experimentation will be delayed and the UK may miss the opportunity of becoming a world leader on this new type of experimentation.

The US Brain Research through Advancing Innovative Neurotechnologies (BRAIN) initiative (Jorgenson et al., 2015) funded projects focused on data analysis with advanced machine learning methods for long-duration and complex experiment that we propose to address in this project. However, no BRAIN initiative project generated the unique long-duration and naturalistic neuroscience experimental data that we are creating at the SWC.

We have focused this proposal on applications of naturalistic and long-duration experimentation to experimental neuroscience. However, this type of experimentation is relevant to other scientific disciplines. For example, it could

²https://github.com/joacorapela/lds_python/blob/master/docs/tracking/tracking.pdf

we useful to improve the health of plants. We could perform long-duration experiments on the wild measuring many quantities from the plant environment (e.g., water, light, temperature, mineral nutrients, humidity) and from the plant physiology (e.g., photosynthesis, bud burst). We could then study how the environment affects plant physiology and health across multiple scales. Close loop interventions could then be performed (e.g., increasing water delivery) to improve the plant health.

Why are we a suitable team to create this resource?

We (the SWC, GCNU and NeuroGEARS) are an excellent team to develop the proposed resource. The SWC is at the forefront of experimental neuroscience research and has been developing mice foraging experiments for five years. The GCNU is a leader in computational neuroscience and machine learning, with ample experience in building methods to characterise neural data, and more recently in distributing openly machine learning methods. And NeuroGEARS has more than a decade of experience building high-quality software for experimental neuroscience. We collaborate extensively in a wide range of projects.

3.2 Approach

Word limit: 4,400

How are you going to deliver your proposed work?

What the assessors are looking for in your response

Explain how you have designed your approach so that it:

- is effective and appropriate to achieve your objectives
- is feasible, and comprehensively identifies any risks to delivery and how they will be managed
- uses a clearly written and transparent methodology (if applicable)
- summarises the previous work and describes how this will be built upon and progressed (if applicable)
- will maximise translation of outputs into outcomes and impacts
- describes how your, and if applicable your team's, research environment (in terms of the place and relevance to the project) will contribute to the success of the work

Include the following when describing your approach:

- measurable targets against which the outcome of the work will be assessed
- significant technical details for the development, maintenance or enhancement of the resource, indicating how this is of internationally exceptional quality
- any proposed research efforts and how they directly facilitate development of the resource (if applicable)
- if the focus is on maintaining an existing resource instead of suggesting further development, provide evidence of why significant upgrades are not required at this time and detail why the resource needs continued support to maintain world-leading functionality (if applicable)

Describe the specific contribution of each applicant to the proposed resource:

- their scientific contributions, for example, research field and specialist knowledge, experience, resource management expertise, technical and data analysis expertise
- their role and responsibilities, for example, managerial, leadership, mentoring
- references to specific work packages are recommended
- highlight where applicants will work collaboratively to deliver specific project requirements

- include clear time commitments for each applicant

There is no need to duplicate information included in the ‘Applicant and team capability to deliver’ section.

References may be included within this section.

You may demonstrate elements of your responses in visual form if relevant. Further details are provided in the Funding Service.

A project Gantt chart is compulsory and should be inserted as an image at the very end of this section. The Gantt chart should identify appropriate deliverables, responsibilities and time points for each objective.

The proposed resource will distribute behavioural and electrophysiological data from our foraging experiments and machine learning methods to simulate and analyse these data. We will initially contribute to the resource machine learning methods that we have used for the analysis of our foraging data. We will later invite method contributions from external method developers. To motivate these contributions we will organise foraging data simulation and data analysis competitions.

3.2.1 Distribution of long-duration and naturalistic foraging data

We will distribute in the **DANDI** archive data and metadata generated in our foraging experiments.

The data will include video recordings from all cameras, foraging wheels positions from all patches, pellet delivery times from all patches, mice weights recorded at the nest and ultrasound recordings of mice vocalisations.

The metadata will include subject information (birth data, sex and genetic strain) and experiment information (start date, duration and type – single or multiple mice).

We will also provide Python routines to access specific data and meta data items from the raw files³.

The data generated by our experiments is very large. Storing one hour of electrophysiological recordings requires xx gigabytes and one hour of video recordings requires yy gigabytes. For this reason we will initially only share publicly in the **DANDI** archive a small number of experimental sessions. Investigating alternative methods to share the large amounts of data generated by our experiments is a research item proposed for this resource (Section 3.2.5).

3.2.2 Methods for data analysis

Below we describe a few methods that we have used, or plan to use, to control/characterize our foraging data. We may or may not use all of these methods, or we may use other methods not listed there. It is difficult to anticipate the challenges that we might encounter when processing our long-duration, multi-modal and complex datasets, specially when adding to the analysis electrophys-

³https://github.com/SainsburyWellcomeCentre/aeon_mecha

iological recordings. Yet we are confident that we will excel on this processing, as we are a team with unique experience to do so (Section 3.4).

Linear dynamical systems models Linear dynamical systems (LDS) models are used to characterise time-varying observations as a function of hidden (i.e., latent) continuous state variables that vary over time. It assumes that both state and observations are Gaussian random vectors. The Kalman filter algorithm can be used to infer the probability distribution of the state given observations. More information about LDS models can be found in (Durbin and Koopman, 2012, part I). They can be used to model behavioural and neural data.

We have developed an implementation of LDS models⁴ and used it to infer denoised positions, velocities and accelerations of foraging mice from noisy and incomplete position measurements offline and online with Bonsai. We have also used this implementation to characterise electrophysiological recordings in mice.

Nonlinear and non-Gaussian dynamical systems models LDS models are very versatile and can be used to model a wide range of time-series observations. However, there are cases where this model does not apply and nonlinear and non-Gaussians extensions are needed, like extended Kalman filter, the unscented Kalman filter and the particle filter (Durbin and Koopman, 2012, part II).

We have created the Poisson Linear Dynamical System (PLDS) model to characterise count observations (Macke et al., 2015) and distributed a Matlab implementation⁵. This is an offline implementation, and we have not yet produced an online one. We have neither use PLDS to model behavioural or neural foraging data.

Gaussian processes models Gaussian process factor analysis (GPFA) models (Yu et al., 2009) are similar to LDS ones in that the probability distribution of observation is a function of latent variable. However, differently from LDS, where latent variables have linear dynamics, in GPFA models latent variables are samples from a Gaussian process and have nonlinear dynamics. We contributed to the development of a Matlab implementation of GPFA⁶.

GPFA is used to characterise neural activity. It requires binning of spikes into spikes counts and assumes that spike counts follow a Gaussian distribution. We have recently introduced sparse variational Gaussian process factor analysis (svGPFA, Duncker and Sahani, 2018) that uses point process observations and does not require to bin spikes times and developed a Python implementation of it⁷.

We have not yet attempted online versions of these algorithms.

⁴https://github.com/joacorapela/lds_python

⁵https://bitbucket.org/mackelab/pop_spike_dyn

⁶<https://users.ece.cmu.edu/~byronyu/software.shtml>

⁷<https://github.com/joacorapela/svGPFA>

Hidden Markov models The hidden Markov model (HMM) is a latent variable model similar to the LDS model, but where the hidden states are discrete (Bishop, 2016, Chapter 13). It is used to assign discrete labels to time series observations.

We have developed an offline implementation⁸ of the HMM, used it to find repeatable states in epileptic seizures of human subjects, from Utah array recordings of their neural activity (Rapela and Todorov, 2019). We have also used the HMM to infer mouse foraging states from kinematic inferences from the LDS model. We are currently building an online implementation of the HMM in Bonsai.

Switching linear dynamical systems models A switching linear dynamical system (SLDS) model is a hybrid/nonlinear system which consists of several linear subsystems and a switching rule that decides which of the subsystems is active at each moment in time (Murphy, 2012, Section 18.6).

Over long periods of time the behaviour and neural activity of mice may alternate between different states, where each state can be well modelled by a linear dynamical system. Long-duration behavioural and neural time series could be well modelled by SLDS models.

We have not yet used SLDS models to characterise our foraging data.

Recognition parameterised models

Generalised linear models Generalised linear models (GLMs) are regression models for observations with diverse noise distributions (i.e., noise distributions in the exponential family). For example, they can be used to estimate a linear regression model with spike count as the observation variable.

We have used a Gaussian GLM (i.e., a standard linear regression model) to study how different regressors (e.g., average speed or acceleration before entering a patch, amount of reward obtained in the previous visit to the current patch, amount of reward obtained in the previous visit to the other patch, weights) influence the length of a foraging bout in a short (three hours) experimental session. The predictive power of this model was poor.

We have used online Bayesian linear regression models in Bonsai (implemented using linear dynamical systems) to estimate receptive fields of cells in primary visual cortex⁹.

Deep neural networks Deep neural networks are powerful nonlinear function approximators used in supervised learning (Goodfellow et al., 2016). This networks require large amounts of training data to achieve good performance. Thus, they are not good models for conventional experiments generating smaller

⁸<https://github.com/joacorapela/hiddenMarkovModels>

⁹<https://ncguilbeault.github.io/machinelearning/examples/examples/LinearDynamicalSystems/LinearRegression/ReceptiveFieldSimpleCell/README.html>

datasets. However, they are promising models for experiments generating large datasets, such as ours.

For example, we plan to use them to predict the duration of mice foraging bouts from a large set of regressors, as mentioned above, but using long experimental sessions. The hope is that deep neural networks trained with datasets from our long-duration foraging experiments will overcome the limitations of linear regression models and achieve excellent predictive power.

We are currently adding to Bonsai functionality to use pre-trained PyTorch networks¹⁰.

Multiple body parts tracking and pose estimation methods Deep neural networks have proved very successful for tracking animal body parts in video recordings. We have used DeepLabCut (Mathis et al., 2018) for tracking body parts in single-animal experiments and SLEAP (Pereira et al., 2022) for tracking body parts in multiple-animal experiments. Other methods for tracking animal body parts have recently become available (e.g., multi-animal DeepLabCut (Lauer et al., 2022) and lighting pose (Biderman et al., 2023)). We will apply these methods to our long-duration foraging recordings and report comparisons of their performance.

MoSeq Wiltchko et al. (2015) is a pose estimation method that attempts to decompose the behaviour of animals into repeatable behavioural motifs. It can use as inputs animal body parts (e.g., outputs from DeepLabCut) and it is based on hidden Markov models. We have used hidden Markov models to find repeatable motifs in foraging mouse, but using as inputs kinematic variables of mice centre of mass instead of multiple body parts. We will compare the motifs estimated by MoSeq and by our HMMs.

An important question for our experiments is the stability of the learned models; i.e., will a model estimated with data from the first experiment day perform well when used to track body part or infer behavioural syllables in data from the fifth experiment day? If not, how frequently should the model parameters be updated in our long-duration naturalistic experiments?

Bonsai can already track online animal body parts in video recording using pre-trained DeepLabCut models¹¹ (Kane et al., 2020). However, the utility of this capability for our foraging experiments depends critically on the stability of the learned DeepLabCut models.

Other methods to estimate neural latent variables Two recent popular methods to estimate neural latent variables are *Latent Factor Analysis via Dynamical Systems* (LFADS Pandarinath et al., 2018), a method based on recurrent neural networks and autoencoders, and *Computational Embedding of Behavioural Response with Adaptive-weights* (Schneider et al., 2023, CEBRA), a method based on contrastive learning. We propose to compare their performance to that of more classical methods to estimate neural latents (e.g., LDS,

¹⁰<https://pytorch.org/vision/stable/models.html>

¹¹<https://github.com/bonsai-rx/deeplabcut>

PLDS, GPFA, svGPFA) on data generated by our long-duration naturalist experiments.

3.2.3 Simulation of foraging data

We will build simulated foraging arenas where virtual agents can forage, run and sleep. Then we will create reinforcement learning agents for this arena. Deep reinforcement learning has been successfully applied to simulate foraging behaviours (Wispiński et al., 2022), and we will use similar approaches in our simulations.

3.2.4 Data competitions

To motivate contributions by external machine learning methods developers of methods to process long-duration naturalistic experiments we will organise neural data analysis and behavioural data simulation competitions.

For the neural data competitions we will provide participants with a neural dataset and ask them to perform a given inference. For example, we could provide them behavioural (e.g., body parts positions, reward delivery, travelled distance) and neural recordings (e.g., spikes in prefrontal and visual areas) and ask them to use the recordings when an animal is in a patch to predict the moment at which it will leave the patch. Competitions will happen online, like the *Neural Latents Challenge* (?); they will be announced, participants will download datasets and work on their solution, they will submit them, and we will evaluate them following pre-specified metrics.

The reinforcement learning competitions will also happen online, like most other reinforcement learning competitions¹². We will build a virtual foraging arena and ask participants to build agents to forage in it as mice forage in real arenas. We will evaluate contributions following pre-specified metrics.

Competitions will finish with a conference, where top participants will present their algorithms to other participants and to experimental user of our resource.

3.2.5 Research items

Below we comment on research items required for the successful creation of the proposed resource.

Distribution of very large experimental data For the first version of the resource we will ask users to download large files to their computers and we will provide Python functions¹³ to extract relevant items (e.g., behavioural videos, travelled distance, pellets delivery times) from these files.

However, this method may not be convenient to all users. Another method is to provide users a data streaming API so that they can stream to their computer

¹²<https://github.com/seungjaeryanlee/awesome-rl-competitions>

¹³https://github.com/SainsburyWellcomeCentre/aeon_mecha

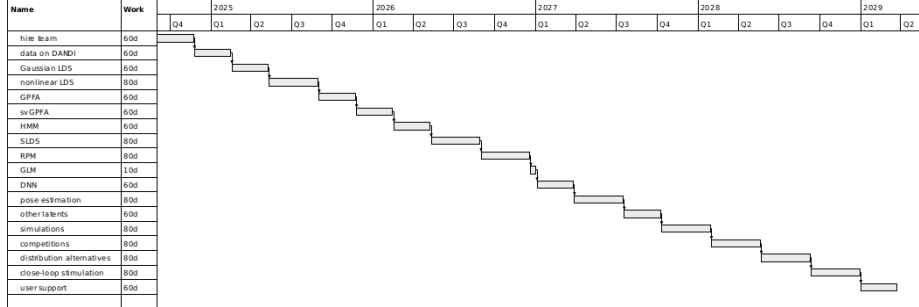


Figure 3: Schedule of proposed tasks. Note that it is difficult to forecast the exact trajectory of the project, as unexpected challenges may appear in the development of methods to process a radically new type of data.

segments of the complete dataset. This is the approach taken by the Open Neurophysiology Environment¹⁴ of the International Brain Laboratory (Bonacchi et al., 2023).

A third method is to store data and code in a cloud. This method has two advantages. First, data is local in the cloud, which saves data transfer time. Second, configuring advanced neural data analysis pipelines can be complex and this method frees users from the complexities of doing so. This configuration will be done on a central cloud location by a system administrator of the resource. Neuroscience in the Cloud Analysis As a Service (NeuroCAAS Abe et al., 2022) uses this method. However, NeuroCAAS focuses on the second previous advantage while our focus is on the first one.

Learning and inference for very large datasets Conventional methods for learning and inference are not feasible for very large datasets, as the ones we are collecting at the SWC. Special methods have been developed for this purpose, like stochastic variational inference (Hoffman et al., 2013). However, some of these methods assume that data is independent or exchangeable, which does not hold for time series. Thus, we should investigate learning and inference methods that can scale to the size and properties of our foraging datasets.

Accelerated and/or distributed computing Even if we can theoretically perform inference with large datasets, this inference could be prohibitively slow. We should research and develop efficient implementations using accelerated and/or distributed computing.

3.2.6 Tasks

Figure 3 shows a gnatt chart for the proposed tasks.

¹⁴<https://int-brain-lab.github.io/ONE/index.html>

3.3 Community demand: letters (or emails) of support

Letters (or emails) of support demonstrating community demand are mandatory for BBR.

Upload a single PDF of maximum 8MB containing a maximum of 10 letters or emails of support. These should be uploaded in English or Welsh only. Enter the words ‘attachment supplied’ in the text box.

What the assessors are looking for in your response

The letters should give an indication of community demand for the resource in question, demonstrating the breadth of research and the high-quality science relevant to BBSRC remit that the resource would underpin.

Add the following details for each letter:

1. the organisation name (searchable via a drop-down list or enter the organisation’s details manually, as applicable)
2. contact name of the signatory

Letters of support aimed at demonstrating community demand should:

1. outline the uniqueness and expected added value of the proposed resource to the UK bioscience research community and infrastructure landscape
2. clearly explain the impact and benefit of the proposed resource on the writer’s research and the associated community
3. if possible, explain where this supported research has already demonstrated or could have potential for particular scientific, economic or societal impact
4. help to demonstrate the breadth of the relevant user community

Letters of support that fail to do so, in particular template letters indicating generic support without identifying a particular usage, are of negligible value for the assessment and should not be submitted. Carefully chosen letters containing relevant evidence of the requirement or benefit to be gained, are of greater value than large numbers of letters.

The Funding Service will provide document upload details when you apply.

3.4 Applicant and team capability to deliver

Word limit: 1,650

Why are you the right individual or team to successfully deliver the proposed work?

What the assessors are looking for in your response

Evidence of how you, and if relevant your team, have:

- the relevant experience (appropriate to career stage) to deliver the proposed work
- the right balance of skills and expertise to cover the proposed work
- the appropriate leadership and management skills to deliver the work and your approach to develop others
- contributed to developing a positive research environment and wider community

You may demonstrate elements of your responses in visual form if relevant. Further details are provided in the Funding Service.

The word count for this section is 1,650 words: 1,150 words to be used for R4RI modules (including references) and, if necessary, a further 500 words for Additions.

Use the Résumé for Research and Innovation (R4RI) format to showcase the range of relevant skills you and, if relevant, your team (project and project co-leads, researchers, technicians, specialists, partners and so on) have and how this will help deliver the proposed work. You can include individuals' specific achievements but only choose past contributions that best evidence their ability to deliver this work.

Complete this section using the R4RI module headings listed. Use each heading once and include a response for the whole team, see the UKRI guidance on R4RI. You should consider how to balance your answer, and emphasise where appropriate the key skills each team member brings:

- contributions to the generation of new ideas, tools, methodologies, or knowledge
- the development of others and maintenance of effective working relationships
- contributions to the wider research and innovation community
- contributions to broader research or innovation users and audiences and towards wider societal benefit

Additions

Provide any further details relevant to your application. This section is optional and can be up to 500 words. You should not use it to describe additional

skills, experiences, or outputs, but you can use it to describe any factors that provide context for the rest of your R4RI (for example, details of career breaks if you wish to disclose them).

Complete this as a narrative. Do not format it like a CV.

There is no need to duplicate information included in the ‘Approach’ section.

UKRI has introduced new role types for funding opportunities being run on the new Funding Service.

For full details, see [Eligibility as an individual](#).

References may be included within this section.

More information can be found in the following [Annotated guidance on preparing a UKRI R4RI](#).

3.5 Contributions to the generation of new ideas, tools, methodologies, or knowledge

In 2021 Dr. Rapela and Dr. Lopes generated the main ideas of the BBSRC funded award [Machine learning for neuroscience experimental control](#) and in collaboration with Dr. Guilbeault they are now building the [Bonsai.ML](#) package, which adds machine learning functionality to the [Bonsai](#) ecosystem.

In 2013 Dr. Rapela was awarded a seed grant from the Center for Brain Activity Mapping of the Kavli Institute for Brain and Mind at UC San Diego titled *Taking the next step toward understanding computations by neural ensembles with high resolution neural recordings, generic data assimilation methods, and increased computational power*.

Dr. Rapela uses advanced statistical methods to understand brain function. He has generated knowledge on the estimation of receptive fields of visual neurons stimulated with natural images ([Rapela et al., 2006, 2010](#)), on the use of the electroencephalogram to understand cognition and mental disorders in humans ([Rapela et al., 2012a,b, 2018](#)), on the use of nonlinear dynamical systems to model neural population activity ([Rapela et al., 2015](#)), on the use of surface cortical recordings in humans to understand the relation between brain oscillations and speech ([Rapela, 2016, 2017, 2018](#)) and on the use of intracortical recordings in humans to understand the neural basis of epilepsy ([Rapela et al., 2019; Rapela and Todorov, 2019](#)).

Dr. Rapela is the lead developer of several open-source statistical algorithms with applications in neuroscience like [Sparse Variational Gaussian Process Factor Analysis](#), [Linear Dynamical Systems](#) and [Hidden Markov Models](#).

3.6 The development of others and maintenance of effective working relationships

In 2011, Dr. Rapela has co-mentored Mr. Tsong-Yan Lin during his MSc studies at UC San Diego ([Rapela et al., 2012b](#)). In 2021 and 2023 he has mentored Ms. Aisha Qureshi and Ms. Zimo Li, respectively, in seven-month-long [Shenoy Undergraduate Research Fellowship in Neuroscience \(SURFiN\)](#) fellowships, sponsored by the [Simons Collaboration on the Global Brain](#).

As a computational neuroscientist Dr. Rapela has build multiple collaborations with experimental neuroscientists. For example, with Dr. Jon Touryan and Dr. Guidon Felsen ([Rapela et al., 2006, 2010](#)). with Prof. Jeanne Townsend ([Rapela et al., 2012a,b, 2018](#)) and with Prof. Edward Chang ([Rapela et al., 2015](#); [Rapela, 2016, 2017, 2018](#)).

3.7 Contributions to the wider research and innovation community

3.8 Contributions to broader research or innovation users and audiences and towards wider societal benefit

3.9 Project partners

Add details about any project partners' contributions. If there are no project partners, you can indicate this on the Funding Service.

A project partner is a collaborating organisation who will have an integral role in the proposed research. This may include direct (cash) or indirect (in-kind) contributions such as expertise, staff time or use of facilities.

Add the following project partner details:

- the organisation name and address (searchable via a drop-down list or enter the organisation's details manually, as applicable)
- the project partner contact name and email address
- the type of contribution (direct or in-direct) and its monetary value

If a detail is entered incorrectly and you have saved the entry, remove the specific project partner record and re-add it with the correct information.

For audit purposes, UKRI requires formal collaboration agreements to be put in place if an award is made.

3.10 Project partners: letters (or emails) of support

Upload a single PDF containing the letters or emails of support from each partner you named in the Project Partner section. These should be uploaded in English or Welsh only.

Enter the words ‘attachment supplied’ in the text box, or if you do not have any project partners enter N/A.

What the assessors are looking for in your response Each letter or email you provide should:

- confirm the partner’s commitment to the project
- clearly explain the value, relevance, and possible benefits of the work to them
- describe any additional value that they bring to the project

The Funding Service will provide document upload details when you apply. If you do not have any project partners, you will be able to indicate this in the Funding Service.

Ensure you have prior agreement from project partners so that, if you are offered funding, they will support your project as indicated in the project partners’ section.

For audit purposes, UKRI requires formal collaboration agreements to be put in place if an award is made.

3.11 Management strategy

Word limit: 500

How do you plan to manage the resource?

What the assessors are looking for in your response

1. resources will be expected to have governance arrangements appropriate for the oversight and successful delivery of the project's complexity
2. provide details about the project's management and advisory structure
3. provide details of the approach to project and risk management, and the monitoring strategy for tracking progress of the proposed programme
4. provide details on how demand and access requests will be managed, and what support will be provided to the users of the resource
5. an advisory board is required for all projects, which is independent from both the academic institutions and project partners involved in the proposal. Provide information on the proposed membership of this advisory board and how it will be used
6. provide details on how the resource user perspective and their needs will be considered, including how feedback will be sought and subsequently used to inform the management of the resource

You may demonstrate elements of your responses in visual form if relevant. Further details are provided in the Funding Service.

3.12 Data management and sharing

Word limit: 1,500

How will you manage and share data collected or acquired as part of the proposed resource?

What the assessors are looking for in your response

Provide a data management plan using the [BBR DMP template \(PDF, 161KB\)](#) structure that clearly details how your proposed resource will comply with UKRI's published [data sharing policy](#), which includes detailed guidance notes.

3.13 Trusted Research and Innovation (TR&I)

Word limit: 500

Does the proposed work involve international collaboration in a sensitive research or technology area?

What the assessors are looking for in your response

Demonstrate how your proposed international collaboration relates to trusted research and innovation, including:

- list the countries your international project co-leads, project partners and visiting researchers, or other collaborators are based in
- if international collaboration is involved, explain whether this project is relevant to one or more of the 17 areas of the UK National Security and Investment (NSI) Act
- if one or more of the 17 areas of the UK National Security and Investment (NSI) Act are involved, please identify which areas

If your proposed work does not involve any international collaboration, answer 'n/a' here.

We may ask you to provide additional information about how your proposed project will comply with our approach and expectation towards TR&I, identifying potential risks and the relevant controls you will put in place to help proportionately reduce these risks.

3.14 Resources and cost justification

Word limit: 1,000

What will you need to deliver your proposed work and how much will it cost?

What the assessors are looking for in your response

Justify the application's more costly resources, in particular:

- project staff
- significant travel for field work or collaboration (but not regular travel between collaborating organisations or to conferences)
- any equipment that will cost more than £10,000
- any consumables beyond typical requirements, or that are required in exceptional quantities
- all facilities and infrastructure costs
- all resources that have been costed as 'Exceptions'

Assessors are not looking for detailed costs or a line-by-line breakdown of all project resources. Overall, they want you to demonstrate how the resources you anticipate needing for your proposed work:

- are comprehensive, appropriate, and justified
- represent the optimal use of resources to achieve the intended outcomes
- maximise potential outcomes and impacts
- evidence appropriate consideration for alternative long-term sustainability options beyond BBSRC funding

3.15 Your organisation’s support

Word limit: 500

Provide details of support from your research organisation and project co-lead research organisations.

What the assessors are looking for in your response

Provide a statement of support from all participating research organisations detailing why they are best placed to support the proposed project. This should include details of any matched funding that will be provided to support the activity and any additional support that might add value to the work.

Assessors will be looking for a strong statement of commitment from your research organisations.

BBSRC recognises that in some instances, this information may be provided by the Research Office, the Technology Transfer Office (TTO) or equivalent, or a combination of both.

You must also include the following details:

- a significant person’s name and their position, from the TTO or Research Office, or both
- office address or web link

Upload details are provided within the service on the actual application.

We do not require separate institutional letters of support as attachments. By submitting your application to us, you are confirming that your institutions are supportive of and committed to your project.

References

- Abe, T., Kinsella, I., Saxena, S., Buchanan, E. K., Couto, J., Briggs, J., Kitt, S. L., Glassman, R., Zhou, J., Paninski, L., et al. (2022). Neuroscience cloud analysis as a service: An open-source platform for scalable, reproducible data analysis. *Neuron*, 110(17):2771–2789.
- Biderman, D., Whiteway, M. R., Hurwitz, C., Greenspan, N., Lee, R. S., Vishnubhotla, A., Warren, R., Pedraja, F., Noone, D., Schartner, M., Huntenburg, J. M., Khanal, A., Meijer, G. T., Noel, J.-P., Pan-Vazquez, A., Socha, K. Z., Urai, A. E., Laboratory, T. I. B., Cunningham, J. P., Sawtell, N., and Paninski, L. (2023). Lightning pose: improved animal pose estimation via semi-supervised learning, bayesian ensembling, and cloud-native open-source tools. *bioRxiv*.
- Bishop, C. M. (2016). *Pattern recognition and machine learning*. Springer-Verlag New York.
- Bonacchi, N., Chapuis, G. A., Churchland, A. K., DeWitt, E. E., Faulkner, M., Harris, K. D., Huntenburg, J. M., Hunter, M., Laranjeira, I. C., et al.

- (2023). A modular architecture for organizing, processing and sharing neurophysiology data. *Nature methods*, 20(3):403–407.
- Duncker, L. and Sahani, M. (2018). Temporal alignment and latent gaussian process factor inference in population spike trains. In *Advances in Neural Information Processing Systems*, pages 10445–10455.
- Durbin, J. and Koopman, S. J. (2012). *Time series analysis by state space methods*, volume 38. OUP Oxford.
- Goodfellow, I., Bengio, Y., and Courville, A. (2016). *Deep learning*. MIT press.
- Hoffman, M. D., Blei, D. M., Wang, C., and Paisley, J. (2013). Stochastic variational inference. *Journal of Machine Learning Research*.
- Jorgenson, L. A., Newsome, W. T., Anderson, D. J., Bargmann, C. I., Brown, E. N., Deisseroth, K., Donoghue, J. P., Hudson, K. L., Ling, G. S., MacLeish, P. R., et al. (2015). The brain initiative: developing technology to catalyse neuroscience discovery. *Philosophical Transactions of the Royal Society B: Biological Sciences*, 370(1668):20140164.
- Juavinett, A. (2022). In foraging tasks, new opportunities to study the brain’s complexity. <https://www.simonsfoundation.org/2022/02/23/in-foraging-tasks-new-opportunities-to-study-the-brains-complexity/>.
- Kane, G. A., Lopes, G., Saunders, J. L., Mathis, A., and Mathis, M. W. (2020). Real-time, low-latency closed-loop feedback using markerless posture tracking. *eLife*, 9:e61909.
- Lauer, J., Zhou, M., Ye, S., Menegas, W., Schneider, S., Nath, T., Rahman, M. M., Di Santo, V., Soberanes, D., Feng, G., et al. (2022). Multi-animal pose estimation, identification and tracking with deeplabcut. *Nature Methods*, 19(4):496–504.
- Macke, J. H., Buesing, L., and Sahani, M. (2015). *Estimating state and parameters in state space models of spike trains*, page 137–159. Cambridge University Press.
- Mathis, A., Mamidanna, P., Cury, K. M., Abe, T., Murthy, V. N., Mathis, M. W., and Bethge, M. (2018). Deeplabcut: markerless pose estimation of user-defined body parts with deep learning. *Nature neuroscience*, 21(9):1281–1289.
- Murphy, K. P. (2012). *Machine learning: a probabilistic perspective*. MIT press.
- Pandarínath, C., O’Shea, D. J., Collins, J., Jozefowicz, R., Stavisky, S. D., Kao, J. C., Trautmann, E. M., Kaufman, M. T., Ryu, S. I., Hochberg, L. R., et al. (2018). Inferring single-trial neural population dynamics using sequential auto-encoders. *Nature methods*, 15(10):805–815.

- Pereira, T. D., Tabris, N., Matsliah, A., Turner, D. M., Li, J., Ravindranath, S., Papadoyannis, E. S., Normand, E., Deutsch, D. S., Wang, Z. Y., McKenzie-Smith, G. C., Mitelut, C. C., Castro, M. D., D’Uva, J., Kislin, M., Sanes, D. H., Kocher, S. D., S-H, S., Falkner, A. L., Shaevitz, J. W., and Murthy, M. (2022). Slep: A deep learning system for multi-animal pose tracking. *Nature Methods*, 19(4).
- Rapela, J. (2016). Entrainment of traveling waves to rhythmic motor acts.
- Rapela, J. (2017). Rhythmic production of consonant-vowel syllables synchronizes traveling waves in speech-processing brain regions.
- Rapela, J. (2018). Traveling waves appear and disappear in unison with produced speech.
- Rapela, J., Felsen, G., Touryan, J., Mendel, J. M., and Grzywacz, N. M. (2010). ePPR: a new strategy for the characterization of sensory cells from input/output data. *Network: Computation in Neural Systems*, 21(1-2):35–90.
- Rapela, J., Gramann, K., Westerfield, M., Townsend, J., and Makeig, S. (2012a). Brain oscillations in switching vs. focusing audio-visual attention. In *Proc. of 2012 IEEE Engineering in Medicine and Biology Society (EMBC’12)*, San Diego, CA.
- Rapela, J., Lin, T.-Y., Westerfield, M., Jung, T.-P., and Townsend, J. (2012b). Assisting autistic children with wireless EOG technology. In *Proc. of 2012 IEEE Engineering in Medicine and Biology Society (EMBC’12)*, San Diego, CA.
- Rapela, J., Mendel, J., and Grzywacz, N. (2006). Estimating nonlinear receptive fields from natural images. *Journal of Vision*, 6(4):441–474.
- Rapela, J., Proix, T., Todorov, D., and Truccolo, W. (2019). Uncovering low-dimensional structures of high-dimensional electrophysiological recordings in epilepsy. In *Engineering in Medicine and Biology Society (EMBC), 2019 41st Annual International Conference of the IEEE*. IEEE.
- Rapela, J., Rowat, P. F., Kostuk, M., Mullen, T., Chang, E. F., and Bouchard, K. E. (2015). Modeling neural activity at the ensemble level.
- Rapela, J. and Todorov, D. (2019). Hidden markov models applied to LFPs from layer two and three of human cortex reveal highly stereotypical discrete states in epileptic seizures separated by more than an hour. Accepted at the 41st Engineering in Medicine and Biology Conference of the IEEE; main text https://sccn.ucsd.edu/~rapela/papers/EMBC19_Rapela_and_Todorov_EMBC_2019.pdf and supplemental information [embc19_hmm_supplemental.pdf](#).

- Rapela, J., Westerfield, M., and Townsend, J. (2018). A new single-trial foreperiod effect on inter-trial phase coherence. Part I: existence and relevance. *Neural Computation*, 30(9):2348–2383.
- Schneider, S., Lee, J. H., and Mathis, M. W. (2023). Learnable latent embeddings for joint behavioural and neural analysis. *Nature*, 617(7960):360–368.
- Wiltschko, A. B., Johnson, M. J., Iurilli, G., Peterson, R. E., Katon, J. M., Pashkovski, S. L., Abaira, V. E., Adams, R. P., and Datta, S. R. (2015). Mapping sub-second structure in mouse behavior. *Neuron*, 88(6):1121–1135.
- Wispinski, N. J., Butcher, A., Mathewson, K. W., Chapman, C. S., Botvinick, M. M., and Pilarski, P. M. (2022). Adaptive patch foraging in deep reinforcement learning agents. *arXiv preprint arXiv:2210.08085*.
- Yu, B. M., Cunningham, J. P., Santhanam, G., Ryu, S. I., Shenoy, K. V., and Sahani, M. (2009). Gaussian-process factor analysis for low-dimensional single-trial analysis of neural population activity. *Journal of neurophysiology*, 102(1):614–635.