Standards and Tools in Neuroscience

A summary of the Open Source Brain workshop, September 2019

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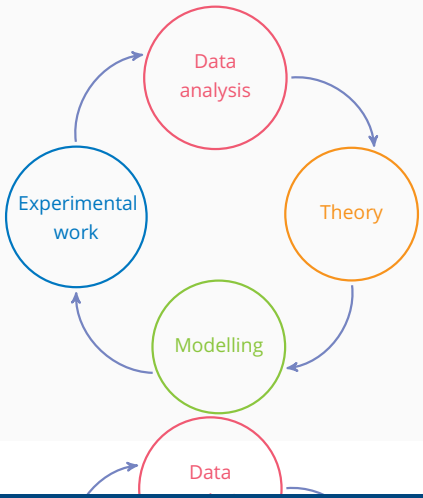
Volunteer: Fedora Project.

Notes

The problem statement

Notes

Neuroscience is complex, and massive



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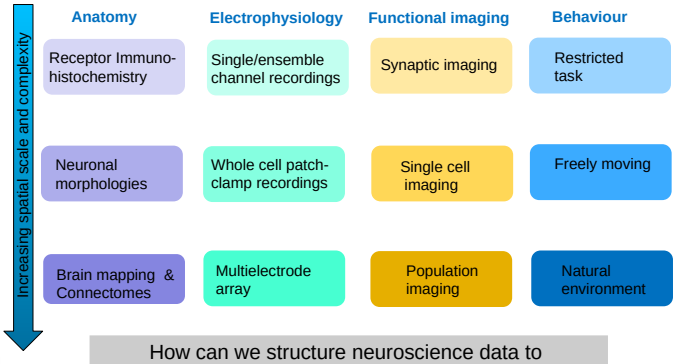
Free/Open Neuroscience

Free/Open science:
Scientific material should be easily, openly accessible to all.

Notes

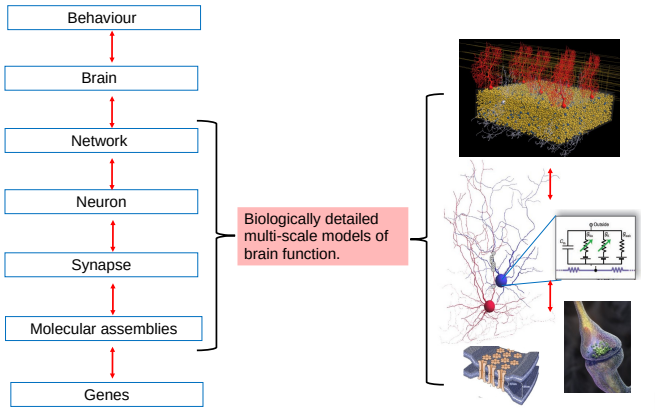
Experimental neuroscience data is heterogeneous, multiscale and analysis is complex



How can we structure neuroscience data to facilitate reuse?

Notes

Models of brain function span multiple spatial scales



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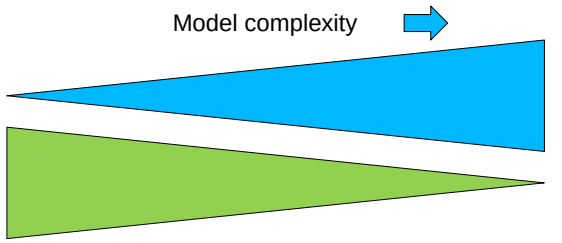
A scaling problem



Transparency, accessibility, reproducibility, reuse…..and utility as a scientific tool

Notes

Standards: the common tongue

Notes

NWB:N 2.0: An Ecosystem for Neurophysiology Data Standardization

Oliver Rübel
Computational Research Division, Lawrence Berkeley National Laboratory

Open Source Brain Workshop
Alghero, Sardinia
September 10, 2019

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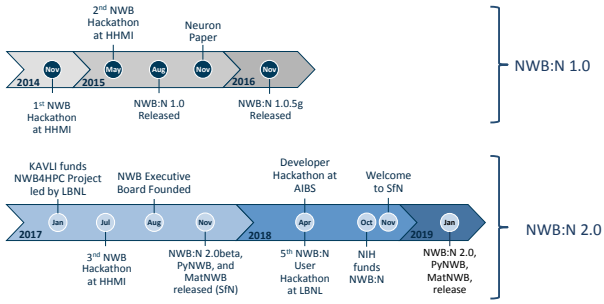
Notes

A brief history of NWB:N



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Overview

Motivation: Lack of standards for neurophysiology data and related metadata is the single greatest impediment to fully extracting return-on-investment from neurophysiology experiments, impeding interchange and reuse of data and reproduction of derived conclusions

NWB:N – An Ecosystem for Neuroscience Data Standardization

- The NWB:N data standard defines a unified data format for neurophysiology data, focused on the dynamics of groups of neurons measured under a large range of experimental conditions
- NWB:N is more than just a file format but it defines an ecosystem of tools, methods, and standards for storing, sharing, and analyzing complex neurophysiology data

Goal: With NWB:N we aim to develop a next generation data format and software ecosystem that will enable standardization, sharing, and reuse of neurophysiology data and analyses, enhancing discovery and reproducibility

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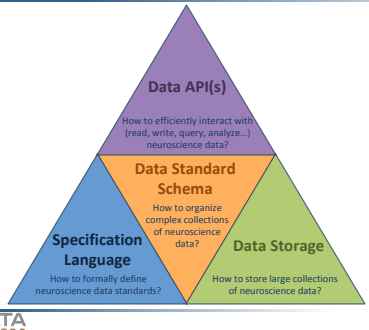
Main components of the NWB:N ecosystem



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Specification language

Goal: Enable the formal definition and programmatic interpretation of neuroscience data standards

Format specification language: Schema for defining hierarchical data schemas

Main primitives of the specification language:

- **Object primitives:**
  - **Group:** A group is a collection of objects i.e. subgroups, datasets, links
  - **Dataset:** n-dimensional array with associated data type, dimensions, etc.
  - **Attribute:** small metadata dataset defined on a Group or Dataset
  - **Link:** Like a POSIX soft link to given target neurodata_type (specifying a Group or Dataset)
- **Data type specifications**
  - **Basic data types:** strings (ascii, utf8), numeric types (float, int, uint, bool etc.) etc.
  - **Compound data types:** build complex data type, similar to structs
  - **Isodatetime:** ISO8061 datetime string, e.g. 2018-09-28T14:43:54.123+02:00
  - **Object reference:** Like a link to given neurodata_type but stored as values of a dataset (or attribute)
  - **Region reference:** Links to regions (i.e., select subsets) of datasets stored as as values of a dataset (or attribute)
- **Namespace specification:**
  - Used to define a namespace for format specifications
  - Needed to define and avoid collisions between extensions and to enable the creation of new data standards

Notes

Data storage

Primary function: Map NWB:N primitives (Groups, Datasets, Attributes etc.) to storage

NWB:N uses HDF5 as its main file storage backend:

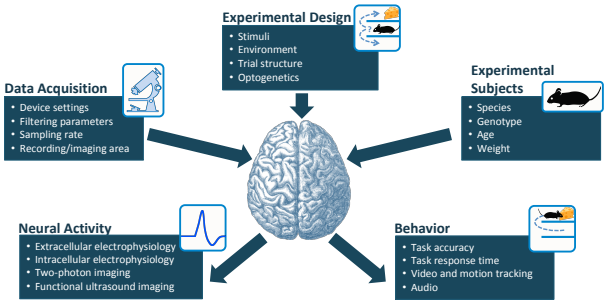
- Supports large-scale storage of complex data collections in a single file
- Optimized for performance (parallel I/O, advanced I/O filters etc.)
- Supported across platforms and programming languages (Matlab, Python, C/C++, Fortran, R…)
- Targets long-term support

NWB:N supports advanced I/O features:

- Lazy data load ➔ Fast file open and efficient memory usage even for very large files
- Chunking ➔ Optimize data layout, storage, and I/O
- I/O filter ➔ E.g, use compression to reduce storage cost
- Self-contained data storage ➔ Store all data in a single file (e.g. for sharing)
- Modular data storage ➔ Store data across multiple files and integrated via external links
- Iterative data write ➔ Support data streaming, reduce memory usage, and optimize I/O

Notes

NWB:N supports complex collections of data required for understanding the brain



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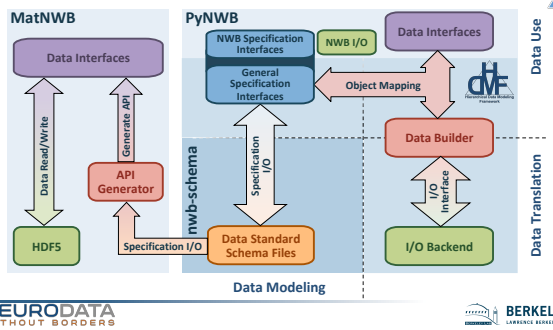
Advanced software architecture for data standardization



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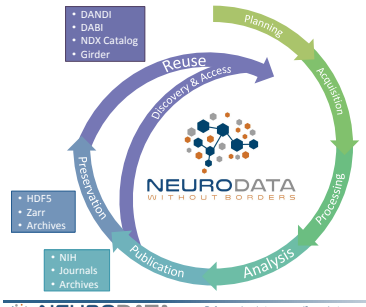
Neurophysiology Tools

NWB:N technologies at the heart of the neurodata lifecycle and applications



Electrophysiology	Optical Physiology
DataJoint, NWB Tutorials	
OpenEphys, Plexon, Neuralynx, Intan, TDT, SpikeGadgets, SpikeGLX	MiniScope ScanImage
PyNWB, MatNWB, HDMF, HDF5 tools/libs	
SpikeInterfaces MountainSort, KiloSort, npsorting BrainStorm WaveClust, UltraMegaSort2000, KiloSort Kluster	NoRMCorre CNMF-E CELLMax EXTRACT
NWB-Explorer (OpenSourceBrain) NWB-JupyterWidgets	
BrainStorm, RAVE, Neo, SpikeWidgets, EcogVis, ephys-viz, …	CaImAn

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Reference herein to any specific product, process, or service does not constitute or imply its endorsement, recommendation, or favoring by NWB or LBNL.

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Notes

Community engagement and outreach

Science Engagement

- **NWB User/Developer Days:**
  - 65+ scientists from 20 major institutions attended in 2018 at LBNL and AIBS
  - 43 users and developers from 29 major labs and research institutions in 2019 at HHMI
- The NIH proposal included 55 letters of support
- Many groups are already adopting NWB:N, e.g., FrankLab, ChangLab (UCSF), BouchardLab (LBNL), SvobodaLab (HHMI), MeisterLab (CalTech), Redwood (UCB), Allen Institute for Brain Science, among others.
- Broad outreach to tool builders:
- Broad outreach to foster publication of data:
- We have reached out to and are working with NIH BRAIN Initiative U19 projects
- KAVLI seed grants
- Simons data pilots

Industry Engagement

- Kitware (Visualization/CI)
- Vidrio (MatNWB)
- MathWorks (MATLAB)
- Vathes (DataJoint)

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Governance

- NWB Executive Board
- NWB Technical Advisory Board
- KAVLI foundation
- NIH BRAIN Initiative

Community Resources

Open Source

GitHub
https://github.com/NeurodataWithoutBorders

Open Documentation

Read the Docs
https://neurodatawithoutborders.github.io

Developer/User Channels

slack
https://nwb-users.slack.com

Google Groups
https://groups.google.com/forum/#!forum/neurodatawithoutborders

User Engagement

- User- and developer hackathons
- Outreach at neuroscience conferences (CoSyNe, SfN, BRAIN Initiative Investigators Meeting)
- Community discussion and surveys

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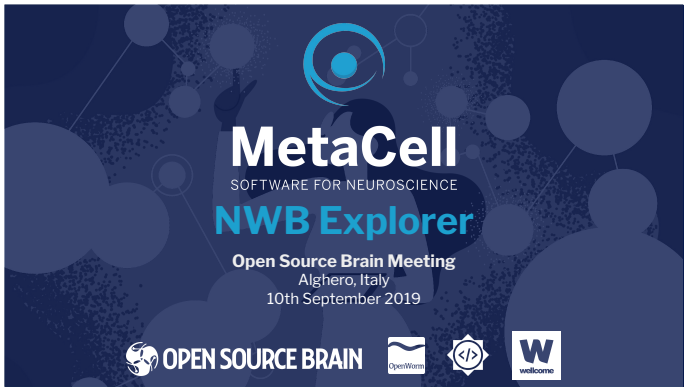

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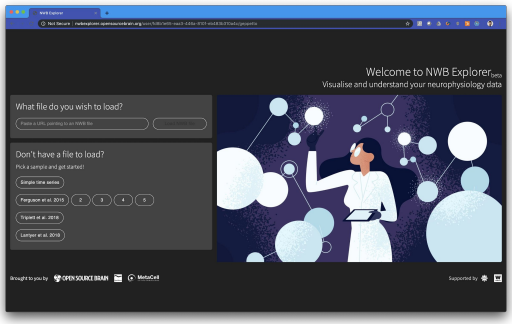


Notes

NWB EXPLORER
DEMO

Try it out at

https://nwbexplorer.opensourcebrain.org

Notes

Creating cortical models across scales in NeuroML

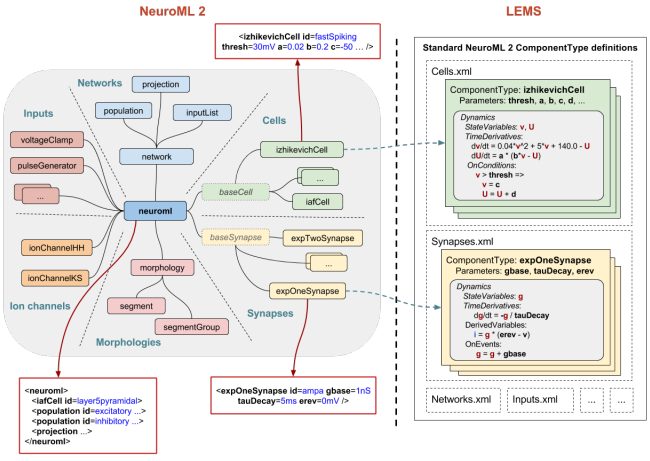
Padraig Gleeson
p.gleeson@ucl.ac.uk
University College London

Open Source Brain workshop 2019

11th Sept 2019

OPEN SOURCE BRAIN NeuroML

Notes



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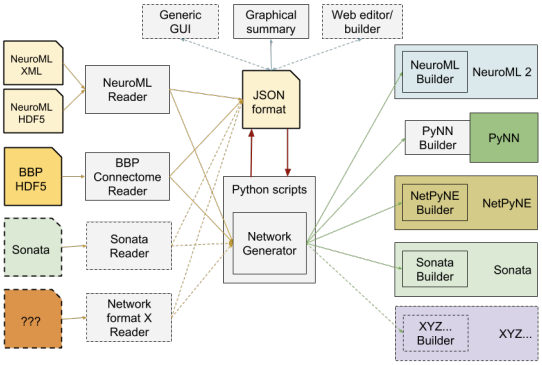

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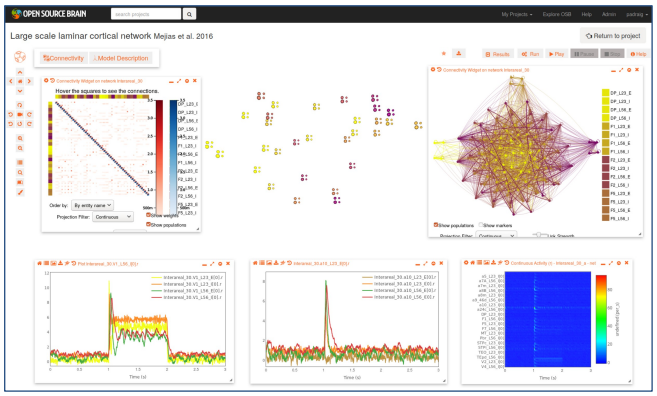
OPEN SOURCE BRAIN NeuroML



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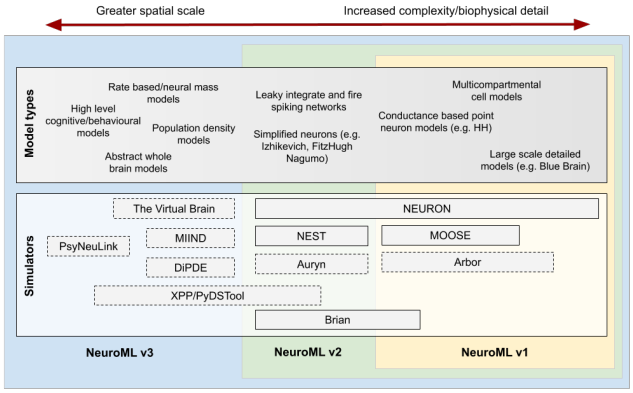

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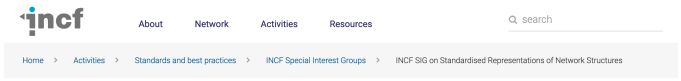
OPEN SOURCE BRAIN NeuroML



INCF SIG on Standardised Representations of Network Structures

This SIG deals with the various tools and formats for creating and sharing representations of biological neuronal networks, and will work towards ensuring these are as interoperable and usable as possible for computational neuroscientists.

Contact info: p.gleeson@uc.ac.uk

Members

Anton Arkhipov, Allen Institute, USA
Tom Close, Monash University, Australia
Sharon Crook, Arizona State University, USA
Kael Dai, Allen Institute, USA
Andrew Davison, UNIC, CNRS, France
Lia Domide, Codemart, Romania & Aix-Marseille Université, France
Salvador Durá-Bernal, SUNY Downstate Medical Center, USA
Viktor Jirsa, Aix-Marseille Université, France
Padraig Gleeson, University College London, UK

Notes

Converting simulator specific formats to NeuroML2

Open Source Brain Meeting 2019

Boris Marin

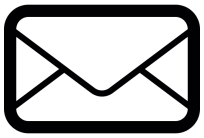
boris.marin@ufabc.edu.br

Universidade Federal do ABC

Notes

Converting NMODL to NeuroML

The Simple™, OSB sponsored way of converting models to NeuroML2



mailto:p.gleeson@ucl.ac.uk

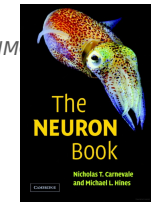
Notes

The *NEURON* simulator

Defining models in *NEURON*

https://www.neuron.yale.edu/neuron/

- Cells, Networks: *hoc* language (accessible from Python)
  - morphologies
  - synaptic connections
  - *.hoc* files
- Ion Channels (membrane mechanisms): *NM*
  - *.mod* files

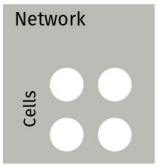


Notes

What is NeuroML, and why should I care?

Why can OSB process any NeuroML2 file?

- NML is *structured* (not unlike a *Type System*)

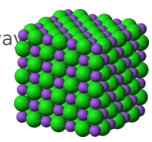


Notes

It is all about **Structure**

Structure in NeuroML / NMODL

- A *Type System* (composability rules) is what grants NML its superpowers
- `nmodl` is also powerful, but can be used as a general purpose language
  - `VERBATIM` blocks
  - many different ways of achieving same goal
  - prone to *unstructuredness*
- OSB could in theory treat `nmodl` the same wa NML...
  - .... if only people stuck to "good practices"!



Notes

Levels of Abstraction

NeuroML2

```
<ionChannelHH id="kChan" conductance="10pS" species="k">

    <gateHHrates id="n" instances="4">
      <forwardRate type="HHExpLinearRate" rate="0.1per_ms" midpoint="-55mV" scale="10mV"/>
      <reverseRate type="HHExpRate" rate="0.125per_ms" midpoint="-65mV" scale="-80mV"/>
    </gateHHrates>

  </ionChannelHH>
```

NMODL

```
BREAKPOINT {
        SOLVE states METHOD cnexp
        gk = gkbar * n ^ 4
        ik = gk * (v-ek)
}
INITIAL{
        n = alpha(v) / (alpha(v) + beta(v))
}
DERIVATIVE states{
        n' = (1 - n) * alpha(v) - n * beta(v)
}
```

```
FUNCTION alpha(Vm(mV))(/ms){
        LOCAL x
        UNITSOFF
        x = (Vm + 55) / 10
        if(fabs(x) > 1e-6){
                alpha=0.1*x/(1-exp(-x))
        }else{
                alpha=0.1/(1-0.5*x)
        }
        UNITSON
}
```

Notes

Levels of Abstraction

Declarative vs Imperative

- NeuroML2 operates (at least syntactically) closer to the level of abstraction employed by electrophysiologists
- The gory details exist, but elsewhere: *LEMS*
  - i.e. what to do with α, β; the definition of an `ExpRate`; how all of that is converted to conductances/currents...
- But we seldom need (want!) to interact with that level (look under the hood)

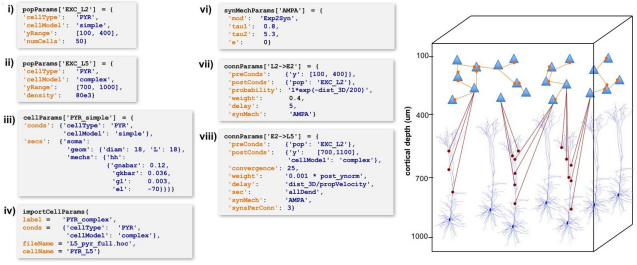
```
<Network ...>
    <Cell ...>
        <Channel ...>
            <Gate ...>
                <Rate ...>
```

```
SOLVE{...} METHOD euler
...
DERIVATIVE {...}
...
FUNCTION trap(v){...}
```

Notes

Levels of Abstraction

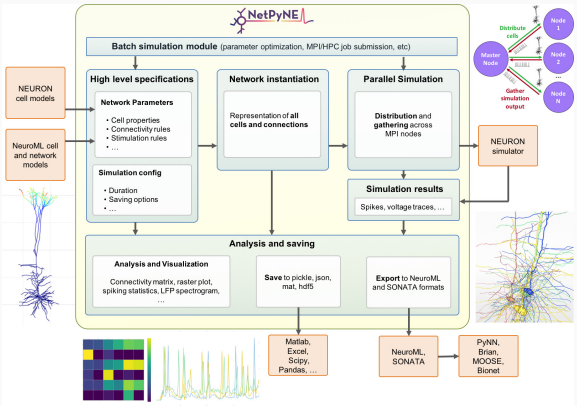
NetPyNE: structured network specification



Dura-Bernal, Salvador, et al. "NetPyNE, a tool for data-driven multiscale modeling of brain circuits." Elife 8

Notes

Netpyne workflow



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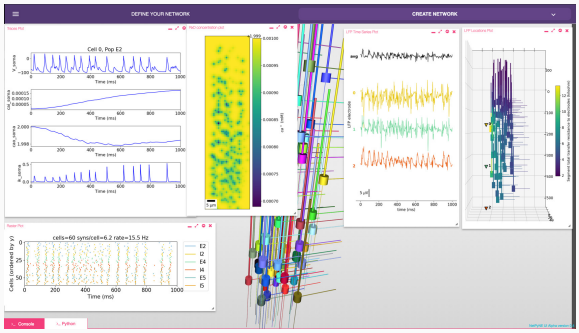
Netpyne GUI



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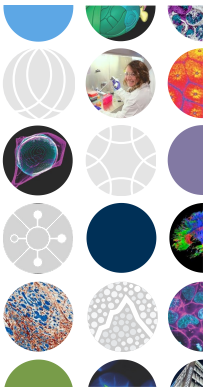
ALLEN INSTITUTE

Large-scale Datasets and Modeling Tools from the Allen Institute for Brain Science

Yazan N. Billeh
yazanb@alleninstitute.org



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ALLEN INSTITUTE | ALLEN INSTITUTE for BRAIN SCIENCE | ALLEN INSTITUTE for CELL SCIENCE | ALLEN INSTITUTE for IMMUNOLOGY | PAUL G. ALLEN FRONTIERS GROUP

- Established 2003 by Paul G. Allen
- South Lake Union, Seattle, WA
- 500 employees++

hard problems
complexity
foundational biology

→

big science

team science

open science

→

data
knowledge
tools

Notes

CORE PRINCIPLES

Team Science

Interdisciplinary teams working towards common goal

Big Science

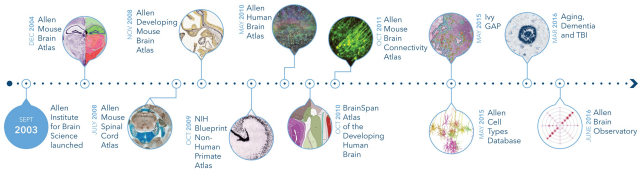
Large-scale projects with robust, massive data

Open Science

All resources available online at brain-map.org or allencell.org

Notes

Allen Institute - Online Public Resources
www.brain-map.org



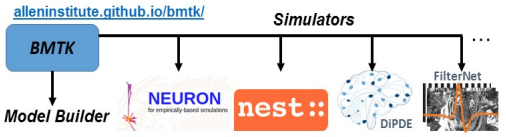
All data are:

- publicly accessible via API as soon as they pass QC
- freely available without any commercial restrictions

Notes

Our Models and Modeling Software Are Freely Available to the Community

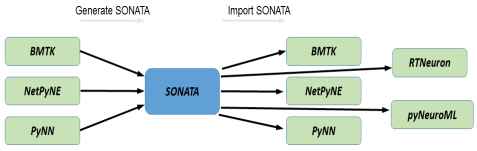
Brain Modeling ToolKit (BMTK): https://alleninstitute.github.io/bmtk/



Notes

Our Models and Modeling Software Are Freely Available to the Community

Scalable Open Network Architecture TemplAte (SONATA):
https://github.com/AllenInstitute/sonata
An interface between SONATA and the NWB format has been developed as well



Notes

Human Brain Project

How model standardization enables new tools and applications in neuroscientific research

Insights from the HBP

Yann Zerlaut
Neuroinformatics team / group of A. Davison
Centre National de la Recherche Scientifique, France

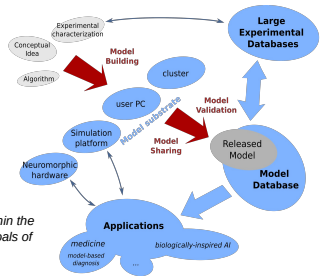
UNIC

Open Source Brain Meeting 2019, Alghero

cnrs

Notes

Motivation



Model production pipeline within the infrastructure and research goals of the Human Brain Project

Notes

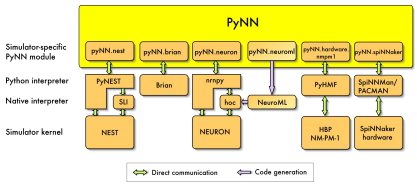
PyNN A unified interface for neuronal network simulators

Simulator-independent environments for developing neuroscience models:

- keep the advantages of having multiple simulators or hardware devices
- but remove the translation barrier.

Three (complementary) approaches:

- GUI (e.g. neuroConstruct)
- XML-based language (e.g. NeuroML, NineML)
- interpreted language (e.g. Python)



Notes

Sonata (pyNN support)

Large-scale simulation of biophysically-detailed neuronal circuits
→ sets specific constraints

the SONATA Data Format emerges as the standard optimized for performance for simulation, analysis and visualization of large-scale circuits
(joint initiative of Blue Brain Project and the Allen Institute for Brain Science)

Export to Sonata format

```
from pyNN.network import Network
from pyNN.serialization import export_to_sonata

sim.setup()
...
# create populations, projections, etc.
...

# add populations and projections to a Network
net = Network(pop1, pop2, ..., prj1, prj2, ...)

export_to_sonata(net, "sonata_output_dir")
```

Import from Sonata format

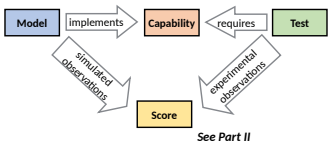
```
from pyNN.serialization import import_from_sonata, load_sonata_simulation_plan
import pyNN.neuron as sim

simulation_plan = load_sonata_simulation_plan("simulation_config.json")
simulation_plan.setup(sim)
net = import_from_sonata("circuit_config.json", sim)
simulation_plan.execute(net)
```

Notes

SciUnit *https://github.com/scidash/sciunit*

Include a validation framework in model development



See Part II

- ✓ **What is SciUnit?**
  *A **Test**-driven framework for formally validating scientific models against data.*
  *It employs the concept of **Capabilities**.*
- ✓ **What are Tests?**
  A procedure intended to establish the quality, performance, or reliability of a model
- ✓ **What are Capabilities?**
  - *interfaces through which the model and the validation framework communicate*
  - *implemented as methods (functions) within the model*

Requires the participation of modellers:

- ✓ support to wrap your models for SciUnit
- ✓ add/request new tests to the library
- ✓ critique existing tests
- ✓ suggest new features

Notes

Test Packages

The overall test suite has been divided into a number of components, some containing validation tests specific to particular brain regions, others more generic. All validation tests are written in Python, using the SciUnit framework. Some of these are listed below:

Test suites for specific brain regions

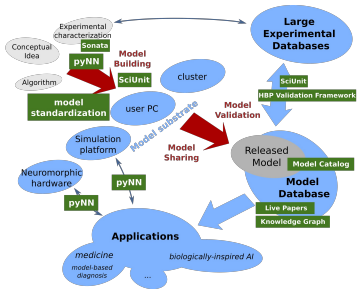
- ***HippoUnit***: `https://github.com/KaliLab/hippounit`
- ***HippoNetworkUnit***: `https://github.com/pedroernesto/HippoNetworkUnit`
- ***CerebUnit***: `https://github.com/lungsi/cerebellum-unit`
- ***BasalUnit***: `https://github.com/appukuttan-shailesh/basalunit`

Test suites for model features, independent of cell type or brain region

- ***MorphoUnit***: `https://github.com/appukuttan-shailesh/morphounit`
- ***NetworkUnit***: `https://github.com/mvonpapen/simrest_validation`
- ***eFELUnit***: `https://github.com/appukuttan-shailesh/eFELunit`

Notes

Summary

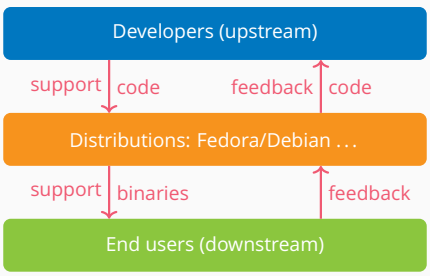


Notes

NeuroFedora: marketing pitch

Notes

Liaison between developers and users



Notes

Search: “NeuroFedora”

“Live” ISO now ready to download (demo)
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