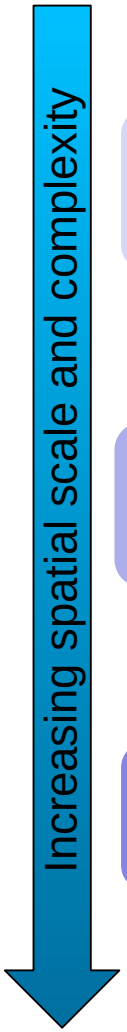
1. A simplified diagram. Actually a lot more complex
2. It is so massive that you can speak to another neuroscientist and not understand a word of what they say—we specialise
3. We won’t even discuss dissemination to a non scientific audience today—a completely different problem.

Experimental neuroscience data is heterogeneous, multiscale and analysis is complex



Anatomy	Electrophysiology	Functional imaging	Behaviour
Receptor Immuno-histochemistry	Single/ensemble channel recordings	Synaptic imaging	Restricted task
Neuronal morphologies	Whole cell patch-clamp recordings	Single cell imaging	Freely moving
Brain mapping & Connectomes	Multielectrode array	Population imaging	Natural environment

How can we structure neuroscience data to facilitate reuse?

Models of brain function span multiple spatial scales

Behaviour

Brain

Network

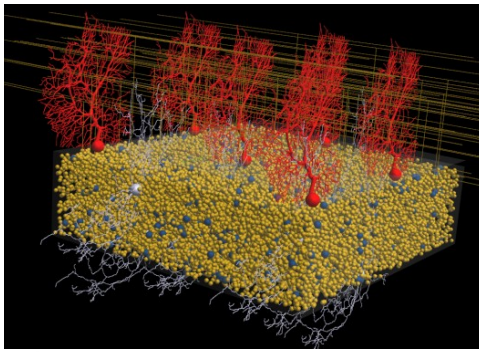
Neuron

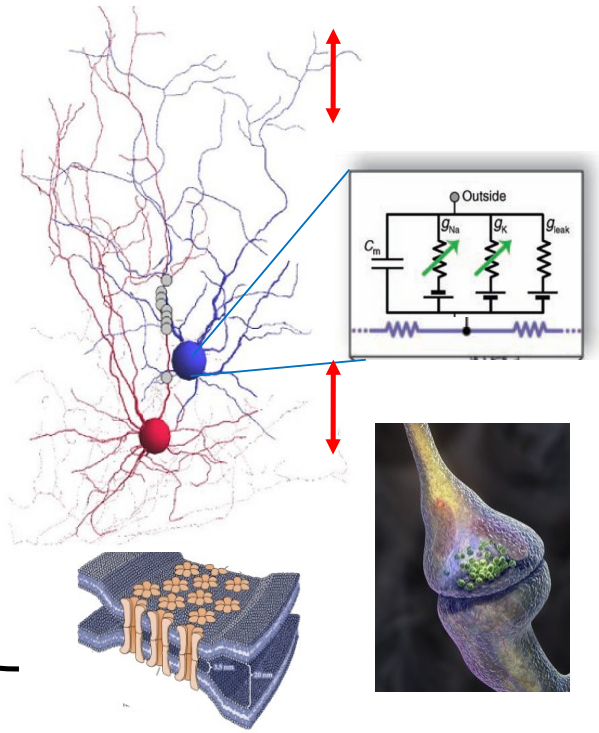
Synapse

Molecular assemblies

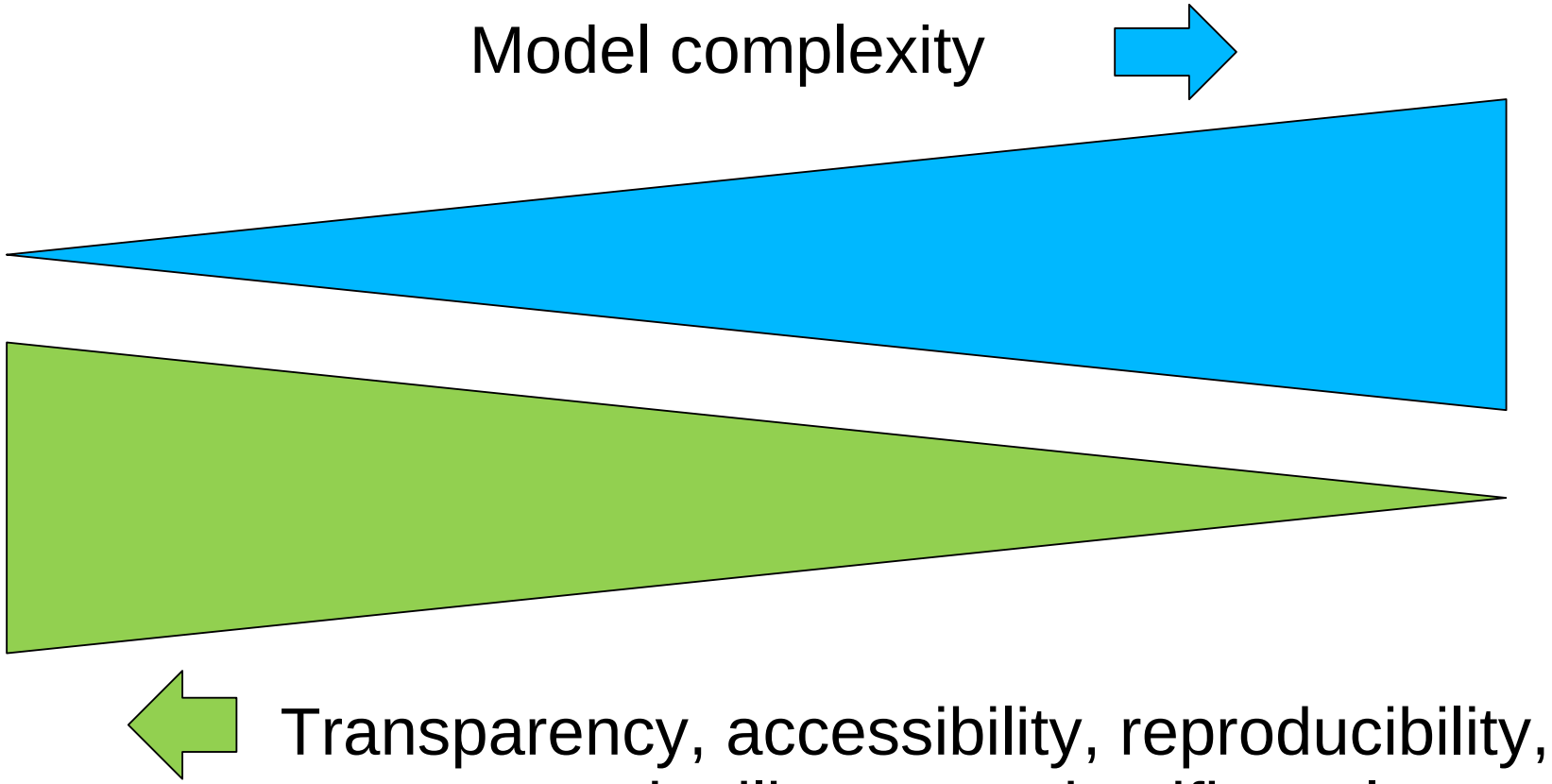
Genes

Biologically detailed multi-scale models of brain function.





A scaling problem



NWB:N 2.0: An Ecosystem for Neurophysiology Data Standardization

Oliver Rübel

Computational Research Division, Lawrence Berkeley National Laboratory

Open Source Brain Workshop

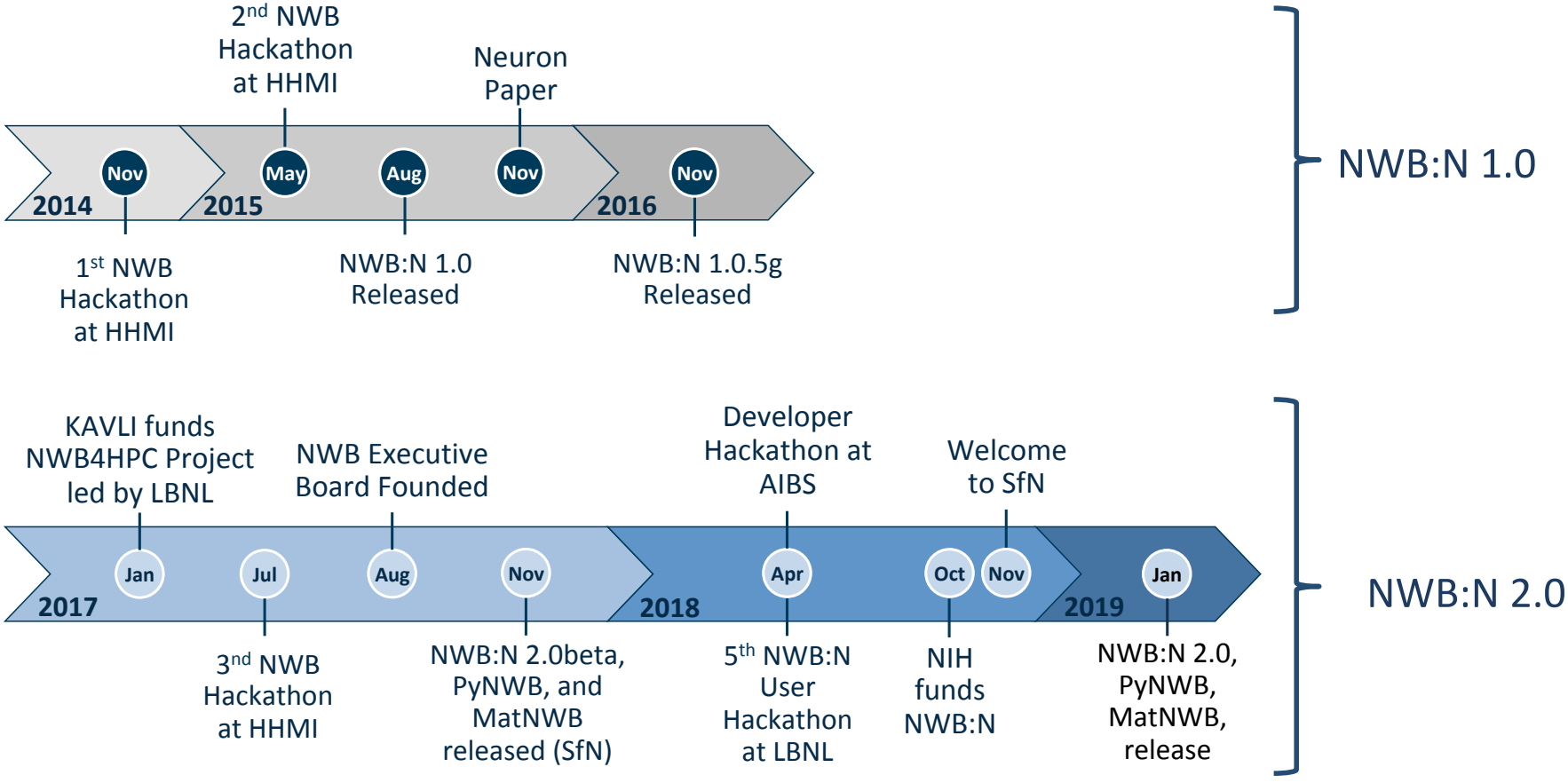
Alghero, Sardinia

September 10, 2019

NEURODATA WITHOUT BORDERS

BERKELEY LAB
LAWRENCE BERKELEY NATIONAL LABORATORY

A brief history of NWB:N



Overview

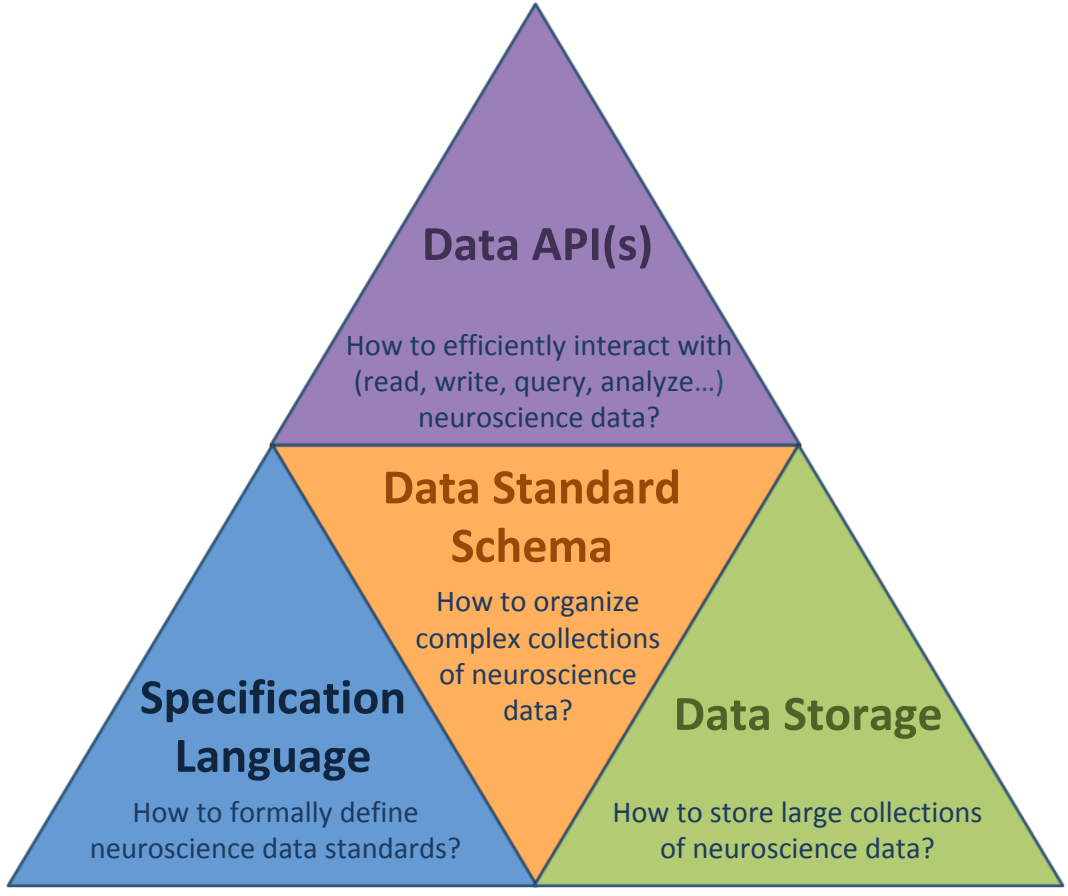
Motivation: Lack of standards for neurophysiology data and related metadata is the single greatest impediment to fully extracting return-on-investment from neurophysiology experiments, impeding interchange and reuse of data and reproduction of derived conclusions

NWB:N – An Ecosystem for Neuroscience Data Standardization

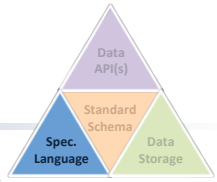
- The NWB:N data standard defines a unified data format for neurophysiology data, focused on the dynamics of groups of neurons measured under a large range of experimental conditions
- NWB:N is more than just a file format but it defines an ecosystem of tools, methods, and standards for storing, sharing,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

Main components of the NWB:N ecosystem



Specification language



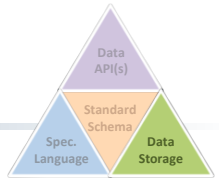
Goal: Enable the formal definition and programmatic interpretation of neuroscience data standards

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

Data storage



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

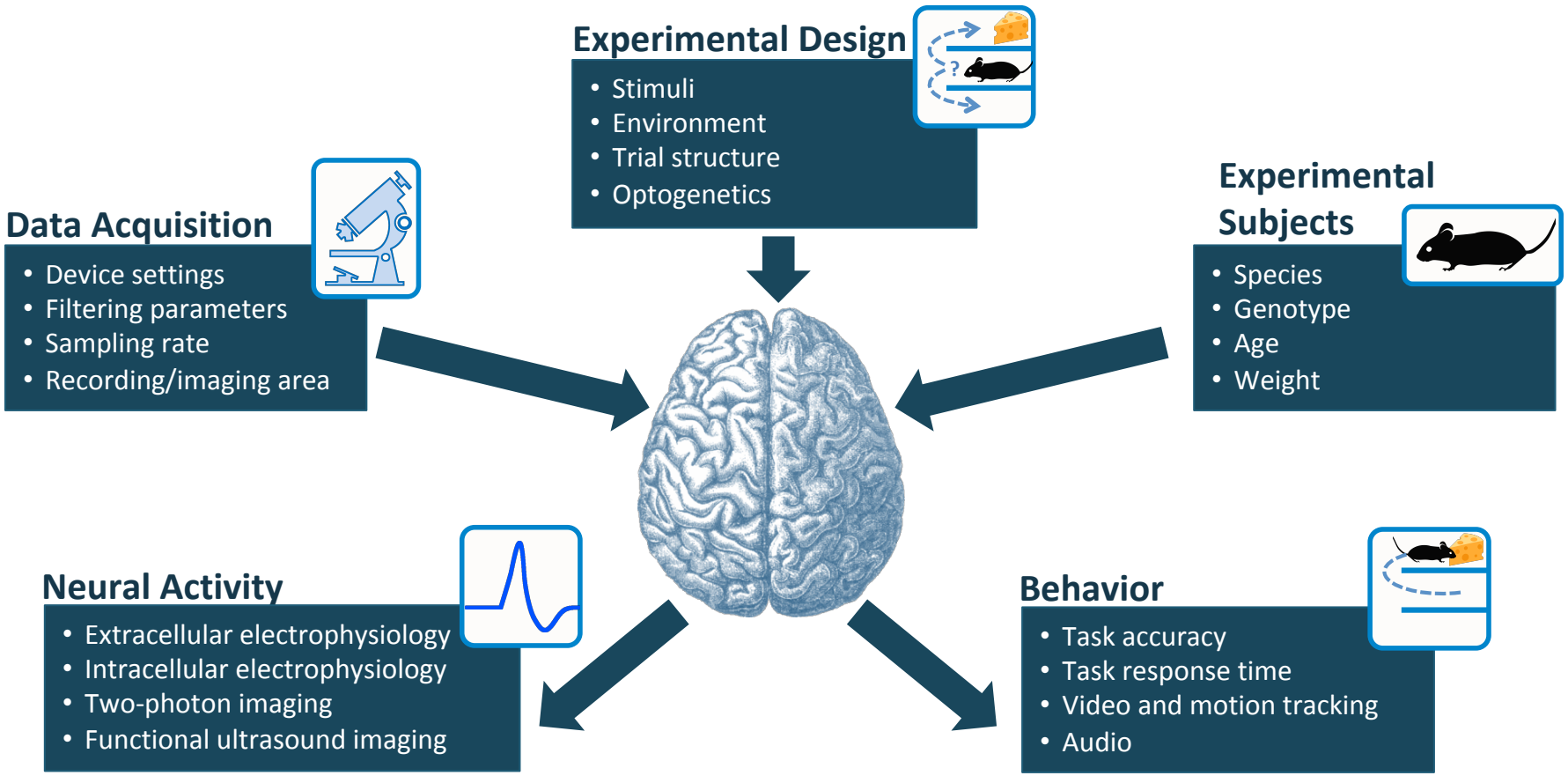
NWB:N uses HDF5 as its main file storage backend:

- 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

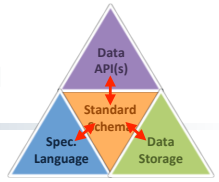
NWB:N supports advanced I/O features:

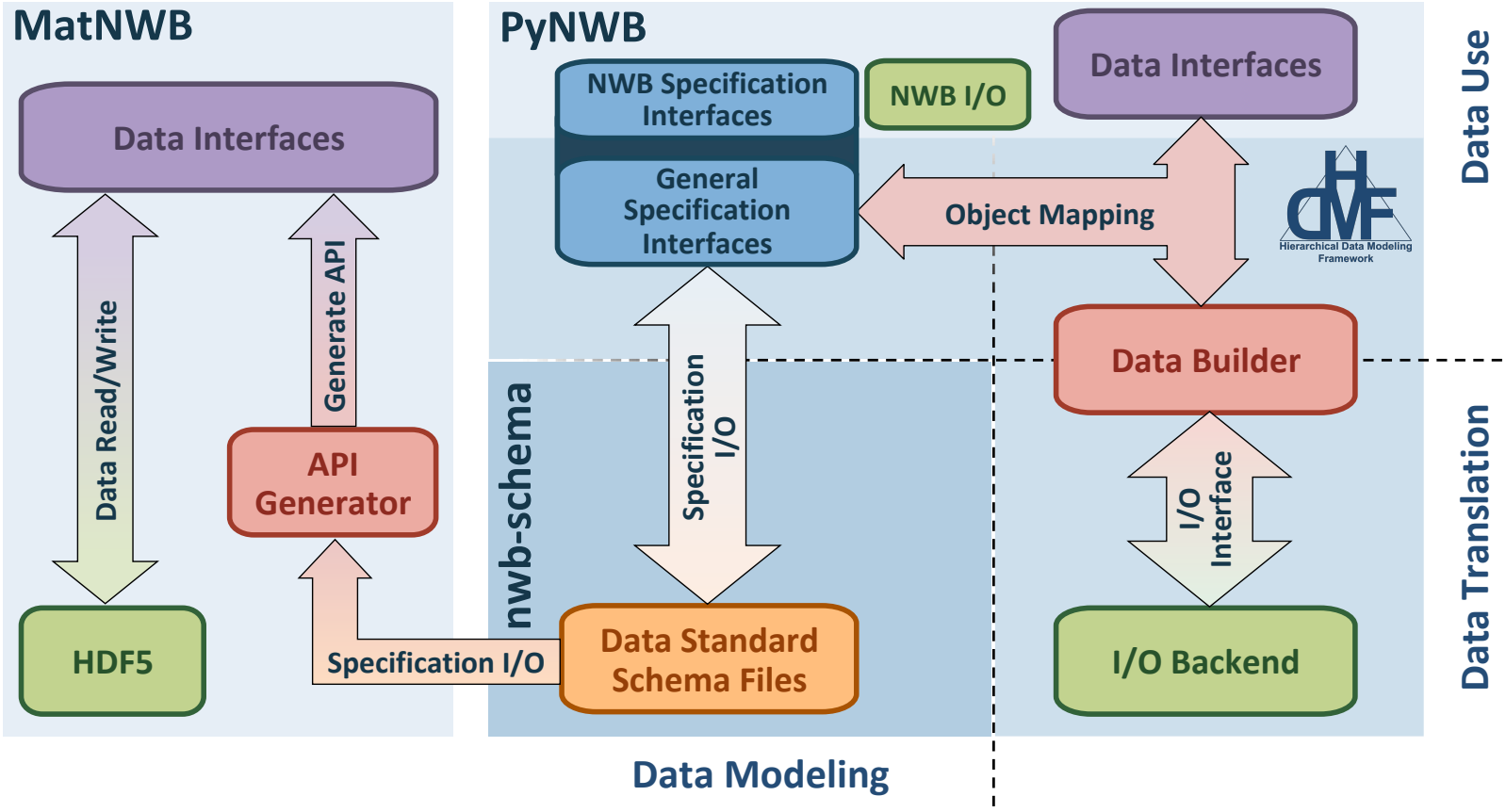
- Lazy data load ➔ Fast file open and efficient memory usage even for very large files
- Chunking ➔ Optimize data layout, storage, and I/O
- I/O filter ➔ E.g, use compression to reduce storage cost
- Self-contained data storage ➔ Store all data in a single file (e.g. for sharing)
- Modular data storage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

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



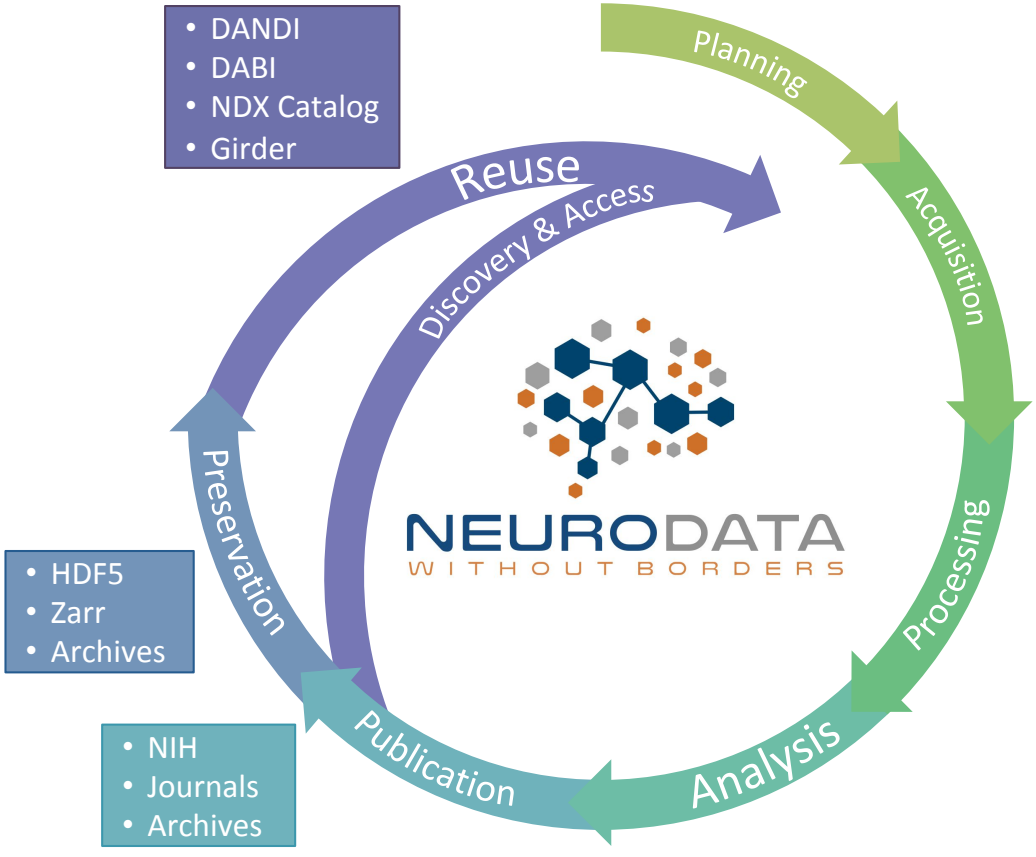
Advanced software architecture for data standardization





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, npzsorting BrainStorm WaveClust, UltraMegaSort2000, KiloSort Kluster	NoRMCorre CNMF-E CELLMax EXTRACT
NWB-Explorer (OpenSourceBrain) NWB-JupyterWidgets	
BrainStorm, RAVE, Neo, SpikeWidgets, EcogVis, ephys-viz, …	CaImAn

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)

NEURODATA WITHOUT BORDERS

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

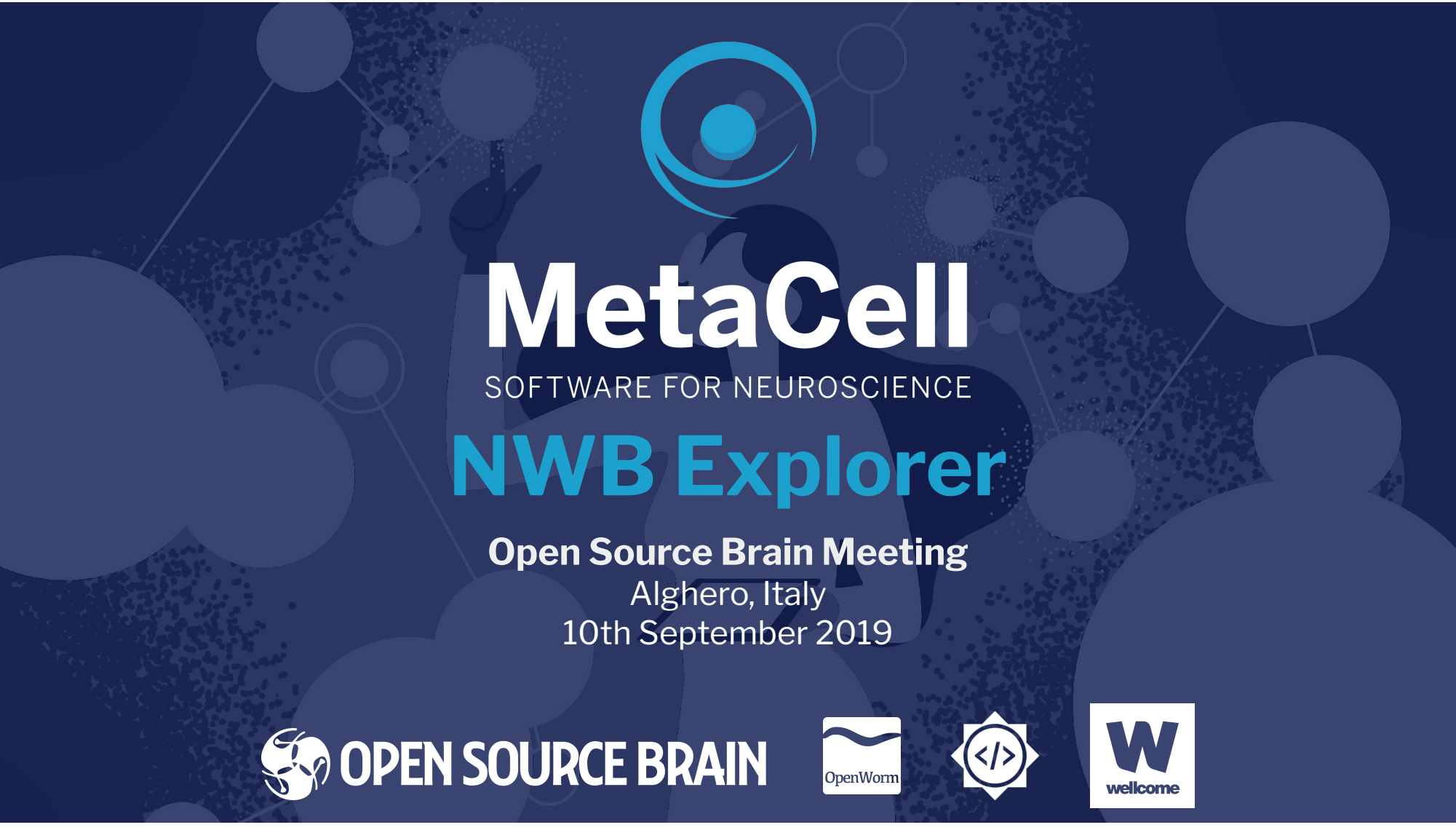
https://nwb-users.slack.com

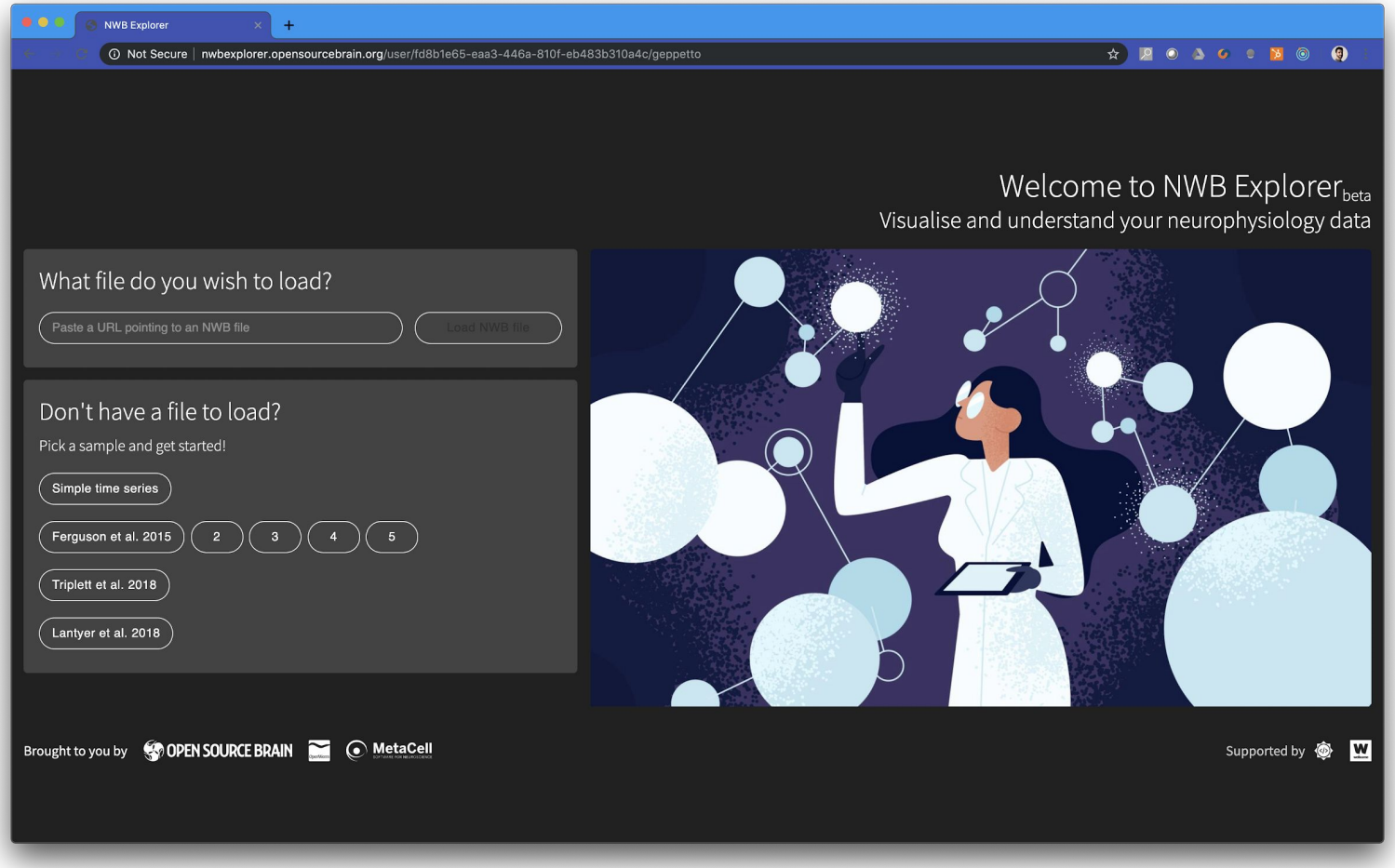
Google Groups

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


MetaCell
SOFTWARE FOR NEUROSCIENCE
NWB Explorer
Open Source Brain Meeting
Alghero, Italy
10th September 2019
OPEN SOURCE BRAIN
OpenWorm
wellcome



NWB EXPLORER

DEMO

Try it out at

https://nwbexplorer.opensourcebrain.org

Creating cortical models across scales in NeuroML

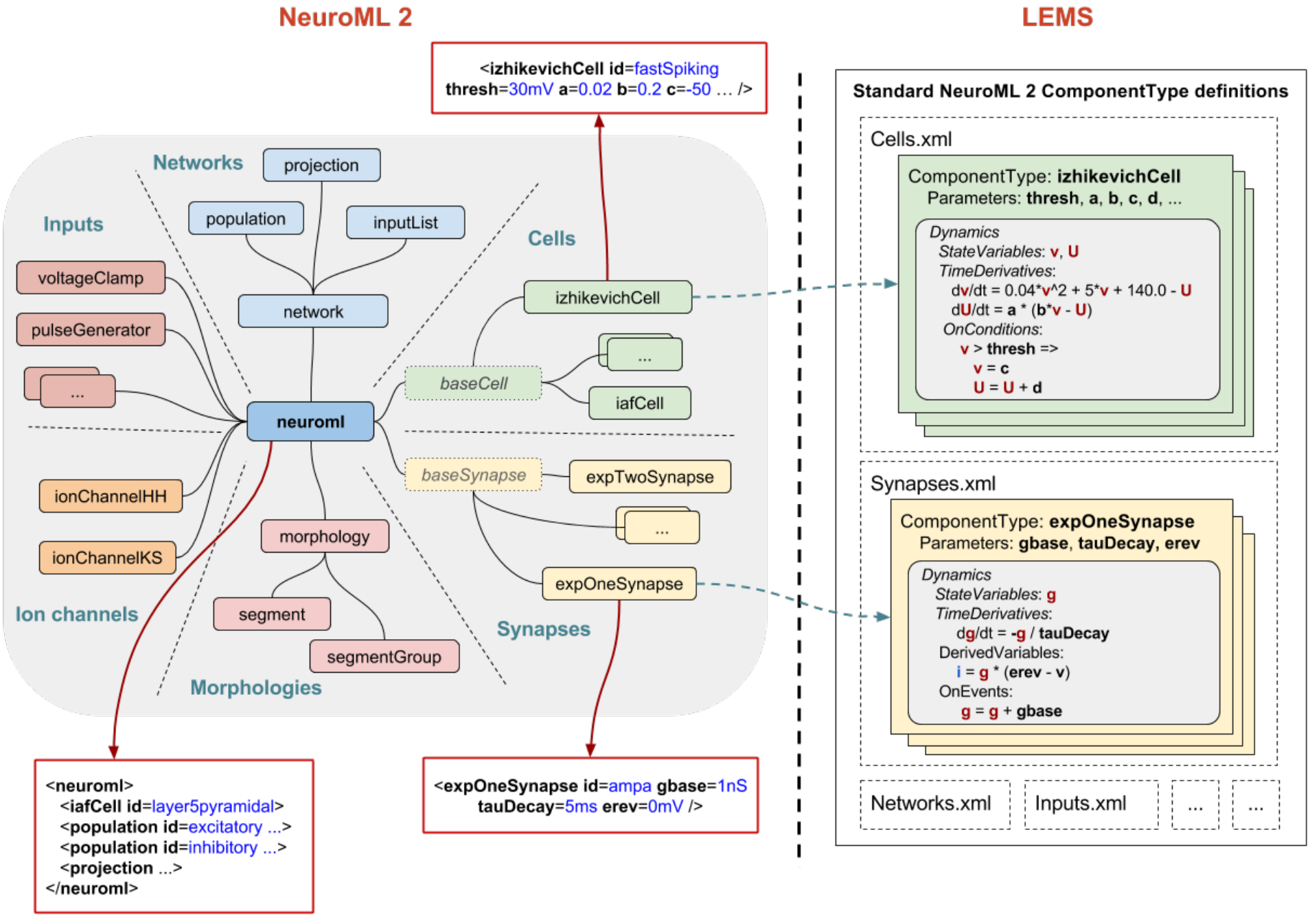
Padraig Gleeson
p.gleeson@ucl.ac.uk
University College London

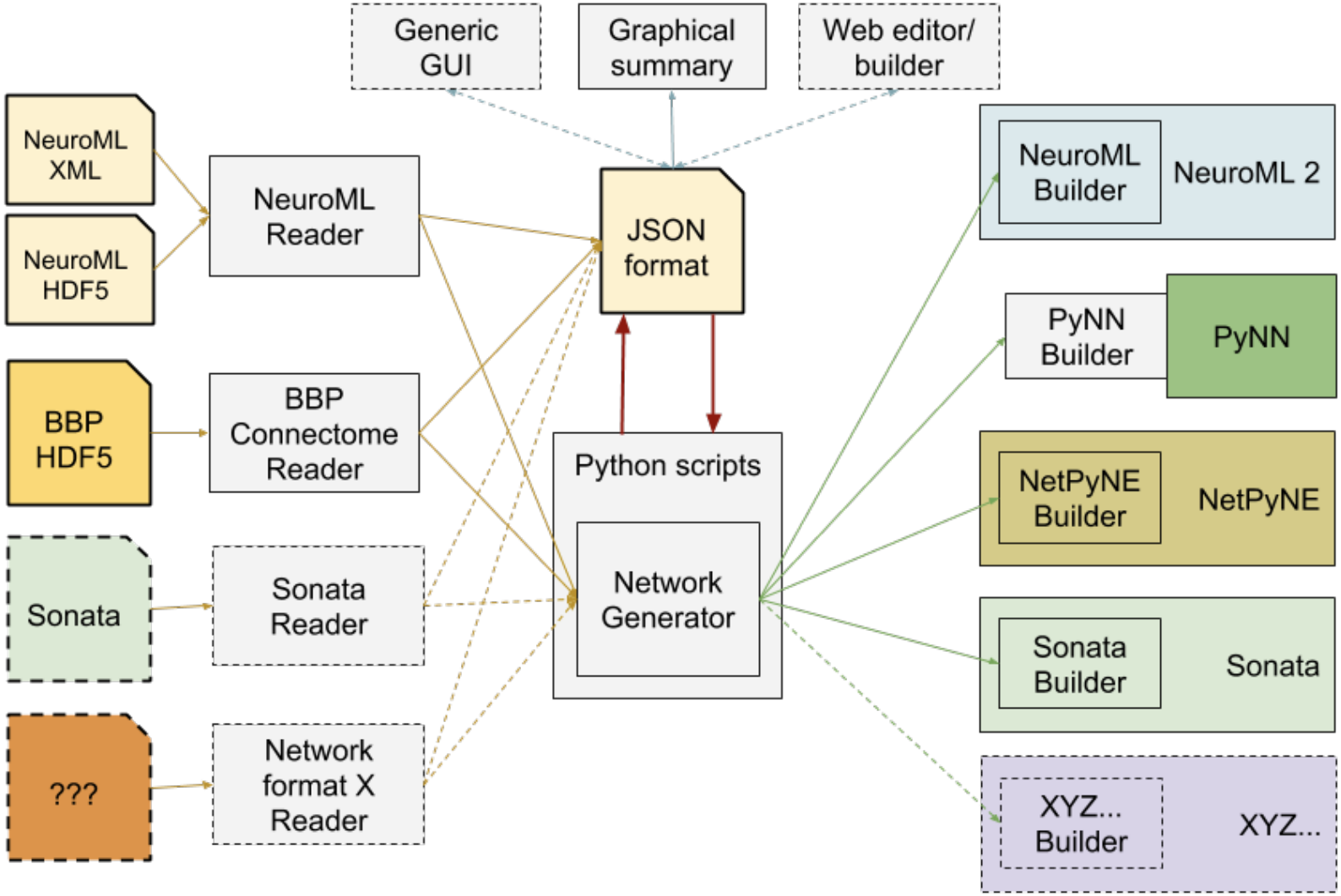
Open Source Brain workshop 2019

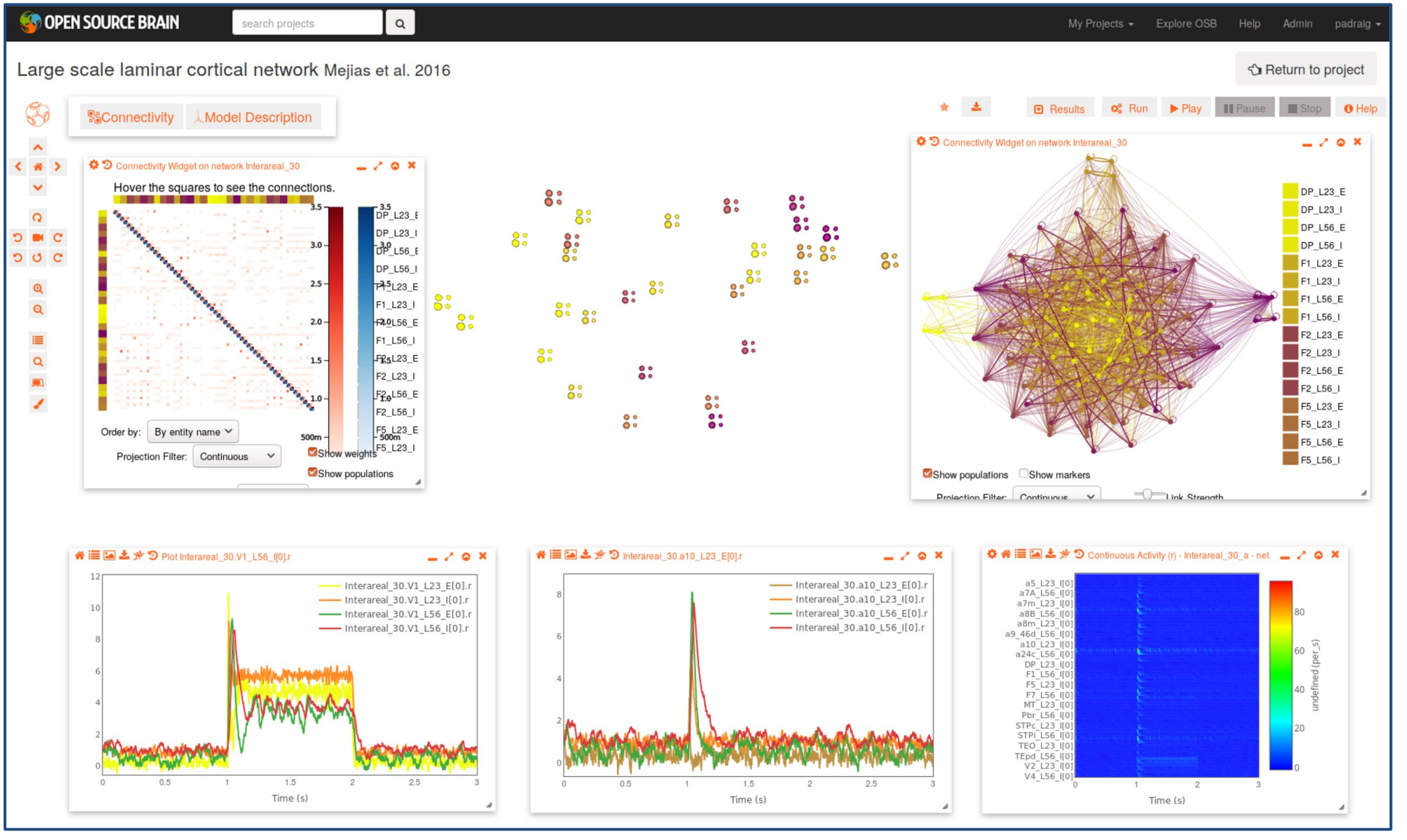
11th Sept 2019

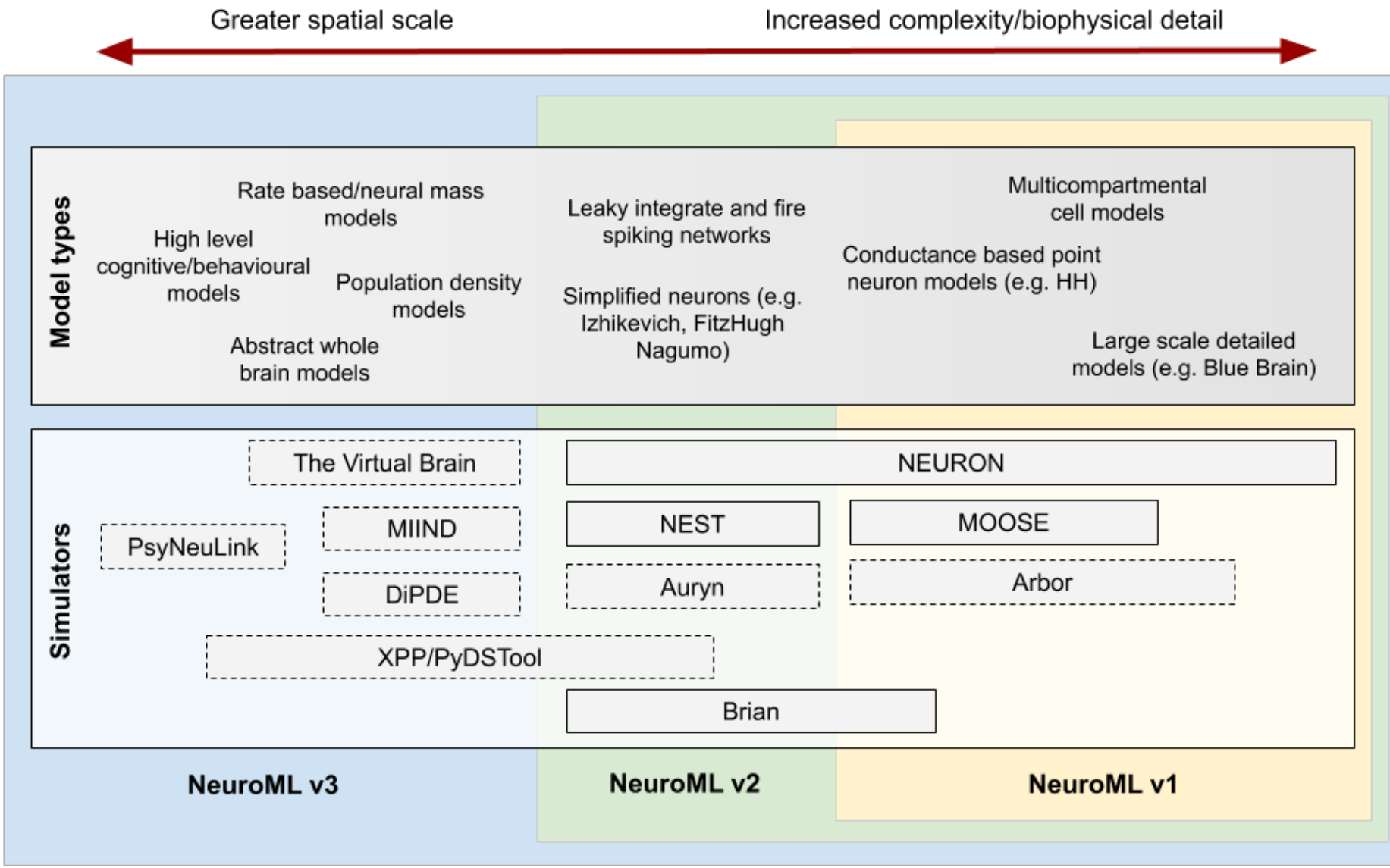
OPEN SOURCE BRAIN [NeuroML]

wellcome


NeuroML 2
LEMS
<izhikevichCell id=fastSpiking thresh=30mV a=0.02 b=0.2 c=-50 … />
Networks
projection
population
inputList
Inputs
Cells
voltageClamp
network
izhikevichCell
pulseGenerator
...
baseCell
...
iafCell
neuroml
baseSynapse
expTwoSynapse
ionChannelHH
morphology
...
ionChannelKS
expOneSynapse
Ion channels
segment
Synapses
segmentGroup
Morphologies
<neuroml>
<iafCell id=layer5pyramidal>
<population id=excitatory ...>
<population id=inhibitory ...>
<projection ...>
</neuroml>
<expOneSynapse id=ampa gbase=1nS tauDecay=5ms erev=0mV />
Standard NeuroML 2 ComponentType definitions
Cells.xml
ComponentType: izhikevichCell
Parameters: thresh, a, b, c, d, ...
Dynamics
StateVariables: v, U
TimeDerivatives:
dv/dt = 0.04*v^2 + 5*v + 140.0 - U
dU/dt = a * (b*v - U)
OnConditions:
v > thresh =>
v = c
U = U + d
Synapses.xml
ComponentType: expOneSynapse
Parameters: gbase, tauDecay, erev
Dynamics
StateVariables: g
TimeDerivatives:
dg/dt = -g / tauDecay
DerivedVariables:
i = g * (erev - v)
OnEvents:
g = g + gbase
Networks.xml
Inputs.xml
...
...


Generic GUI
Graphical summary
Web editor/ builder
NeuroML XML
NeuroML HDF5
NeuroML Reader
JSON format
NeuroML Builder
NeuroML 2
PyNN Builder
PyNN
BBP HDF5
BBP Connectome Reader
Python scripts
NetPyNE Builder
NetPyNE
Sonata
Sonata Reader
Network Generator
Sonata Builder
Sonata
???
Network format X Reader
XYZ... Builder
XYZ...


OPEN SOURCE BRAIN
search projects
My Projects
Explore OSB
Help
Admin
padraig
Large scale laminar cortical network Mejias et al. 2016
Return to project
Connectivity
Model Description
Results
Run
Play
Pause
Stop
Help
Connectivity Widget on network Interareal_30
Hover the squares to see the connections.
Order by:
By entity name
Projection Filter:
Continuous
Show weights
Show populations
Show markers
Link Strength
DP_L23_E
DP_L23_I
DP_L56_E
DP_L56_I
F1_L23_E
F1_L23_I
F1_L56_E
F1_L56_I
F2_L23_E
F2_L23_I
F2_L56_E
F2_L56_I
F5_L23_E
F5_L23_I
F5_L56_E
F5_L56_I
Plot Interareal_30.V1_L56_I[0].r
Interareal_30.V1_L23_E[0].r
Interareal_30.V1_L23_I[0].r
Interareal_30.V1_L56_E[0].r
Interareal_30.V1_L56_I[0].r
Time (s)
Interareal_30.a10_L23_E[0].r
Interareal_30.a10_L23_I[0].r
Interareal_30.a10_L56_E[0].r
Interareal_30.a10_L56_I[0].r
Time (s)
Continuous Activity (r) - Interareal_30_a - net
a5_L23_I[0]
a7A_L56_I[0]
a7m_L23_I[0]
a8B_L56_I[0]
a8m_L23_I[0]
a9_46d_L56_I[0]
a10_L23_I[0]
a24c_L56_I[0]
DP_L23_I[0]
F1_L56_I[0]
F5_L23_I[0]
F7_L56_I[0]
MT_L23_I[0]
Pbr_L56_I[0]
STPc_L23_I[0]
STPi_L56_I[0]
TEO_L23_I[0]
TEpd_L56_I[0]
V2_L23_I[0]
V4_L56_I[0]
undefined (per_s)
Time (s)


Greater spatial scale
Increased complexity/biophysical detail
Model types
Rate based/neural mass models
High level cognitive/behavioural models
Population density models
Abstract whole brain models
Leaky integrate and fire spiking networks
Simplified neurons (e.g. Izhikevich, FitzHugh Nagumo)
Multicompartmental cell models
Conductance based point neuron models (e.g. HH)
Large scale detailed models (e.g. Blue Brain)
Simulators
The Virtual Brain
PsyNeuLink
MIIND
DiPDE
XPP/PyDSTool
NEURON
NEST
MOOSE
Auryn
Arbor
Brian
NeuroML v3
NeuroML v2
NeuroML v1

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

Converting simulator specific formats to NeuroML2

Open Source Brain Meeting 2019

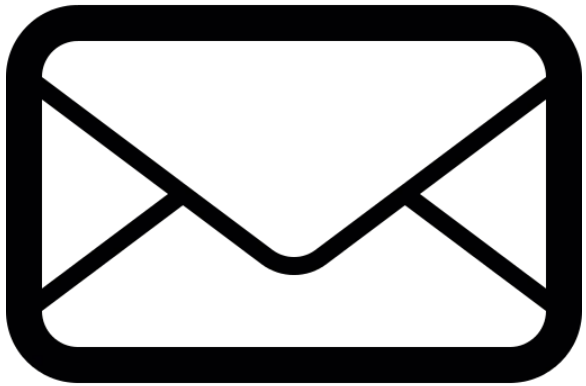
Boris Marin

boris.marin@ufabc.edu.br

Universidade Federal do ABC

Converting NMODL to NeuroML

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



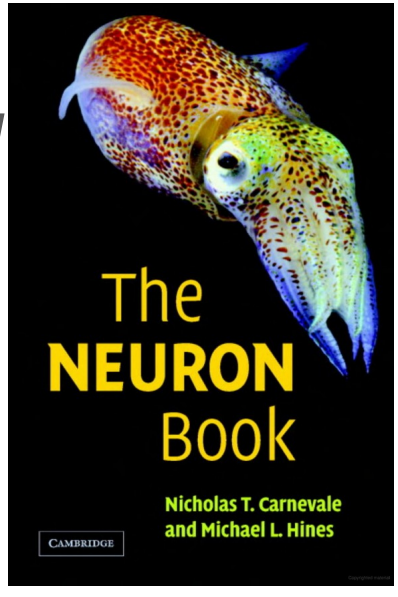
mailto:p.gleeson@ucl.ac.uk

The *NEURON* simulator

Defining models in *NEURON*

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

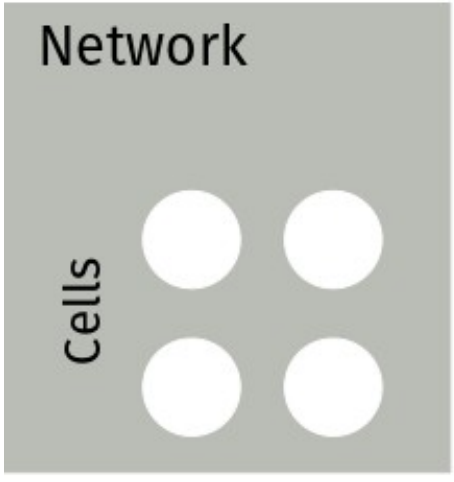
- Cells, Networks: *hoc* language (accessible from Python)
  - morphologies
  - synaptic connections
  - *.hoc* files
- Ion Channels (membrane mechanisms): *NM*
  - *.mod* files



What is NeuroML, and why should I care?

Why can OSB process any NeuroML2 file?

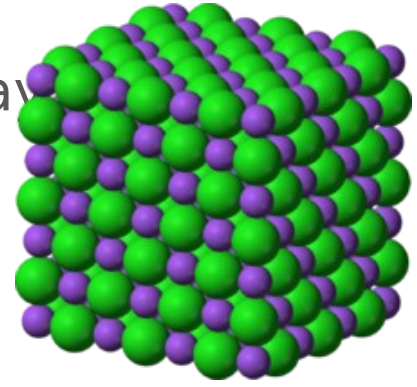
- NML is *structured* (not unlike a *Type System*)



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 way as NML...
  - .... if only people stuck to "good practices"!



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
}
```

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){...}
```

Levels of Abstraction

NetPyNE: structured network specification

i)
```
popParams['EXC_L2'] = {
'cellType':   'PYR',
'cellModel':  'simple',
'yRange':     [100, 400],
'numCells':   50}
```

ii)
```
popParams['EXC_L5'] = {
'cellType':   'PYR',
'cellModel':  'complex',
'yRange':     [700, 1000],
'density':    80e3}
```

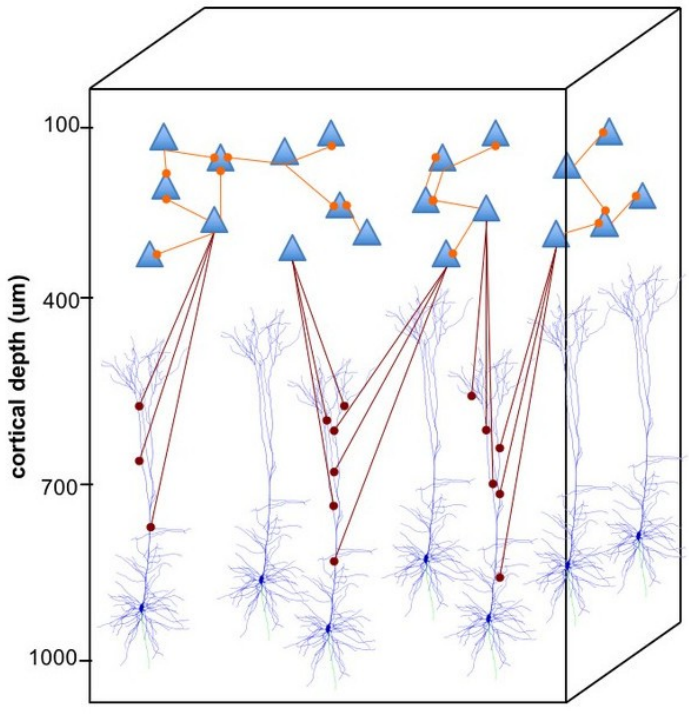
iii)
```
cellParams['PYR_simple'] = {
'conds': {'cellType': 'PYR',
          'cellModel': 'simple'},
'secs':  {'soma':
           'geom': {'diam': 18, 'L': 18},
           'mechs': {'hh':
                     {'gnabar': 0.12,
                      'gkbar': 0.036,
                      'gl':    0.003,
                      'el':    -70}}}}
```

iv)
```
importCellParams(
label =    'PYR_complex',
conds =   {'cellType':   'PYR',
           'cellModel': 'complex'},
fileName = 'L5_pyr_full.hoc',
cellName = 'PYR_L5')
```

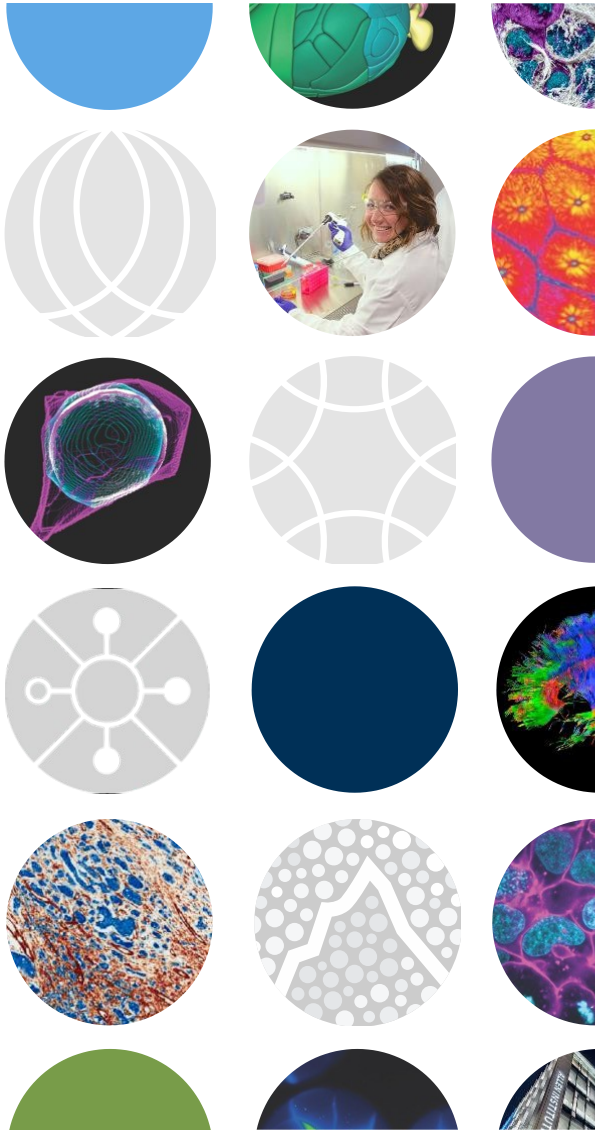
vi)
```
synMechParams['AMPA'] = {
 'mod':   'Exp2Syn',
 'tau1':   0.8,
 'tau2':   5.3,
 'e':      0}
```

vii)
```
connParams['L2->E2'] = {
 'preConds':    {'y': [100, 400]},
 'postConds':   {'pop': 'EXC_L2'},
 'probability': '1*exp(-dist_3D/200)',
 'weight':       0.4,
 'delay':        5,
 'synMech':      'AMPA'}
```

viii)
```
connParams['E2->L5'] = {
 'preConds':    {'pop': 'EXC_L2'},
 'postConds':   {'y':    [700,1100],
                 'cellModel': 'complex'},
 'convergence': 25,
 'weight':      '0.001 * post_ynorm',
 'delay':       'dist_3D/propVelocity',
 'sec':         'allDend',
 'synMech':     'AMPA',
 'synsPerConn': 3}
```



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



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

Yazan N. Billeh
yazanb@alleninstitute.org

ALLEN INSTITUTE

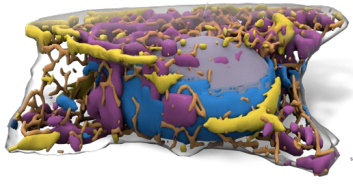
ALLEN INSTITUTE for BRAIN SCIENCE

ALLEN INSTITUTE for CELL SCIENCE

ALLEN INSTITUTE for IMMUNOLOGY

THE 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

CORE PRINCIPLES

Team Science

Interdisciplinary teams working towards common goal

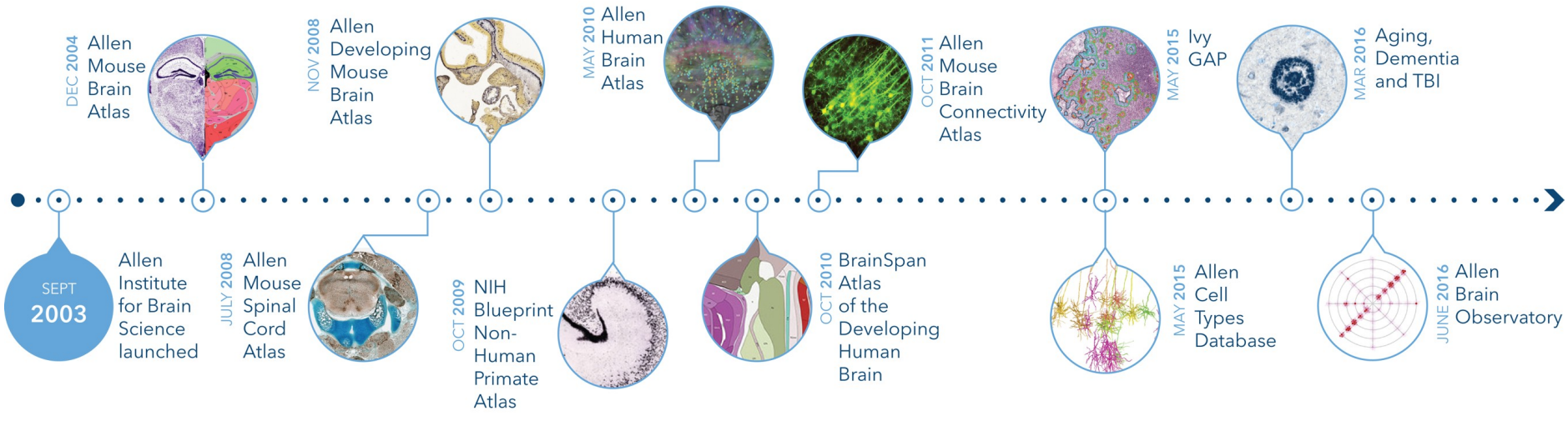
Big Science

Large-scale projects with robust, massive data

Open Science

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

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

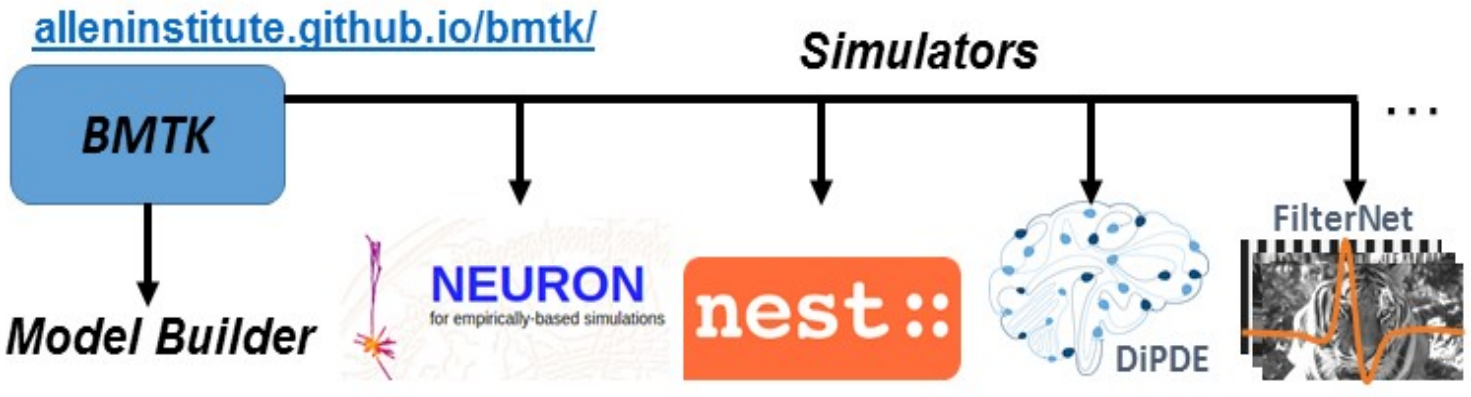


All data are:

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

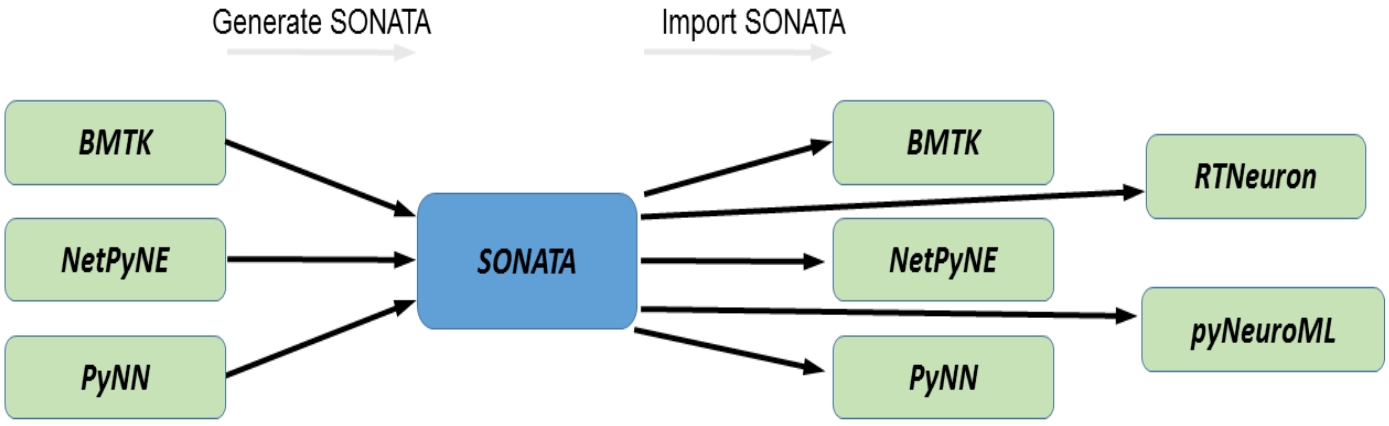
Our Models and Modeling Software Are Freely Available to the Community

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



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



Human Brain Project

How model standardization enables new tools and applications in neuroscientific research

Insights from the HBP

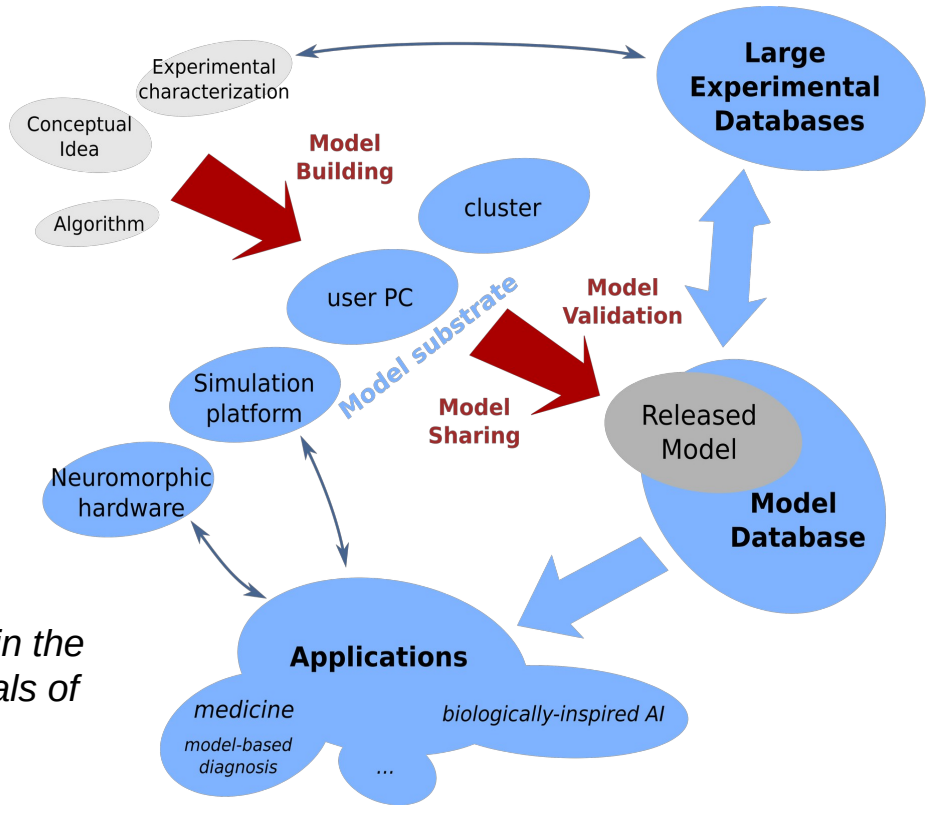
Yann Zerlaut

Neuroinformatics team / group of A. Davison

Centre National de la Recherche Scientifique, France

Open Source Brain Meeting 2019, Alghero

Motivation



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

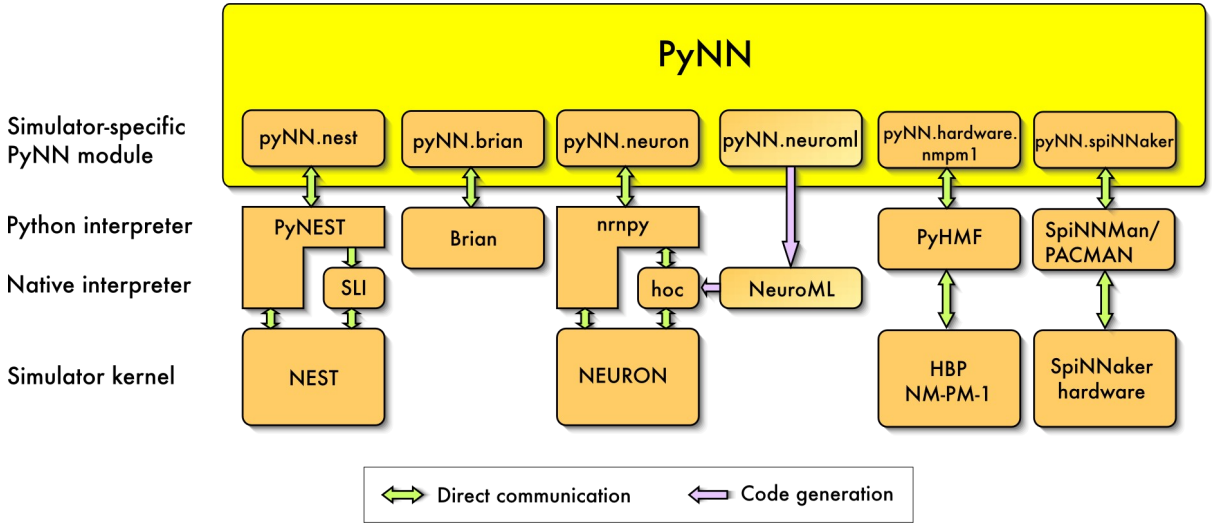
PyNN — A unified interface for neuronal network simulators

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)



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

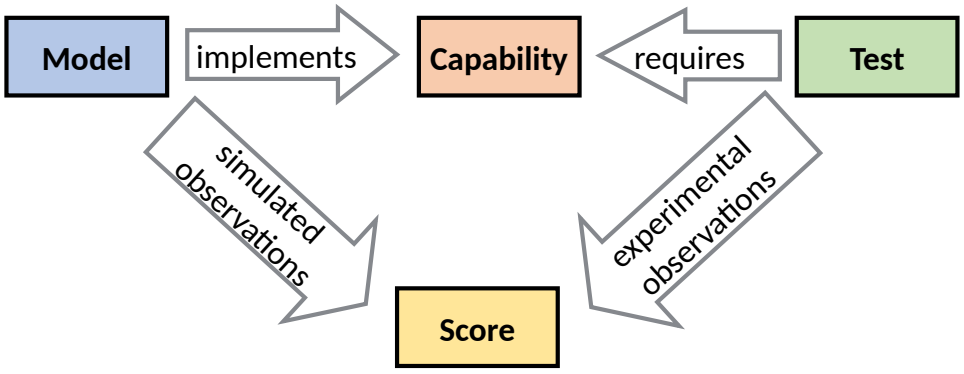
```
from pyNN.serialization import import_from_sonata, load_sonata_simulation_plan
import pyNN.neuron as sim

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)
```

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

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:

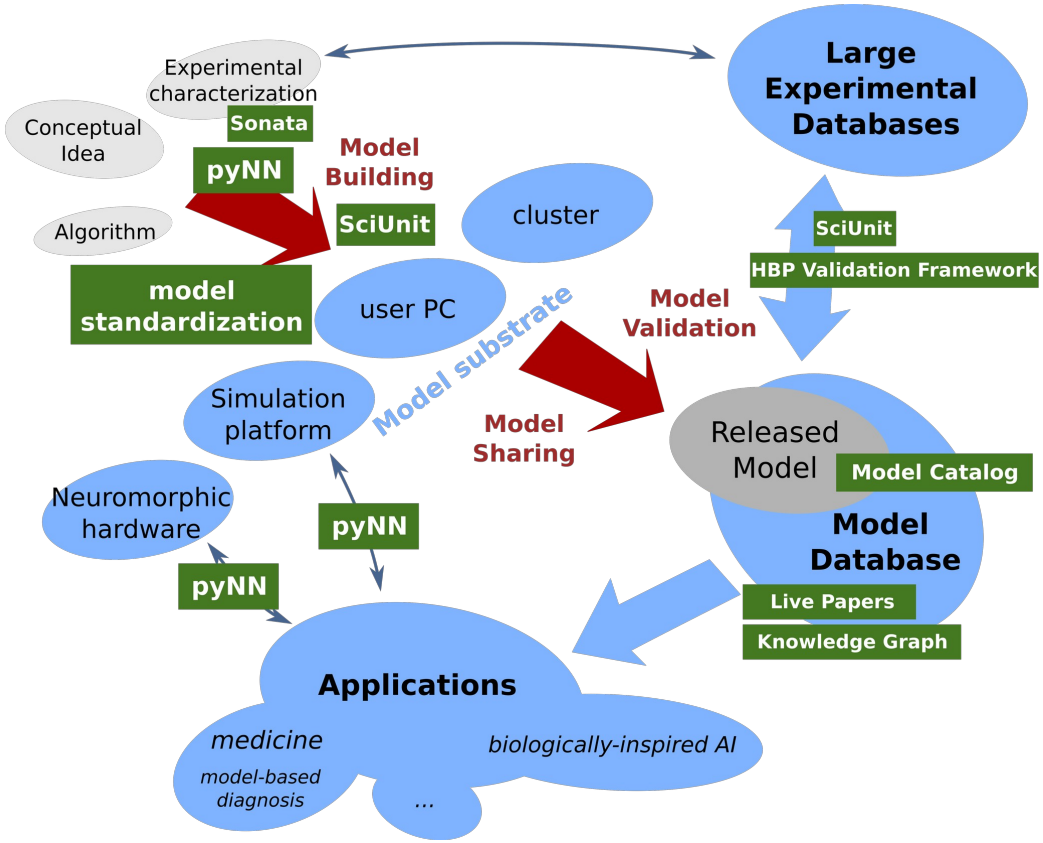
Test suites for specific brain regions

- ***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`

Summary


Large Experimental Databases
Experimental characterization
Sonata
Conceptual Idea
pyNN
Model Building
SciUnit
Algorithm
cluster
model standardization
user PC
SciUnit
HBP Validation Framework
Model Validation
Model substrate
Simulation platform
Model Sharing
Released Model
Model Catalog
Neuromorphic hardware
pyNN
pyNN
Model Database
Live Papers
Knowledge Graph
Applications
medicine
model-based diagnosis
...
biologically-inspired AI