

# LARGE-SCALE MODEL DEVELOPMENT FROM THE NEST PERSPECTIVE

17 July 2018

CNS\*2018 workshop “Developing, standardising and sharing large scale cortical network models”

Seattle, USA

SACHA VAN ALBADA

Institute of Neuroscience and Medicine (INM-6)

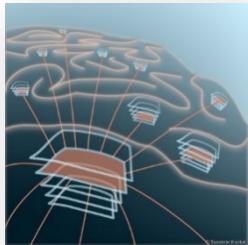
Institute for Advanced Simulation (IAS-6)

JARA Brain Institute I (INM-10)

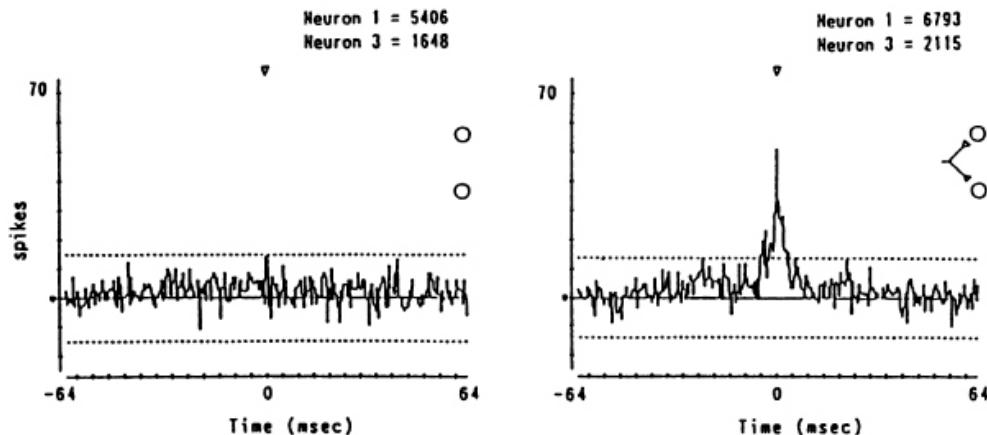
Jülich Research Centre

[www.csn.fz-juelich.de](http://www.csn.fz-juelich.de)

[www.nest-initiative.org](http://www.nest-initiative.org)



# NEED FOR LARGE-SCALE MODELS: IRREDUCIBILITY



Sakurai (1999) *Neurosci Biobehav Rev*

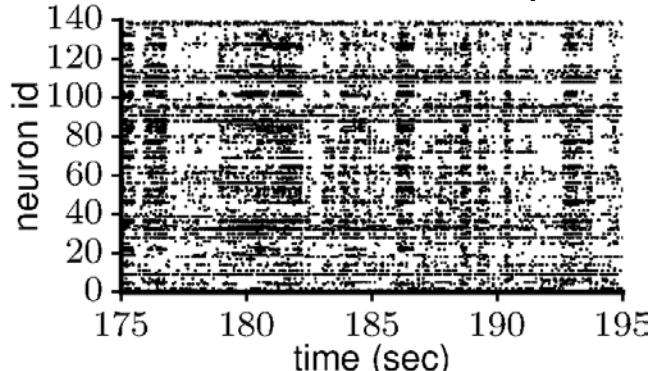
- two simultaneously recorded neurons in CA1 of a rat performing an auditory or visual discrimination task
- task-related correlation only for visual task

- downscaling can preserve certain dynamical properties, but never all; e.g.,
  - reducing number of neurons tends to increase correlations
  - simultaneously preserving mean activities and correlations is only possible over a limited range of network sizes
- testing goodness of approximation requires full-scale simulations

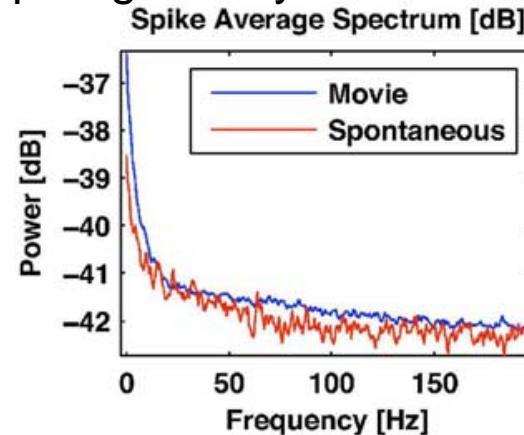
van Albada SJ, Helias M, Diesmann M (2015) *PLOS CB*

# NEED FOR MULTI-SCALE MODELS: LOCAL AND GLOBAL DYNAMICS

low frequencies in spiking activity

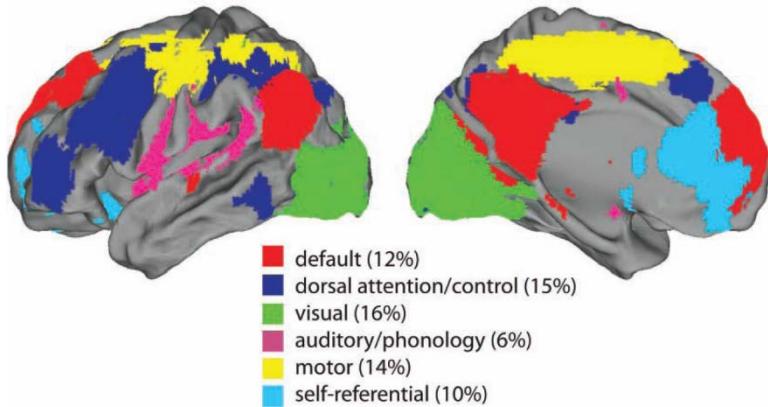


Chu et al. (2014) *Vision Res*



Belitski et al.  
(2008) *J Neurosci*

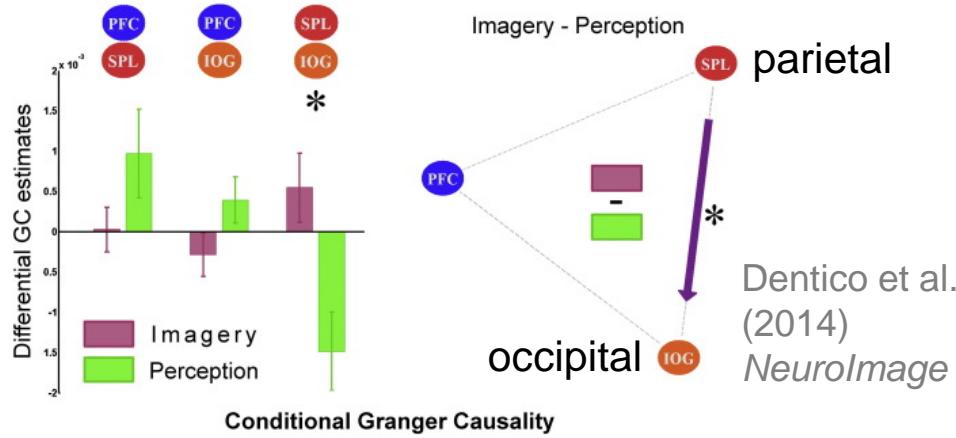
resting-state networks



Deco and Corbetta (2011) *The Neuroscientist*

Member of the Helmholtz Association

inter-area propagation



parietal

occipital

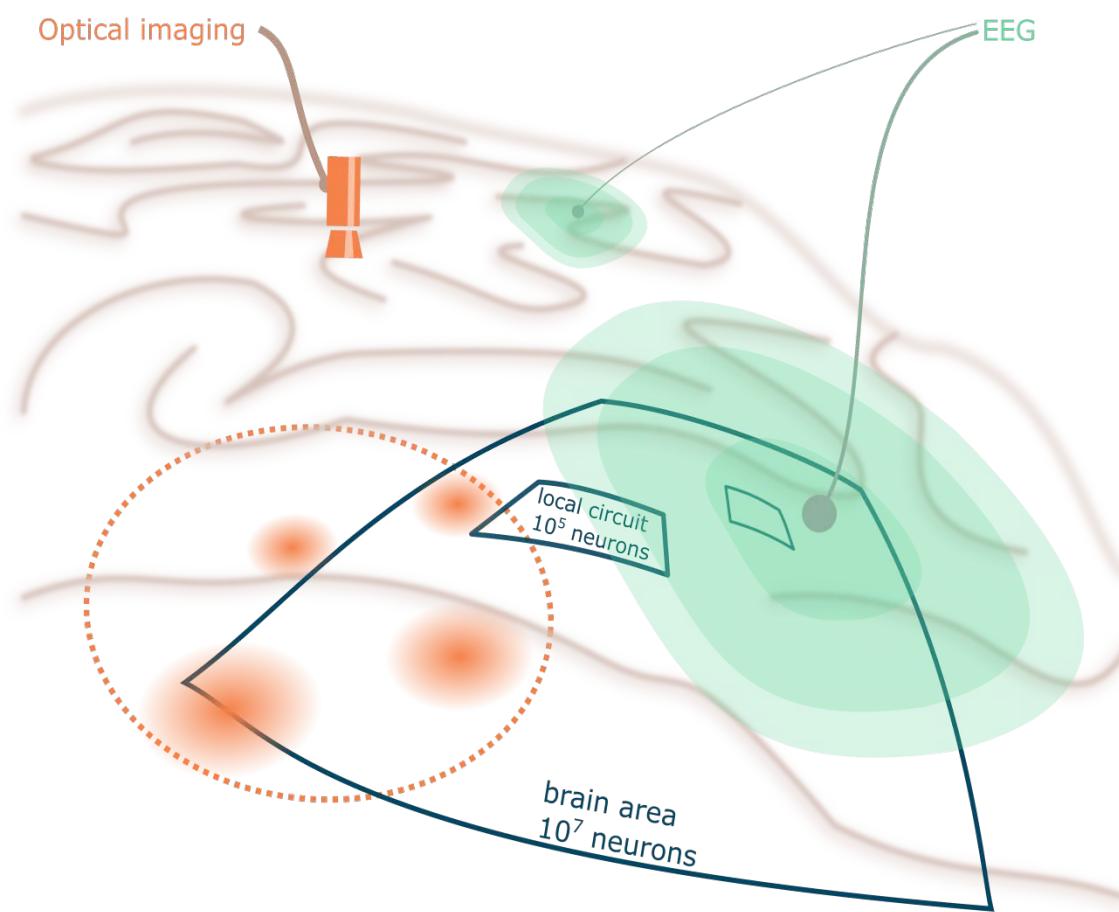
Dentico et al.  
(2014)  
*NeuroImage*

# NEED FOR MULTI-SCALE MODELS: LINKS TO EXPERIMENT

brain-scale networks basis  
for various measures by  
forward modeling

mesoscopic measures  
• local field potential (LFP)  
• voltage sensitive dyes (VSD)

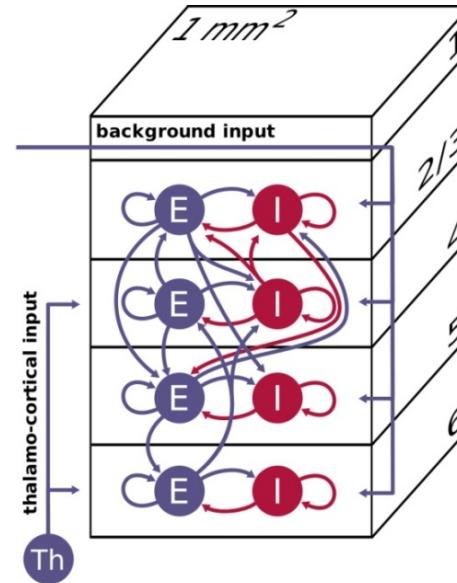
and macroscopic measures  
• EEG, MEG  
• fMRI resting-state networks



S Kunkel

# MINIMAL LAYERED CORTICAL NETWORK MODEL

- 1 mm<sup>3</sup>
- 300 million synapses, 80,000 neurons
- 2 populations of neurons per layer:
  - E: excitatory
  - I: inhibitory
- E and I identical neuronal dynamics
- laterally homogeneous connectivity
- layer- and type-specific connection probability



## Cerebral CORTEX

The Cell-Type Specific Cortical Microcircuit: Relating Structure and Activity in a Full-Scale Spiking Network Model 

Tobias C. Potjans , Markus Diesmann

*Cerebral Cortex*, Volume 24, Issue 3, 1 March 2014, Pages 785–806,  
<https://doi.org/10.1093/cercor/bhs358>

# MINIMAL LAYERED CORTICAL NETWORK MODEL

integrates knowledge of more than  
50 experimental papers

taking into account layer and neuron-type  
specific connectivity is sufficient to  
reproduce experimentally observed:

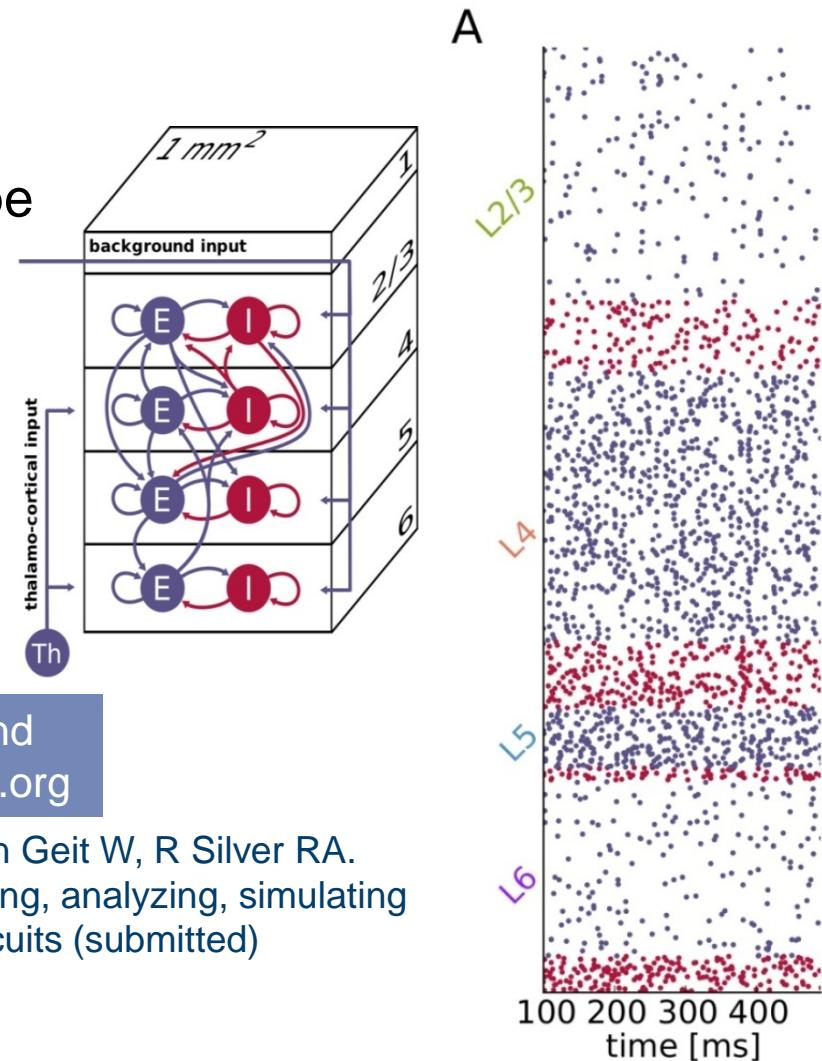
- asynchronous irregular spiking
- higher rate of inhibitory neurons
- correct distribution of spike rates  
across layers

made available as a part of NEST and  
PyNN, and at [www.opensourcebrain.org](http://www.opensourcebrain.org)

Gleeson P, Cantarelli M, Marin B, . . . van Albada SJ, van Geit W, R Silver RA.  
Open Source Brain: a collaborative resource for visualizing, analyzing, simulating  
and developing standardized models of neurons and circuits (submitted)

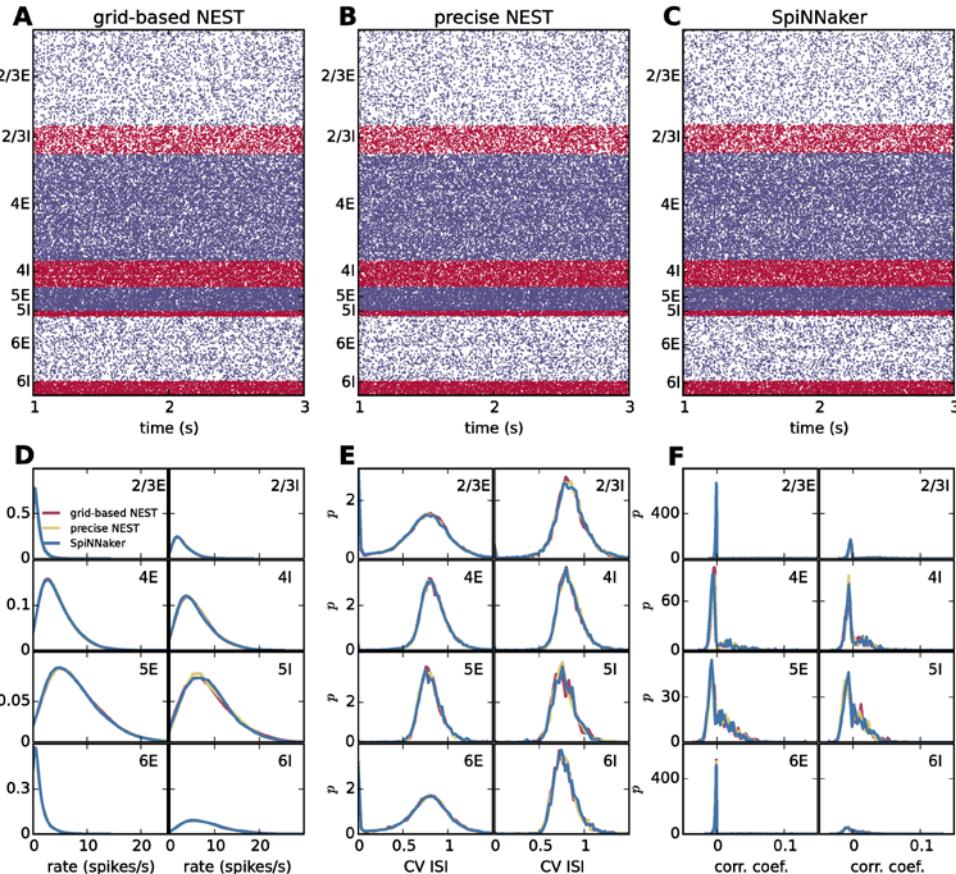
reproduced with Brian

Shimoura RO, Kamiji NL, Pena RFO, Cordeiro VL,  
Ceballos CC, Romaro C, Roque AC (2018) *ReScience*



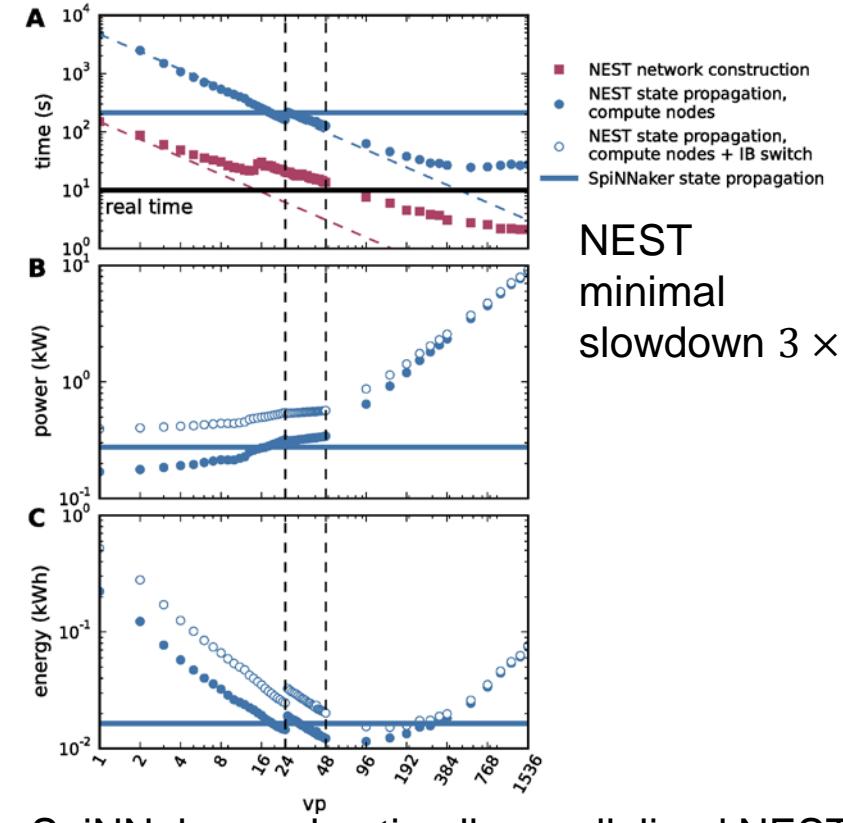
# MICROCIRCUIT PORTED TO SPINNAKER

largest SpiNNaker simulation to date; 6 SpiNNaker boards; 20 × slowdown to accommodate fast dynamics (0.1 ms time steps) and high input spike rates



van Albada, Rowley, Senk, Hopkins, Schmidt, Stokes, Lester, Diesmann, Furber (2018) *Front Neurosci*

Member of the Helmholtz Association



SpiNNaker and optimally parallelized NEST have comparable total energy consumption during run phase

# BUILDING BLOCK FOR FURTHER STUDIES

- used in 17 peer-reviewed studies
- cited in 62 peer-reviewed publications

Neuron  
Volume 72, Issue 5, 8 December 2011, Pages 859–872  
Cell Press

Article  
Modeling the Spatial Reach of the LFP  
Henrik Lindén<sup>1,2</sup>, Tom Tetzlaff<sup>1,3</sup>, Tobias C. Potjans<sup>3,4,5</sup>, Klas H. Pettersen<sup>1,6</sup>, Sonja Grün<sup>3,7,8</sup>, Markus Diesmann<sup>3,4,8,9</sup>, Gáute T. Einevoll<sup>1,6</sup>,   
+ Show more  
<http://doi.org/10.1016/j.neuron.2011.11.006>  
Under an Elsevier user license

frontiers in Computational Neuroscience  
ORIGINAL RESEARCH ARTICLE  
Front. Comput. Neurosci., 08 July 2011 | <https://doi.org/10.3389/fncom.2011.00031>

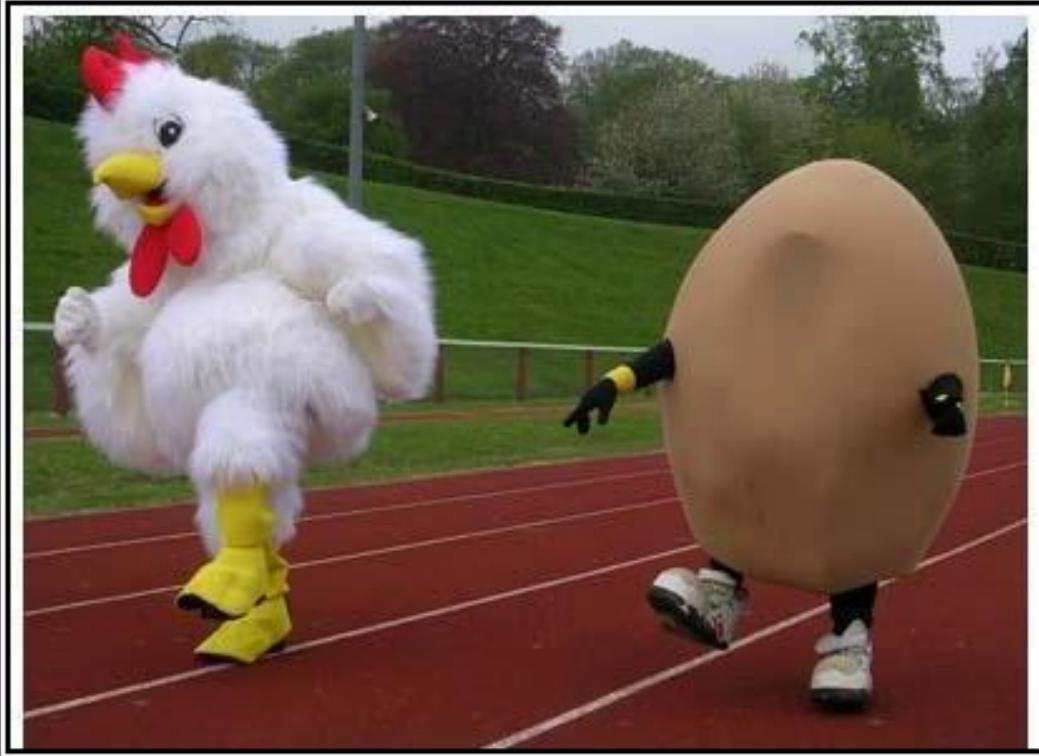
Layer-dependent attentional processing by top-down signals in a visual cortical microcircuit model  
Nobuhiko Wagatsuma<sup>1,2\*</sup>, Tobias C. Potjans<sup>3,4,5</sup>, Markus Diesmann<sup>1,3,4</sup> and Tomoki Fukai<sup>1,4,6</sup>

Cornell University Library  
arXiv.org > q-bio > arXiv:1511.09364  
Quantitative Biology > Neurons and Cognition  
Full-density multi-scale account of structure and dynamics of macaque visual cortex  
Maximilian Schmidt, Rembrandt Bakker, Kelly Shen, Gleb Bezgin, Claus-Christian Hilgetag, Markus Diesmann, Sacha J. van Albeda  
(Submitted on 20 Nov 2015 (v1), last revised 15 Apr 2016 (this version, v6))  
We present a multi-scale structural and population-resolved model of the macaque visual cortex. Simulations reproduce longer intrinsic and cortico-cortical interactions. Detailed connectivity of cortical layers is shown.  
Subjects: Neurons and Cognitive Models  
Cite as: arXiv:1511.09364 [q-bio] (or arXiv:1511.09364v6)  
PLOS COMPUTATIONAL BIOLOGY

RESEARCH ARTICLE  
The Computational Properties of a Simplified Cortical Column Model  
Nicholas Cain, Remake Krishnan Iyer, Christof Koch, Stefan Mihalas\*  
Allen Institute for Brain Science, Seattle, Washington, United States of America  
starfarm@alleninstitute.org  
Abstract  
The mammalian neocortex has a repetitive, laminar structure and performs functions integral to higher cognitive processes, including sensory perception, memory, and coordinated motor output. What computations does this circuitry subsserve that link these unique structural elements to their function? Potjans and Diesmann (2014) parameterized a four-layer, two-cell-type (i.e. excitatory and inhibitory) model of a cortical column with homogeneous

frontiers in Computational Neuroscience  
A Computational Analysis of the Function of Three Inhibitory Cell Types in Contextual Visual Processing  
Jung H. Lee\*, Christof Koch and Stefan Mihalas\*  
Allen Institute for Brain Science, Seattle, WA, USA

Towards a theory of cortical columns: From spiking neurons to interacting neural populations of finite size  
Tilo Schwalger<sup>1,\*</sup>, Moritz Deger<sup>1,2</sup>, Wulfram Gerstner<sup>1</sup>,  
1 Brain Mind Institute, School of Computer and Communication Sciences and School of Life Sciences, École Polytechnique Fédérale de Lausanne (EPFL), Lausanne, Switzerland  
2 Institute for Zoology, Faculty of Mathematics and Natural Sciences, University of Cologne, Cologne, Germany  
\* [tilo.schwalger@epfl.ch](mailto:tilo.schwalger@epfl.ch)



# THIS RACE

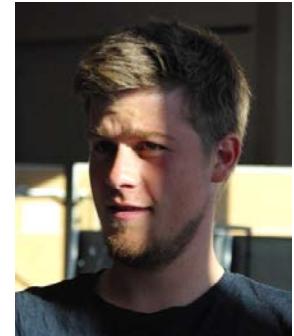
could be interesting

VERY DEMOTIVATIONAL .com

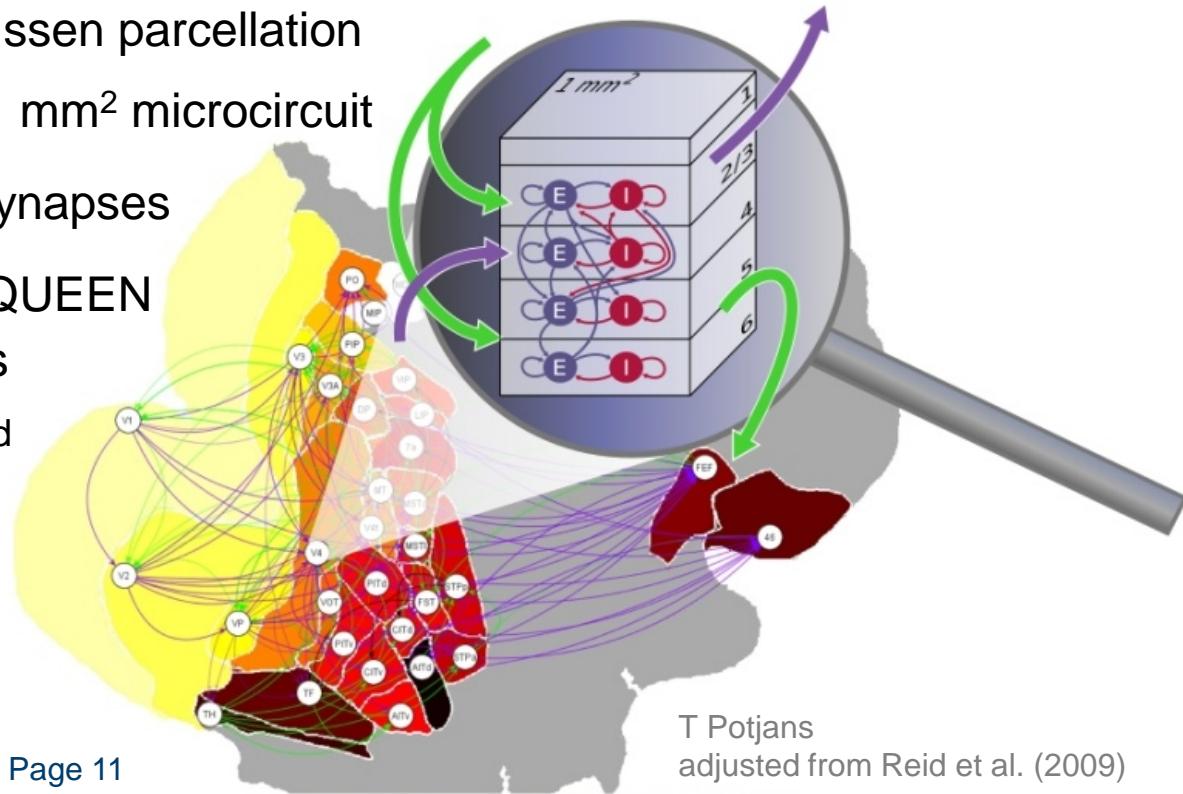


# MULTI-AREA MODEL OF MACAQUE VISUAL CORTEX

- rich anatomical and physiological data sets available
- relatively close to human
- 800 million neurons in one hemisphere
- 32 areas in Felleman & Van Essen parcellation
- representing each area by a  $1 \text{ mm}^2$  microcircuit
- $4 \cdot 10^6$  neurons and  $2.4 \cdot 10^{10}$  synapses
- simulated using NEST on JUQUEEN and JURECA supercomputers
  - $\sim 10^3$  MPI processes, multi-threaded
  - a few minutes for 1 s of simulation



Maximilian Schmidt



# SIMULATION TECHNOLOGY FOR EXASCALE



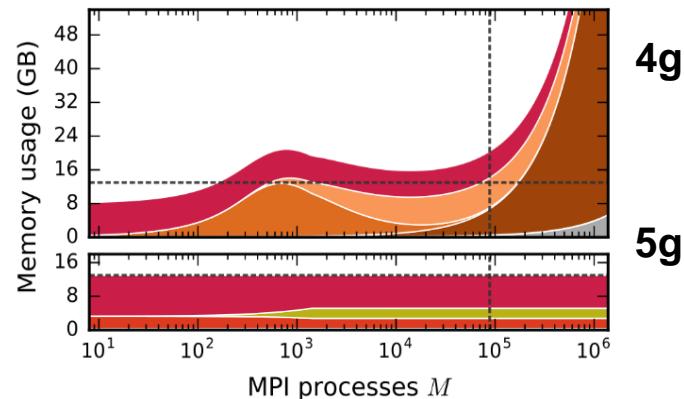
ORIGINAL RESEARCH  
published: 16 February 2018  
doi: 10.3389/fninf.2018.00002

## Extremely Scalable Spiking Neuronal Network Simulation Code: From Laptops to Exascale Computers

Jakob Jordan<sup>1\*</sup>, Tammo Ippen<sup>1,2</sup>, Moritz Helias<sup>1,3</sup>, Itaru Kitayama<sup>4</sup>, Mitsuhsisa Sato<sup>4</sup>, Jun Igarashi<sup>5</sup>, Markus Diesmann<sup>1,3,6</sup> and Susanne Kunkel<sup>7,8</sup>

<sup>1</sup> Institute of Neuroscience and Medicine (INM-6) and Institute for Advanced Simulation (IAS-6) and JARA Institute Brain Structure-Function Relationships (INM-10), Jülich Research Centre, Jülich, Germany, <sup>2</sup> Faculty of Science and Technology, Norwegian University of Life Sciences, Ås, Norway, <sup>3</sup> Department of Physics, Faculty 1, RWTH Aachen University, Aachen, Germany, <sup>4</sup> Advanced Institute for Computational Science, RIKEN, Wako, Japan, <sup>5</sup> Computational Engineering Applications Unit, RIKEN, Wako, Japan, <sup>6</sup> Department of Psychiatry, Psychology and Psychotherapy, Institute of Psychosomatics, Medical Faculty, RWTH Aachen University, Aachen, Germany, <sup>7</sup> Department of Computer Science, Royal Institute of Technology (KTH), Stockholm, Sweden, <sup>8</sup> Department of Communication, KTH Royal Institute of Technology, Stockholm, Sweden, <sup>9</sup> Institute for Parallel Computing, Institute for Simulation and Database Technology, Jülich, Germany

shows how to make memory consumption of compute nodes independent of network size



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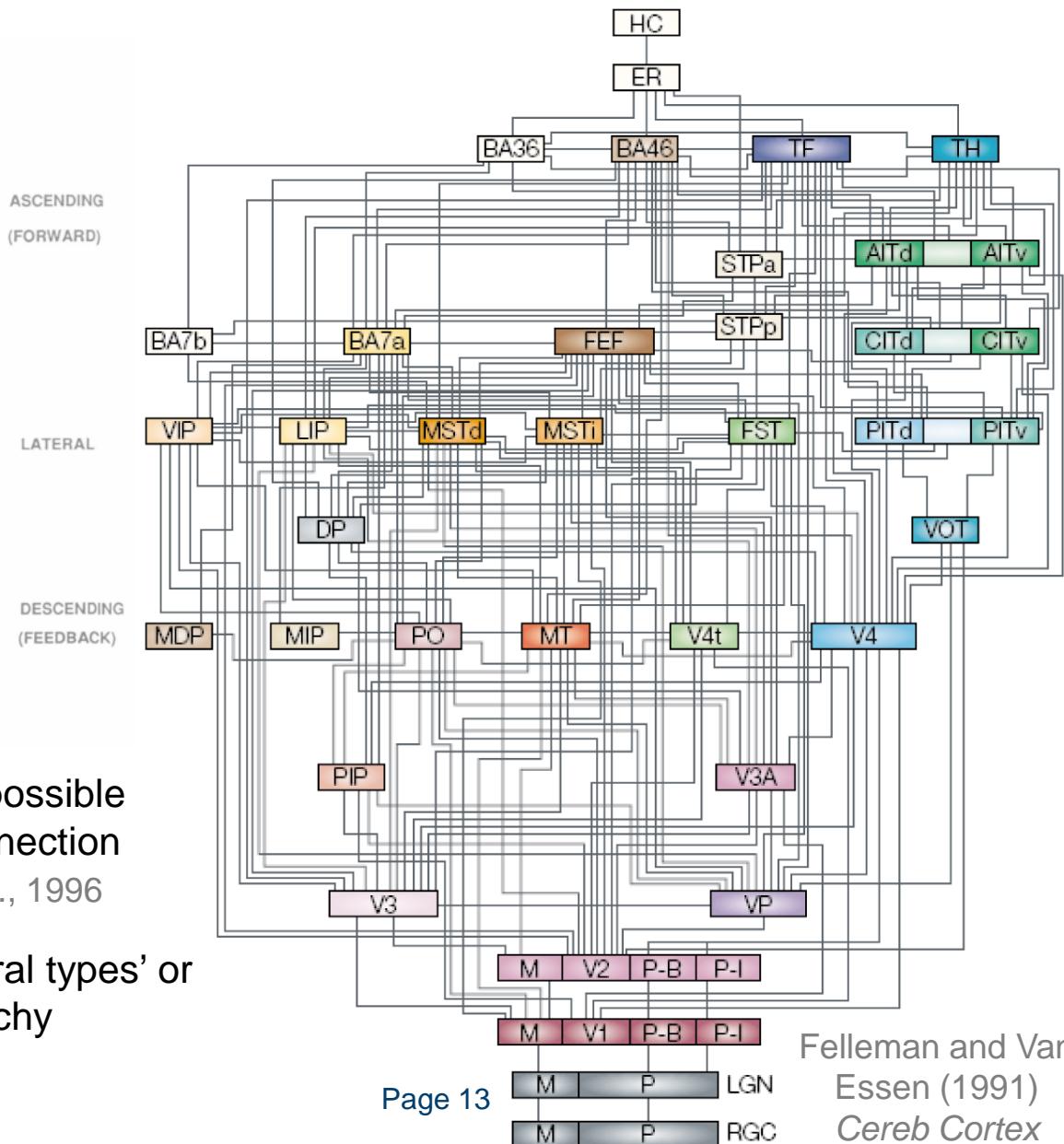
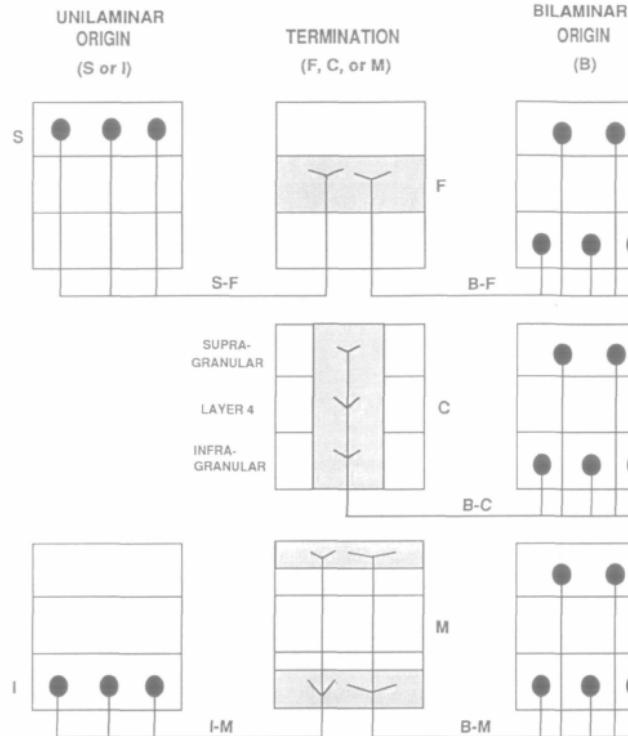
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[2018-07-16]

parallel network construction Ippen et al. (2017) *Front Neuroinform*

# HIERARCHY OF VISUAL CORTICAL AREAS

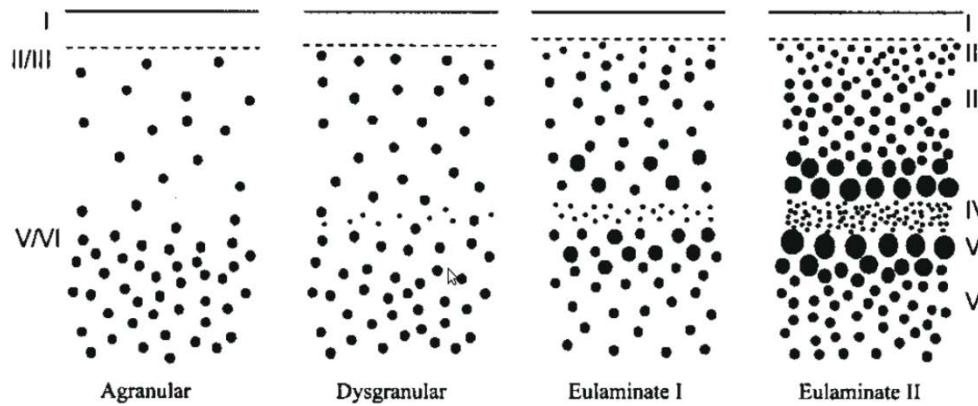


- this is only one of thousands of possible orderings fitting the pairwise connection patterns equally well Hilgetag et al., 1996
- alternatively, can use `architectural types' or neuron densities to define hierarchy

# CORTICAL AREAS ARE NOT ALL THE SAME

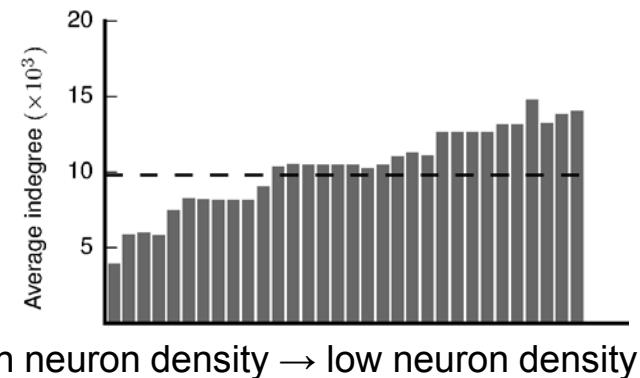
- architectural types
  - neuron densities and layer 4 thickness decrease up the hierarchy
  - total cortical thickness increases up the hierarchy

examples of architectural types



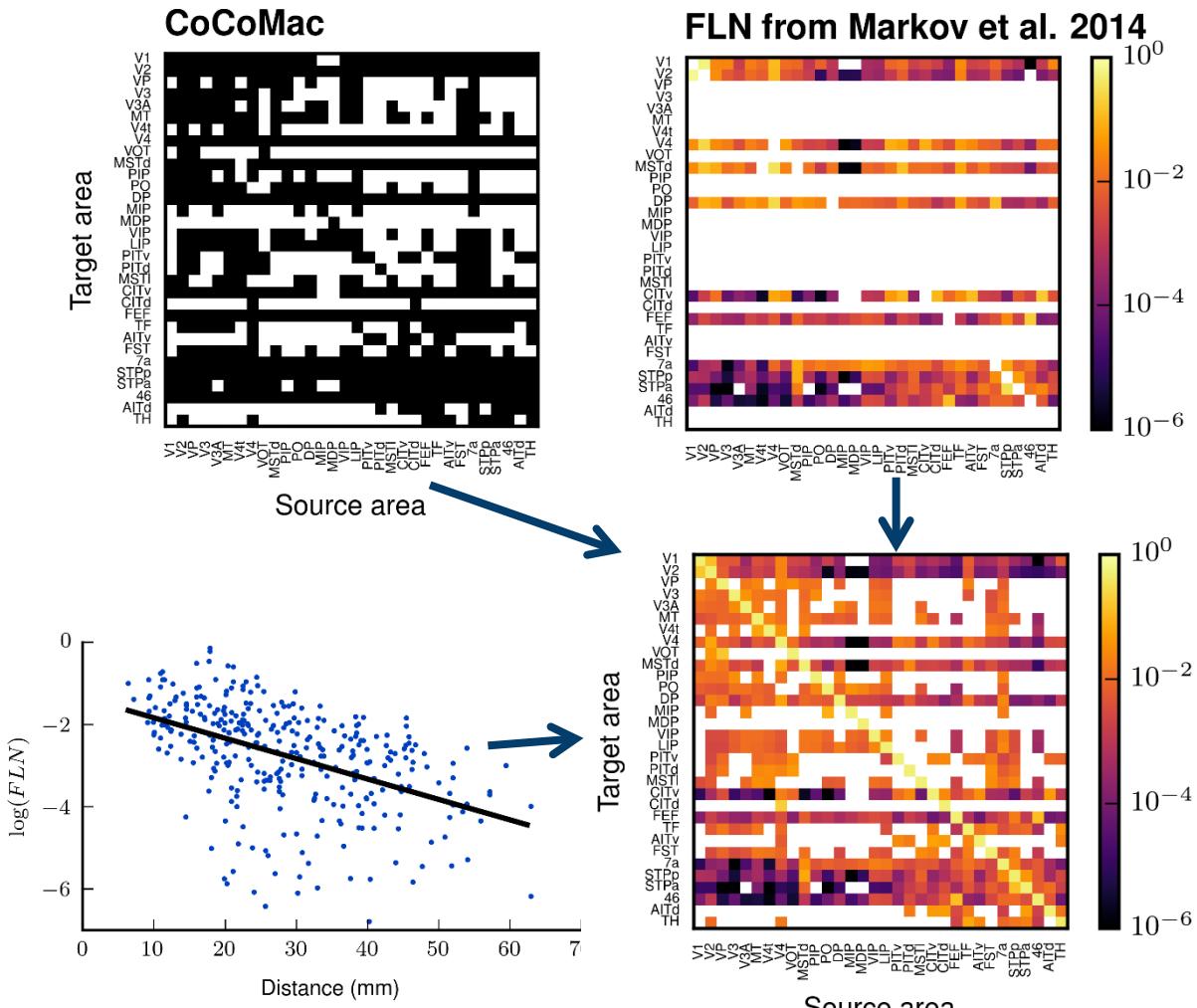
Dombrowski et al. (2001) *Cereb Cortex*

- synapse density remains roughly constant
- → higher areas receive more synapses per neuron



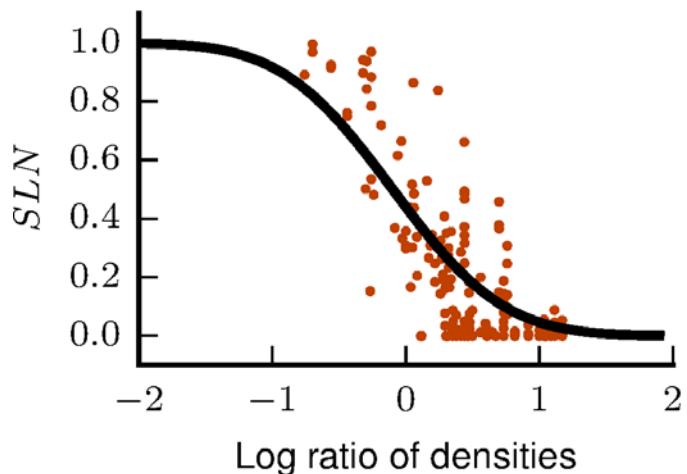
# CONNECTIVITY MAP FROM TRACING DATA

- partly binary, partly quantitative data
- connection probability decays with distance also for inter-area connections
- use this decay to estimate missing data based on distance between areas
- roughly 2/3 of area pairs are connected
- more important: connection density, spanning ~6 orders of magnitude

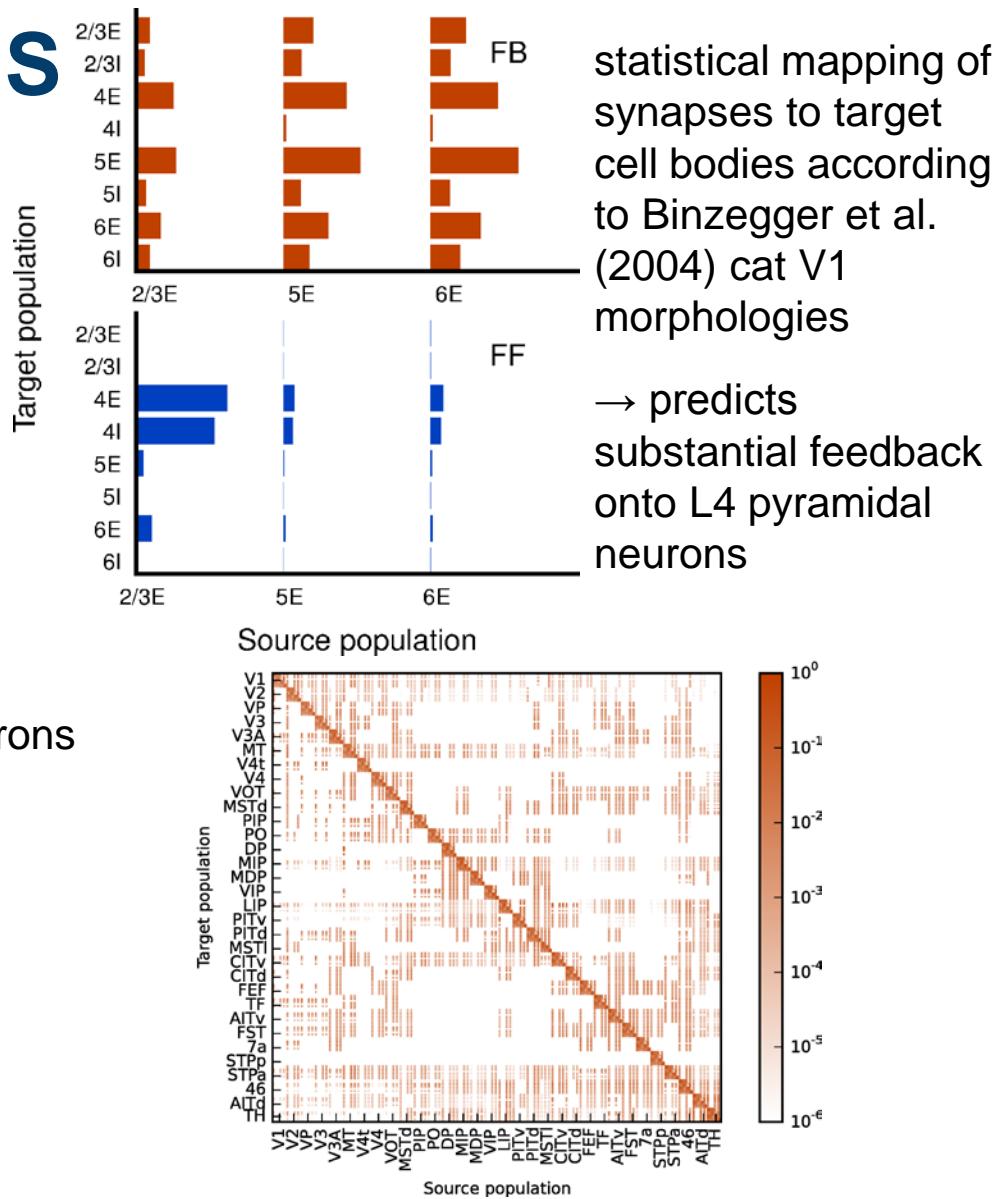
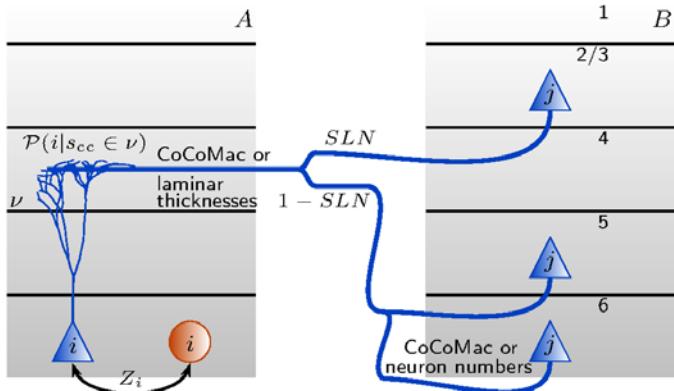


Ercsey-Ravasz et al. (2013) *Neuron*

# LAMINAR PATTERNS

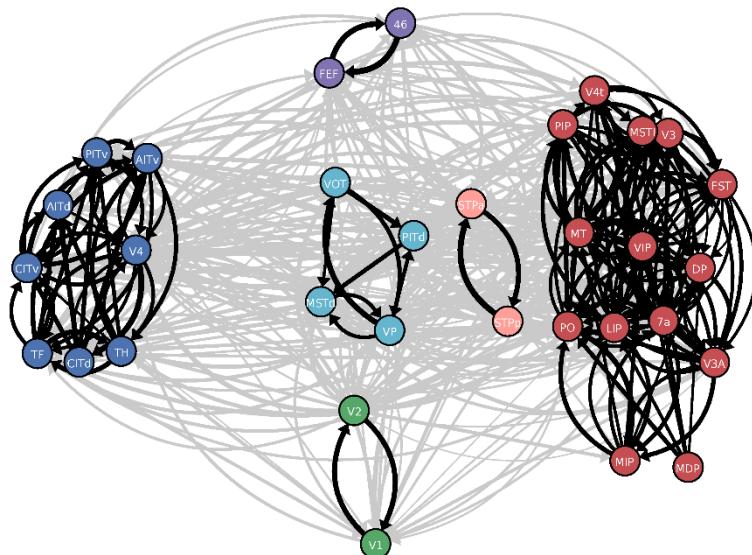
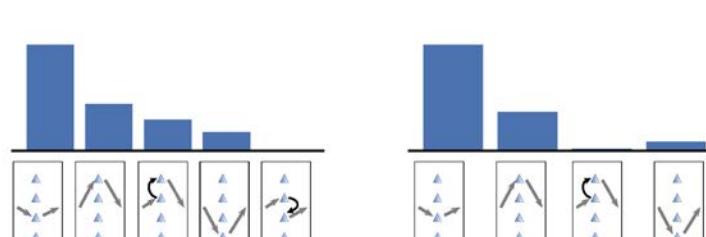
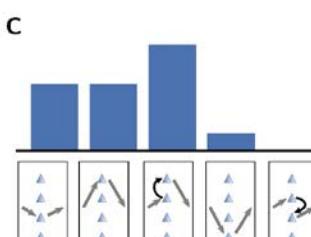
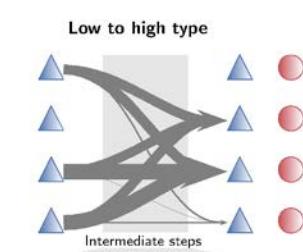
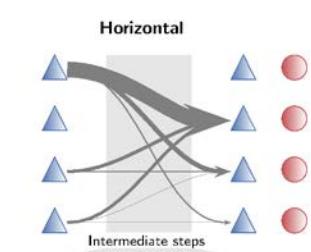
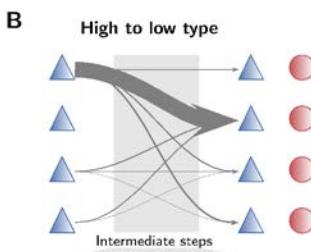
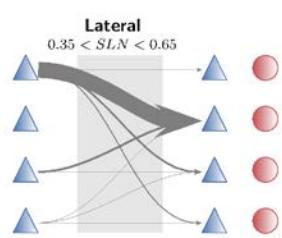
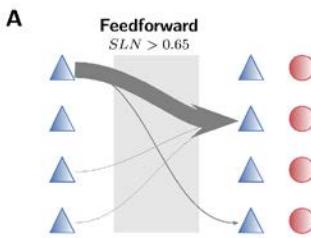


SLN = fraction of supragranular labeled neurons



# CONNECTIVITY ANALYSIS

data integration enables novel characterization of laminar pathways

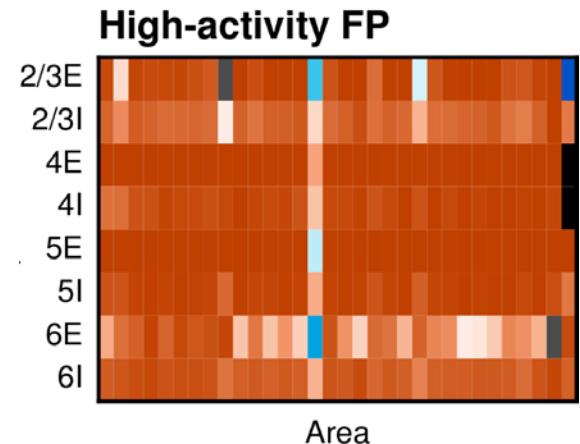
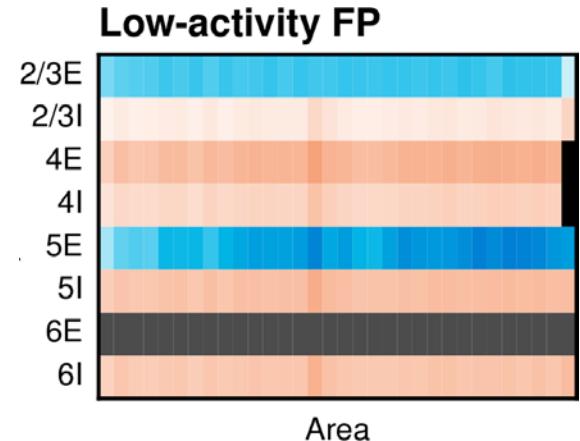
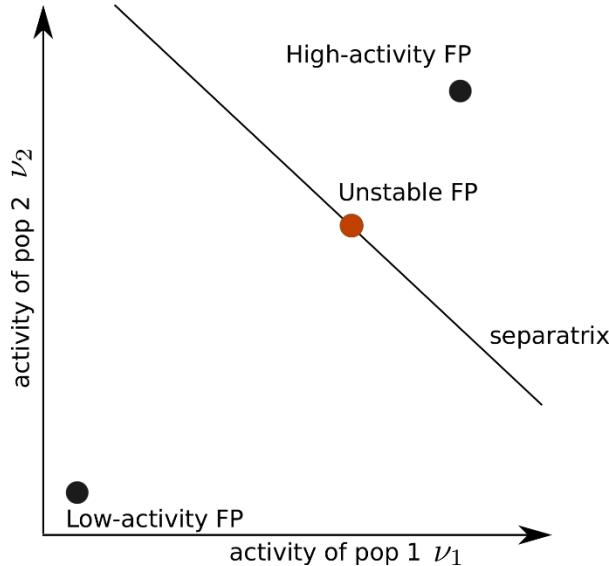
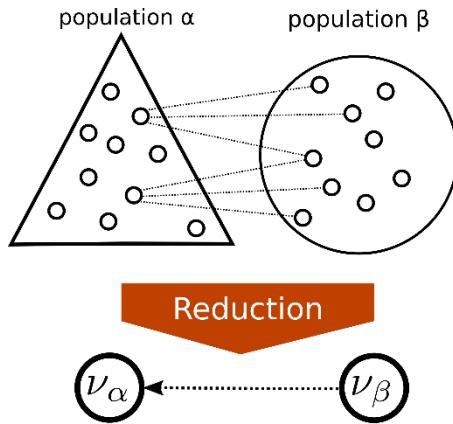


structural connectivity reveals functionally relevant community structure

Schmidt M, Bakker R, Hilgetag CC,  
Diesmann M, van Albada SJ  
(2018) *Brain Struct Func*

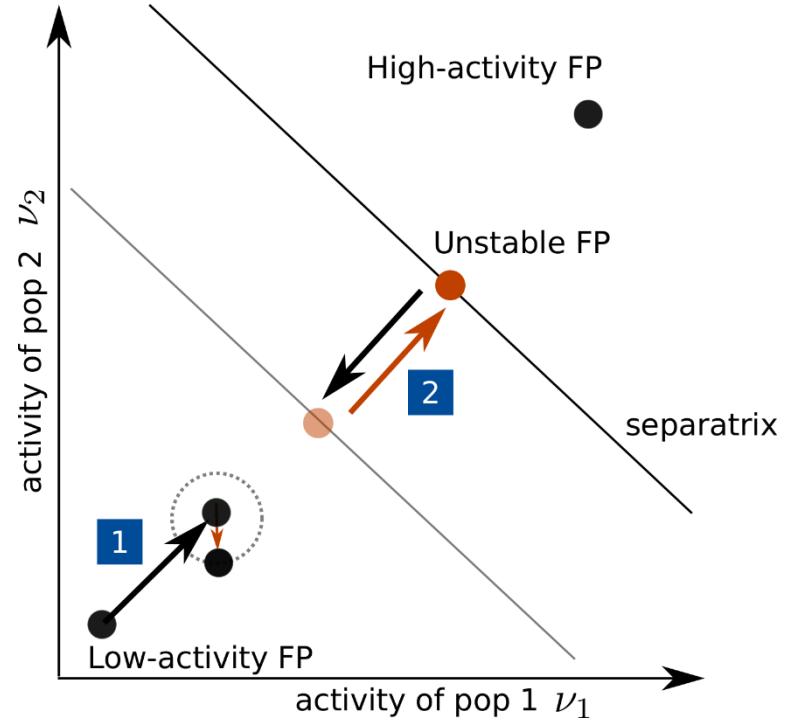
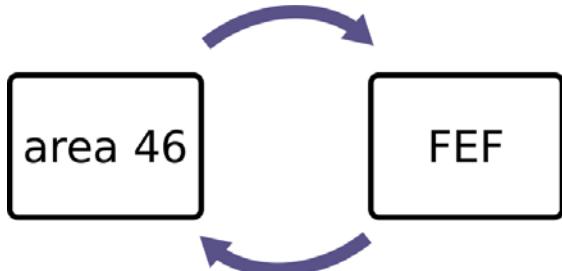
# BISTABLE GROUND STATE

- low rates of excitatory neurons in infragranular layers
- increasing external drive leads to high-activity state with excessive rates



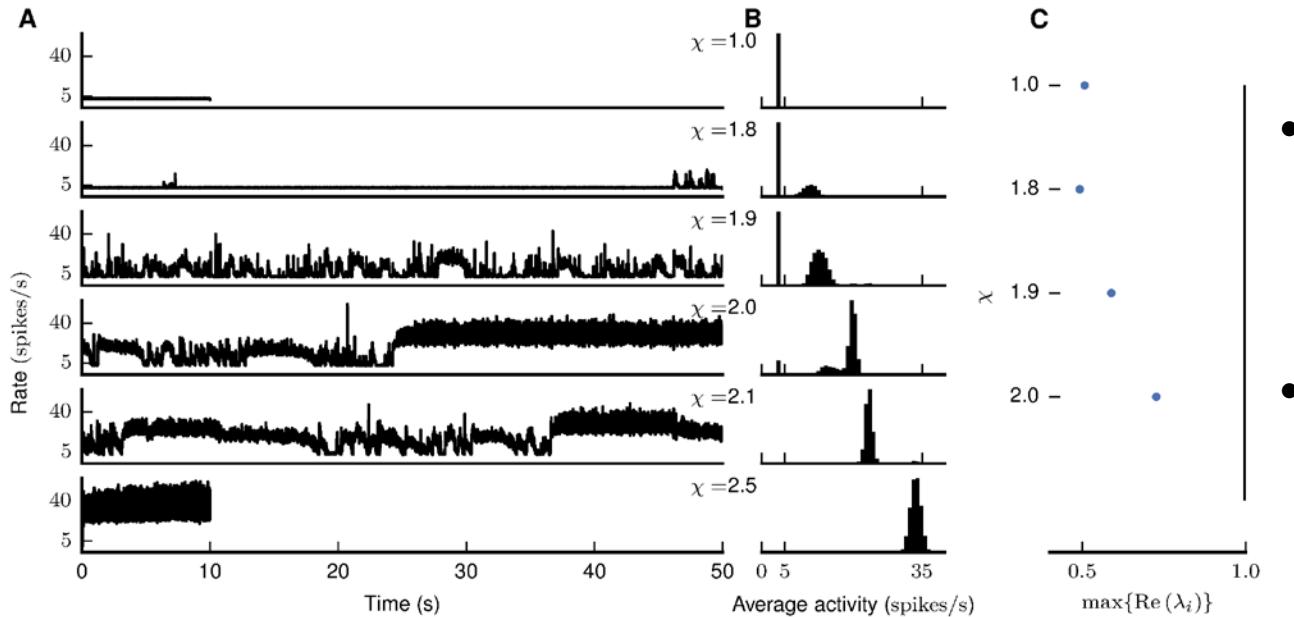
# STABILIZATION USING MEAN-FIELD THEORY

- 1) increase in external drive shifts both low-activity fixed point and separatrix
  - 2) compensation by change in connectivity shifts back separatrix
- yields reasonable rates in all populations
  - reduced connectivity in frontal loop

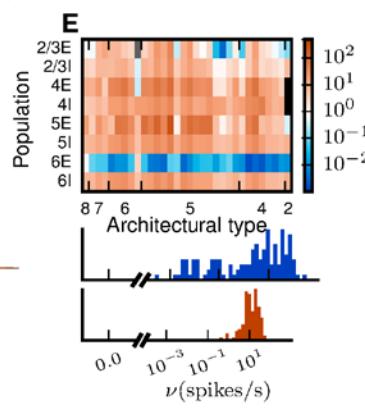
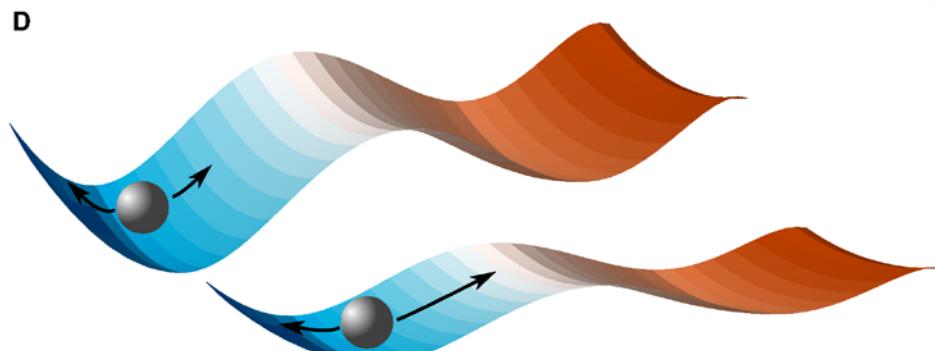


Schuecker J, Schmidt M, van Albada SJ, Diesmann M, Helias M (2017) *PLOS CB*

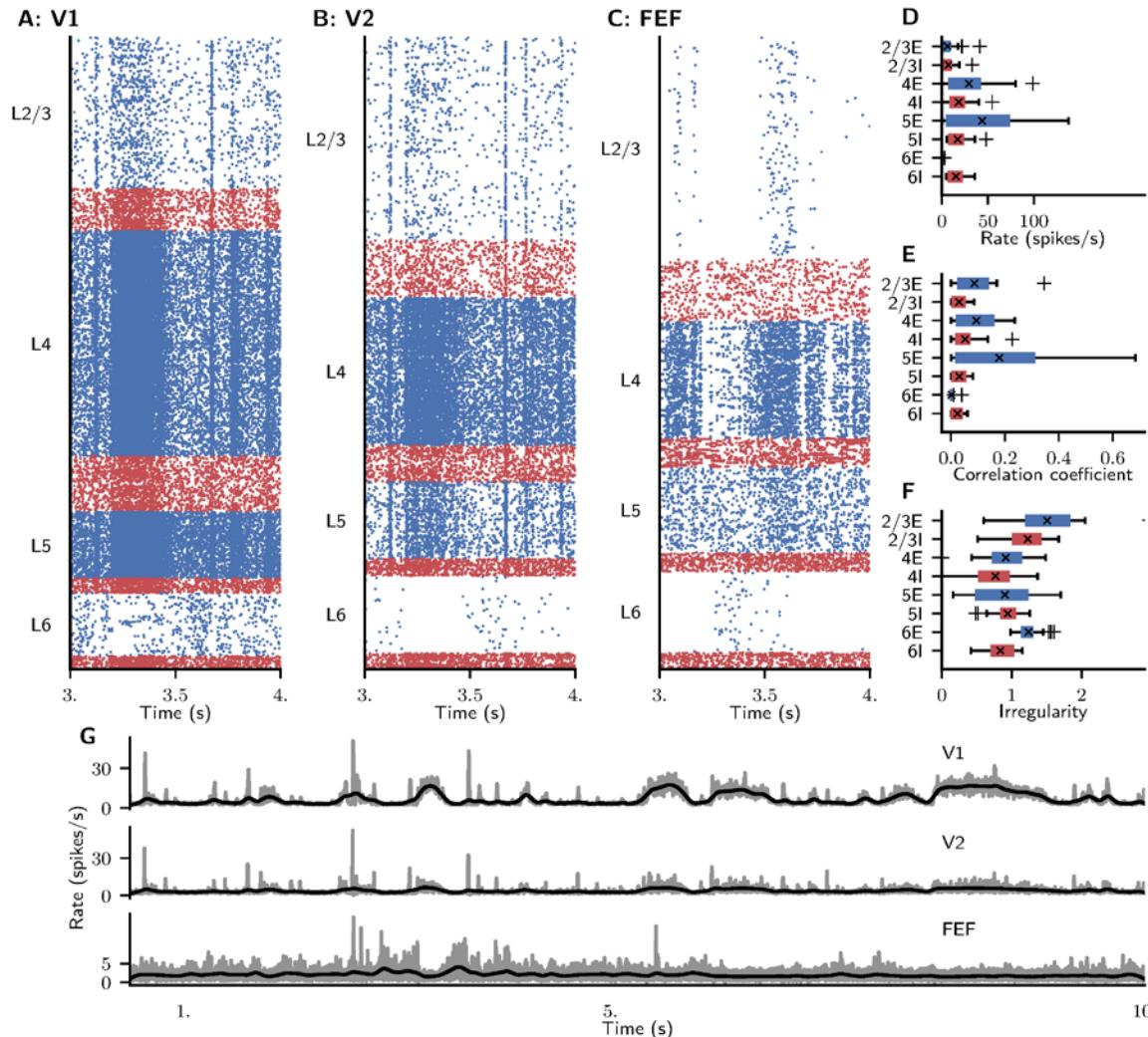
# EMERGENCE OF META-STABLE NETWORK DYNAMICS



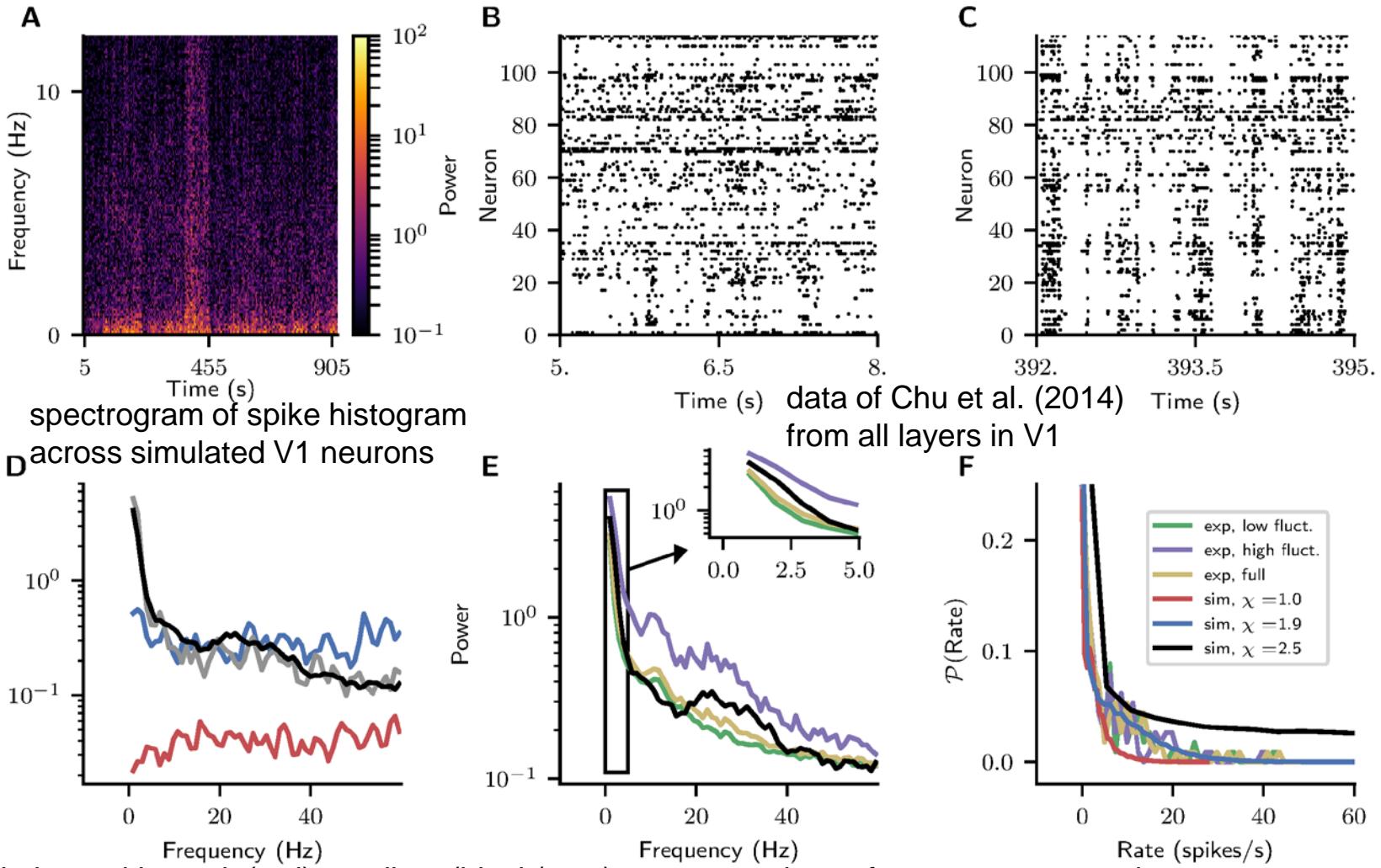
- well-known phenomenon of dynamical slowing near instability
- confirms metastability previously found in less detailed simulation studies Deco et al. 2009 PNAS, Cabral et al. 2011 NeuroImage; Deco and Jirsa 2012 J Neurosci; Cabral et al. 2014 Prog Neurobiol



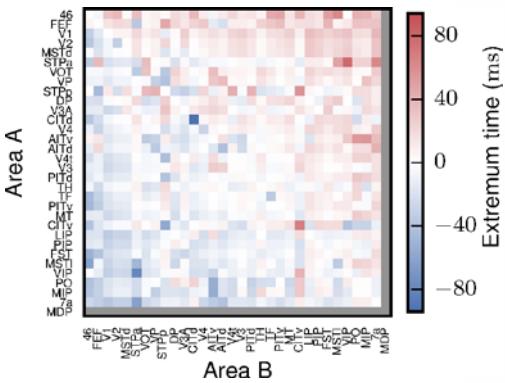
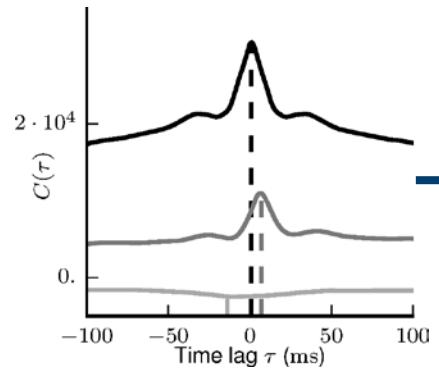
# GROUND STATE WITH MULTIPLE TIME SCALES



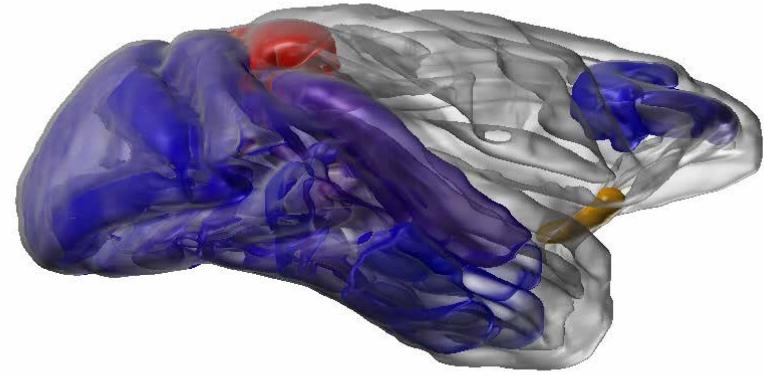
# V1 SPIKING STATISTICS



# TEMPORAL HIERARCHY

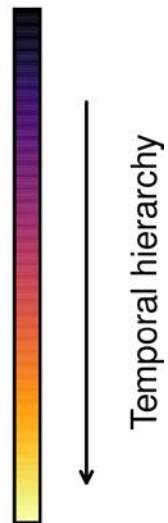
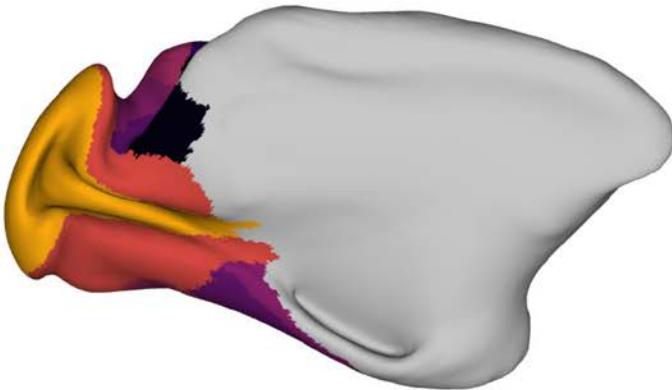
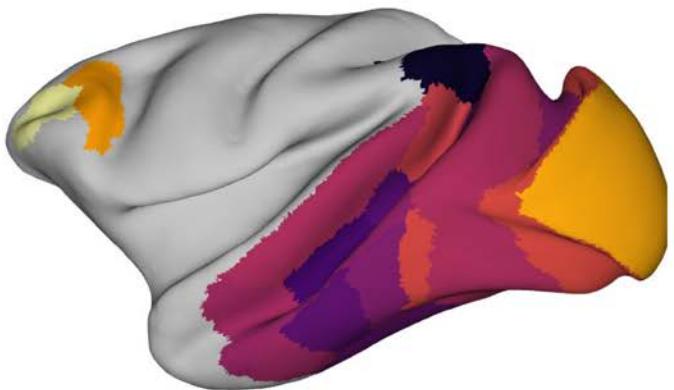


Lateral (left) view



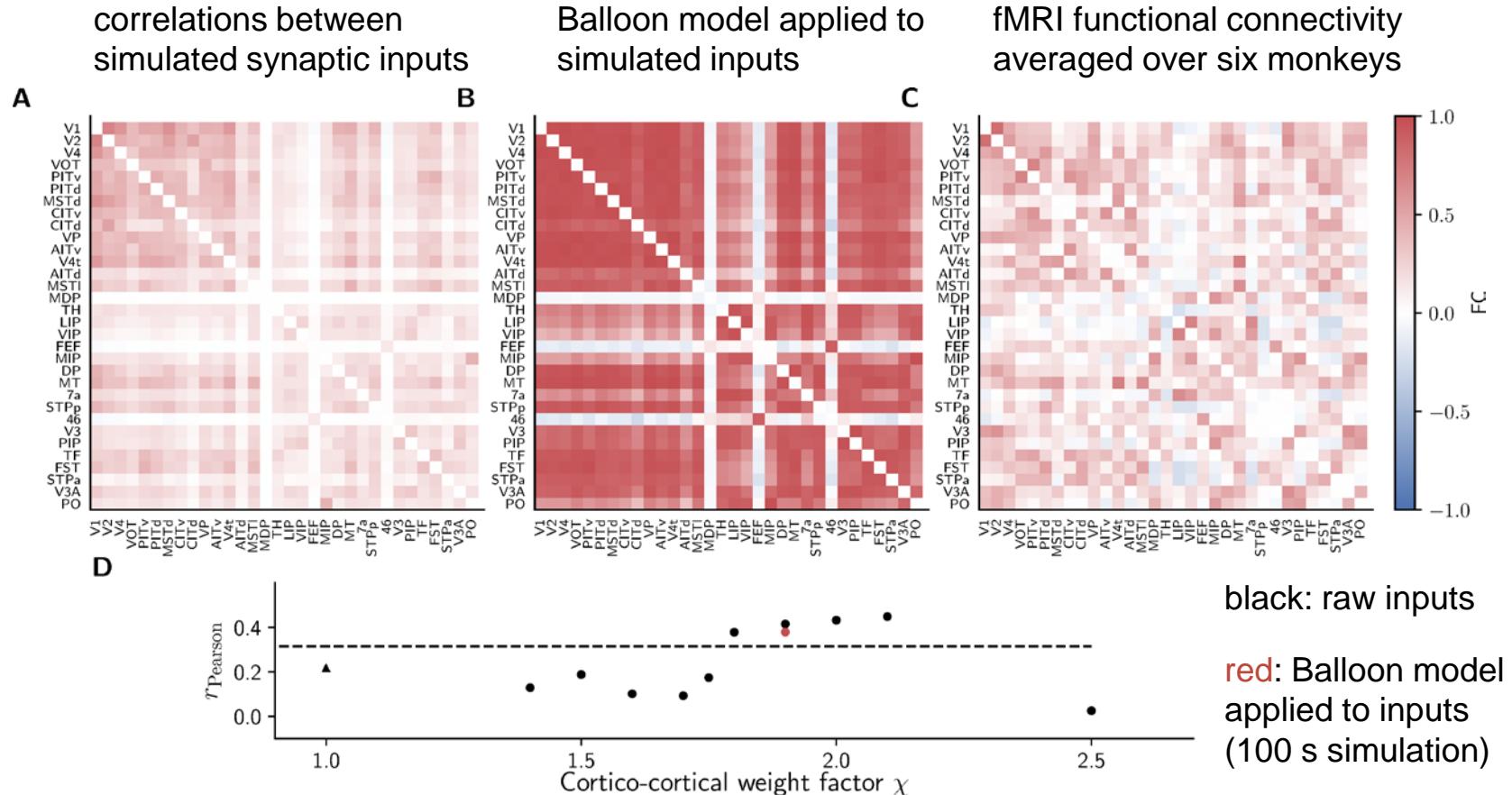
Nowke C, Schmidt M, van Albada SJ, Eppler JM,  
Bakker R, Diesmann M, Hentschel B, Kuhlen T  
(2013) *IEEE BioVis*

Medial (right) view



# FUNCTIONAL CONNECTIVITY

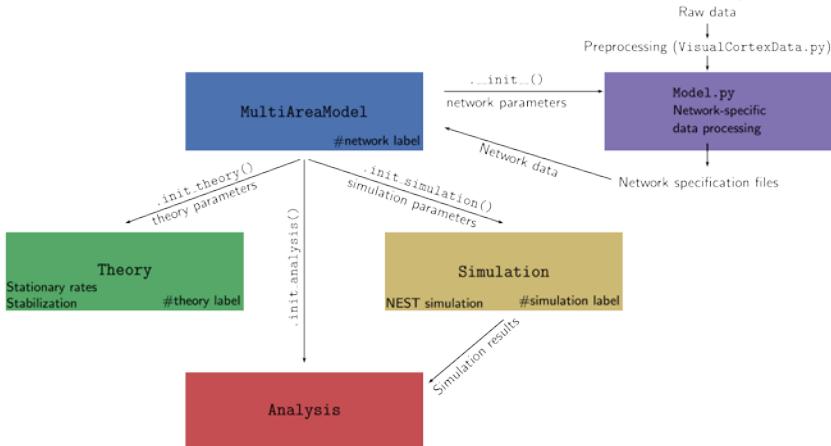
inter-area interactions in metastable state resemble experimental resting-state fMRI



Schmidt M, Bakker R, Shen K, Bezgin G, Diesmann M,  
van Albada SJ, *PLOS CB*, accepted

# IMPLEMENTATION AND HURDLES

- M ported SLI code to PyNEST → more readable; new implementation from scratch enabled clean, modular setup



Maximilian  
Schmidt



Alexander  
van Meegen

- M created private GitHub repository, added files; created README; extended documentation
- M requested permission to publish data; *not yet obtained* → start from processed data
- S & especially A reviewed code → knowledge about model transferred from M to A
- mean-field integration gave different results on different machines
  - problem with package versions? → use Conda
  - problem persisted → check OS environment, incl. GSL
    - problem persisted → create minimal reproducer
      - NEST turned out not to combine well with Conda → *under investigation*

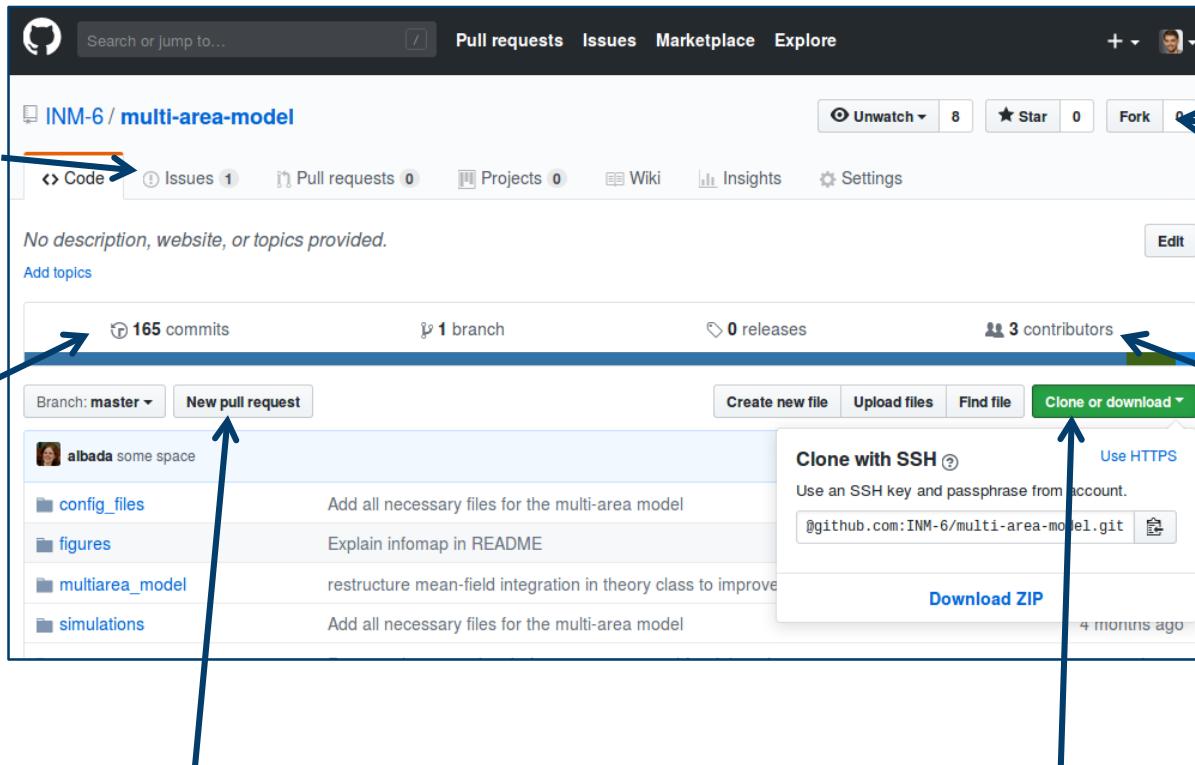
# GITHUB REPOSITORY

<https://github.com/INM-6/multi-area-model>

landing page

code is actively maintained,  
issues and solutions are tracked

code is alive,  
frequent new commits



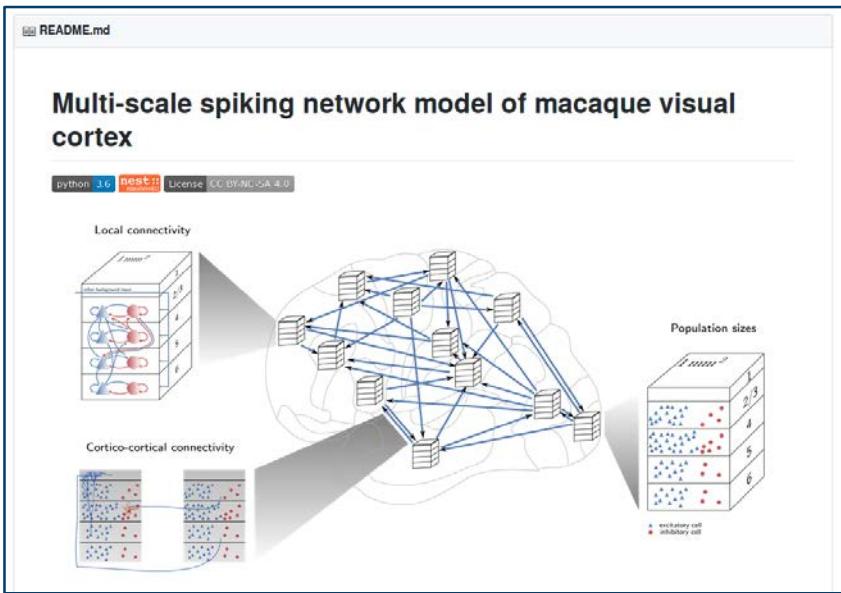
enhance the model via pull requests

clone to get local copy  
of the code

fork to start building your own work on top of the model

multiple contributors, detailed knowledge beyond main author

# EXTENSIVE README



README provides all technical information needed to instantiate and run the model

## instructions for running a simulation

### Running a simulation

A simple simulation can be run in the following way:

1. Define custom parameters `custom_params = ...` `custom_simulation_params = ...`
2. Instantiate the model class together with a simulation class instance.

```
M = MultiAreaModel(custom_params, simulation=True, sim_spec=custom_simulation_params)
```

3. Start the simulation.

```
M.simulation.simulate()
```

## software requirements

### Requirements

`python_dichash` (<https://github.com/INM-6/python-dichash>), `correlation_toolbox` (<https://github.com/INM-6/correlation-toolbox>), `pandas`, `numpy`, `nested_dict`, `matplotlib` (2.1.2), `scipy`, `NEST 2.14.0`

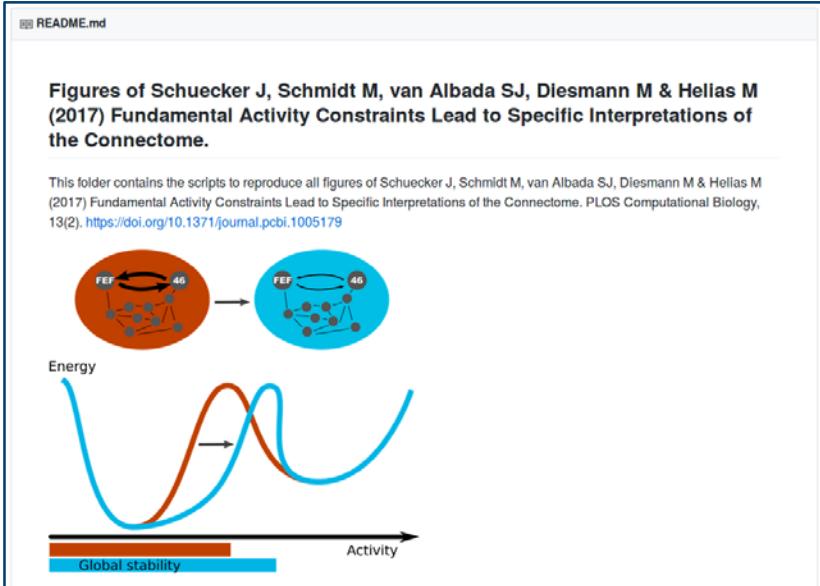
Optional: `seaborn`, `Sumatra`

To install the required packages with pip, execute:

```
pip install -r requirements.txt
```

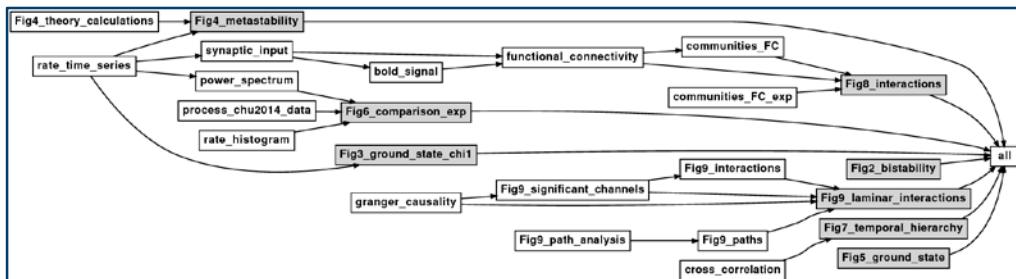
Note that NEST needs to be installed separately, see <http://www.nest-simulator.org/installation/>.

# REPRODUCING FIGURES



additional README with specific instructions for each publication

**Snakemake\*-based workflow to reproduce figures of Schmidt et al. *PLOS CB* (accepted)**



code snippet from the above Snakemake file

```
rule Fig4_metastability:
    input:
        expand(os.path.join(DATA_DIR, '{simulation}', 'Analysis', 'rate_time_series_full', 'rate_time_series_full_V1.npy'),
               simulation=SIM_LABELS['Fig4']),
        expand('Fig4_theory_data/results_{cc_weights_factor}.npy',
               cc_weights_factor=[1.0, 1.8, 1.9, 2., 2.1, 2.5])
    output:
        'Fig4_metastability.eps'
    shell:
        'python3 Fig4_metastability.py'
```

\* Köster J, Rahmann S (2012) *Bioinformatics*

# EXECUTABLE WORKFLOW

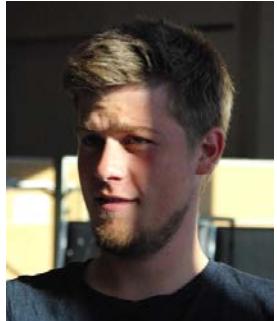
open-source material should:

- be correct
  - testing by main developer(s) and others  
great opportunity for knowledge transfer
- enable testing dependence on underlying data
  - include raw data; obtain permission early
- trace entire workflow from model construction to execution and visualization
  - include figure scripts; Snakemake removes manual steps
- be easy to understand and use
  - readable code; modularity; documentation; open-source packages
- fully specify package versions
  - Sumatra; Conda
- be provenance-tracked; enable continued development
  - Sumatra; GitHub

# ACKNOWLEDGMENTS

## Jülich

modeling,  
theory



Maximilian  
Schmidt

testing



Alexander  
van Meegen

anatomical  
data



Rembrandt  
Bakker



Jannis  
Schuecker

theory



Moritz  
Helias

performance  
measurements



Johanna  
Senk



Markus  
Diesmann

fMRI data



Kelly Shen  
Baycrest, Toronto  
McGill, Montreal



anatomy



Claus Hilgetag  
UKE Hamburg

## Manchester (SpiNNaker)

Andrew Rowley  
Michael Hopkins  
Alan Stokes  
Dave Lester  
Steve Furber