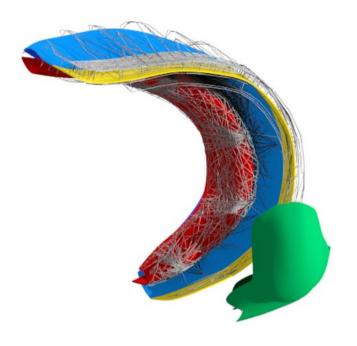




Creating an anatomically constrained model of the hippocampus



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Introduction

Outline

Motivation for anatomical modeling

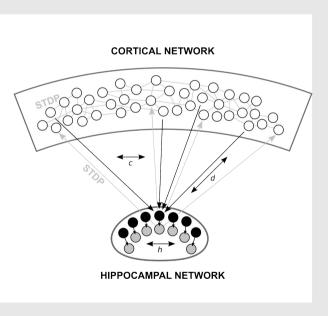
Parametric Anatomical Modeling (PAM)

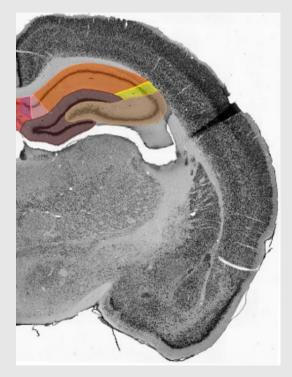
Projects based on PAM



Ways to test the validity

Does the real neural network implement the properties of the abstract one?



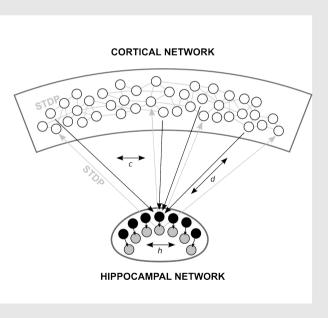


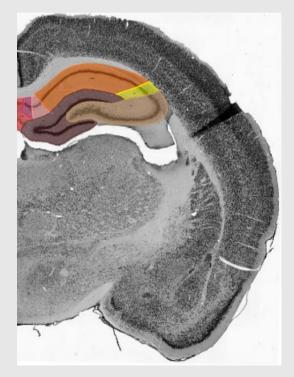
Rat Hippocampus



Ways to test the validity

 Test predictions of the abstract model in real networks



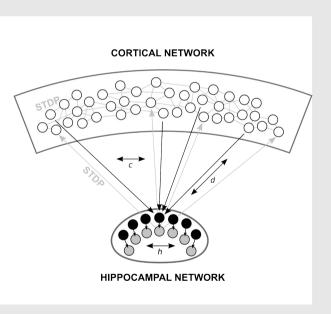


Rat Hippocampus

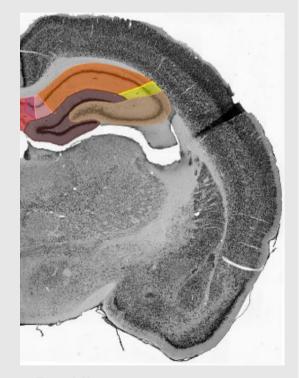


Ways to test the validity

 Test predictions of the abstract model in real networks



similar results may emerge from different mechanistic reasons

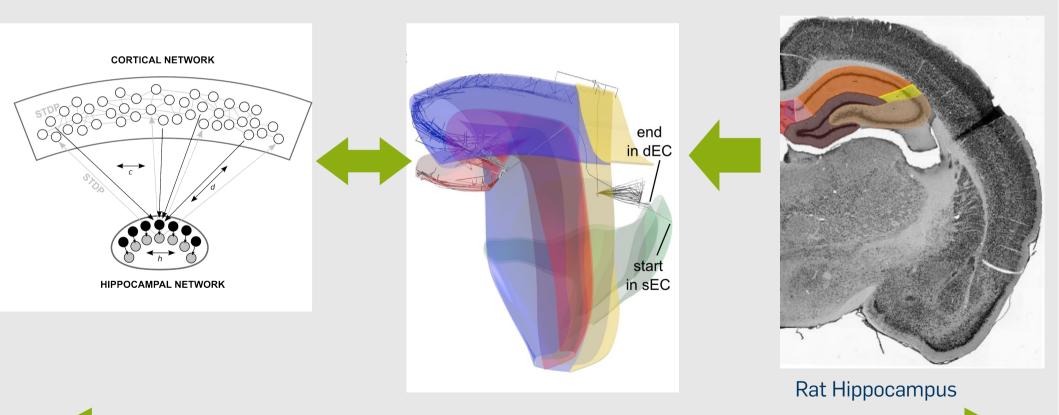


Rat Hippocampus



Ways to test the validity

- Test predictions of the abstract model in real networks
- Test a more detailed version of the target network





What does "more detail" mean?

- Connectivity
 - topology
 - connection spread

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Ropireddy and Ascoli. 2011

Temporal

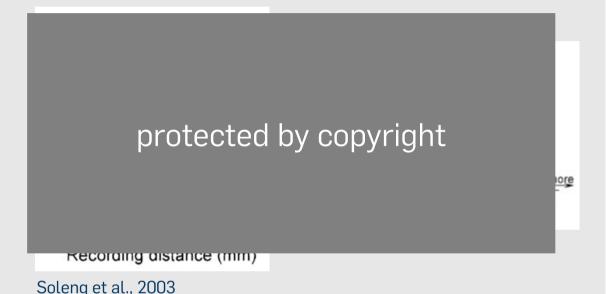
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Andersen et al., 2007



What does "more detail" mean?

- Connectivity
 - topology
 - connection spread
- Delays
 - Connection length
 - axon diameter



protected by copyright Axon diameter (µm) Waxman, 1980

Buzsaki and Mizuseki, 2014



Influence of structure on experimental results

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Blumberg, 1989

"A potential argument for the decreasing frequency and irregularity of hippocampal theta oscillations in mammals as brain size increases is that the hippocampus is a single cortical module and its growth is limited by the axon conduction delays. Pyramidal neurons of the CA3 region innervate a very large volume of the hippocampus; they connect distant peer neurons and require long axonal lengths and, consequently, have longer delays."

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brain weight (g)

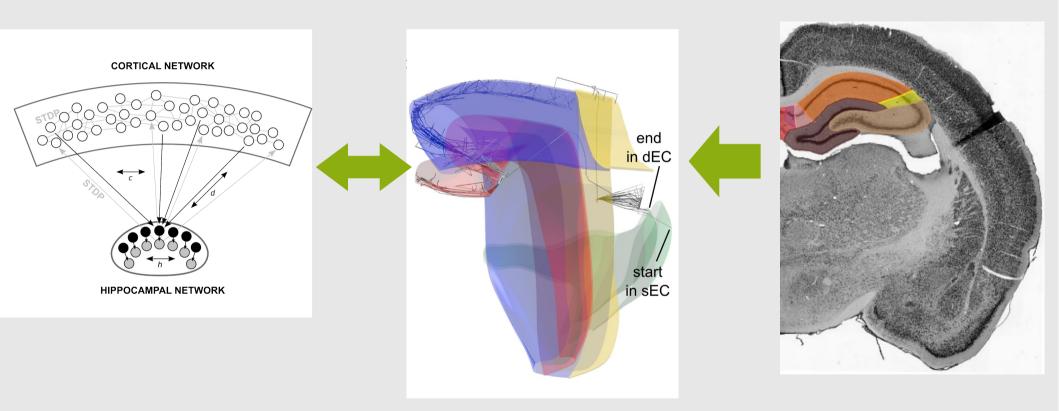
Buzsaki et al., 2014

Jakobs, 2014

Winson, 1972



Motivation for comparative studies

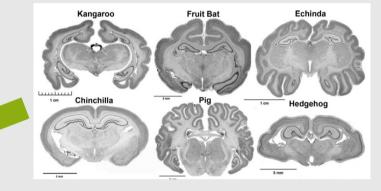


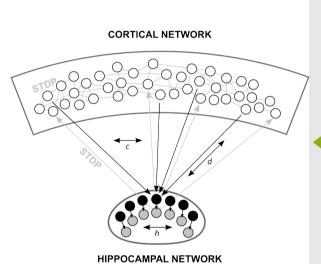
- connectivity
- delays

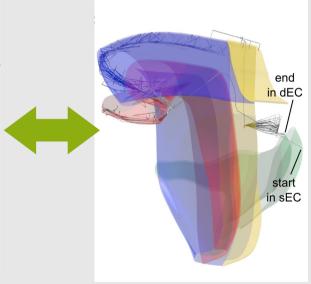


Motivation for comparative studies

Mammals

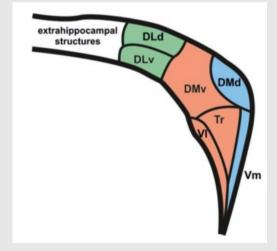








Birds





Motivation for comparative studies

Brain, Behavior and Evolution **Highlights and Perspectives on Evolutionary Neuroscience**

Brain Behav Evol 2014;83:1–8 DOI: 10.1159/000360152 Published online: February 28, 2014

Striedter et al., 2014

NSF Workshop Report: Discovering General Principles of Nervous System Organization by Comparing Brain Maps across Species

Georg F. Striedter^a T. Grant Belgard^b Chun-Chun Chen^c Fred P. Davis^d
Barbara L. Finlay^e Onur Güntürkün^f Melina E. Hale^g Julie A. Harris^h Erin E. Hechtⁱ
Patrick R. Hof^j Hans A. Hofmann^k Linda Z. Holland^l Andrew N. Iwaniuk^m Erich D. Jarvis^c
Harvey J. Kartenⁿ Paul S. Katz^o William B. Kristan^p Eduardo R. Macagno^q Partha P. Mitra^r
Leonid L. Moroz^s Todd M. Preuss^t Clifton W. Ragsdale^u Chet C. Sherwood^v
Charles F. Stevens^w Maik C. Stüttgen^f Tadaharu Tsumoto^x Walter Wilczynski^o





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PHYSICS of LIFE reviews

Physics of Life Reviews 11 (2014) 329-364

www.elsevier.com/locate/plrev

Review

Toward a computational framework for cognitive biology:
Unifying approaches from cognitive neuroscience and comparative cognition

Fitsch 2014

W. Tecumseh Fitch

Neuro View

Hale 2014

Mapping Circuits beyond the Models: Integrating Connectomics and Comparative Neuroscience

Manns and Eichenbaum, 2006

HIPPOCAMPUS 16:795-808 (2006)

Evolution of Declarative Memory

Joseph R. Manns* and Howard Eichenbaum

ABSTRACT: The present review considers research on the hippocampus and related areas from humans and experimental animals and makes three main points. First, many of the anatomical details of the hippocampus and adjacent cortical areas in the parahippocampal region are conserved across mammals. Second, the functional role of these areas in declarative memory is also conserved across species. Third, an evolutionary approach will be key to understanding exactly how the local circuitry of the hippocampus and parahippocampal region supports declarative memory. To highlight the utility of this approach, a schematic model is de-

The evolution of episodic memory

Timothy A. Allen and Norbert J. Fortin¹

regions responsible for episodic memory in humans have anatomical and functional homologs in other species. We propose that episodic memory capacity depends on a fundamental neural circuit that is similar across mammalian and avian species, suggesting that protoepisodic memory systems exist across amniotes and, possibly, all vertebrates. The implication is that episodic memory in diverse species may primarily be due to a shared underlying neural ancestry, rather than the result of evolutionary convergence. We also discuss

Allen and Fortin, 2013

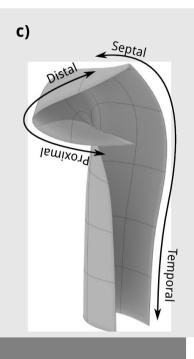


How to use computational models in comparative neuroscience?

- How to create anatomically motivated neural networks?
- How to compare artificial neural networks derived from various species?



Connections in the brain



Main Problem: the formal description of

- mappings on local and global axes
- nonlinear relationships between topological projections and connection distance

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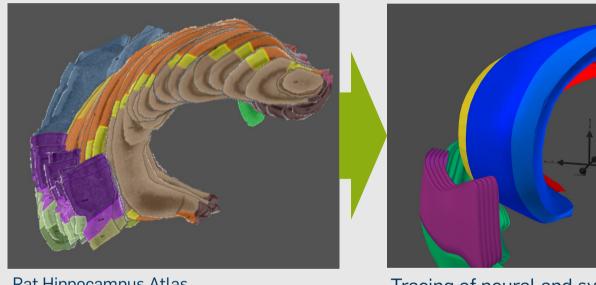
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van Strien et al., 2009



Main ideas

1. Recreate neural layers in 3D



Rat Hippocampus Atlas

van Strien et al., 2009

Andersen et al. 2007

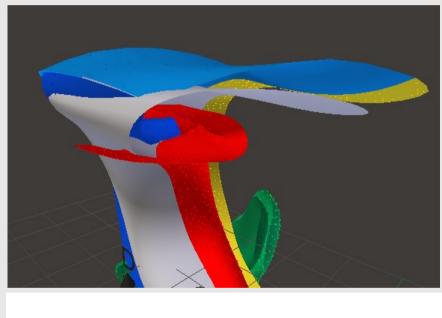
Allen Brain Atlas

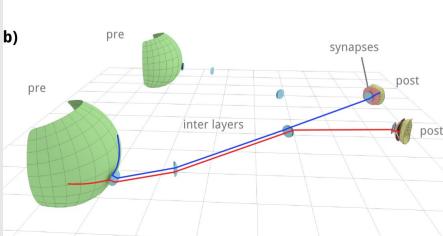
Tracing of neural and synaptic layers



Main ideas

2. Add synaptic and intermediate layers

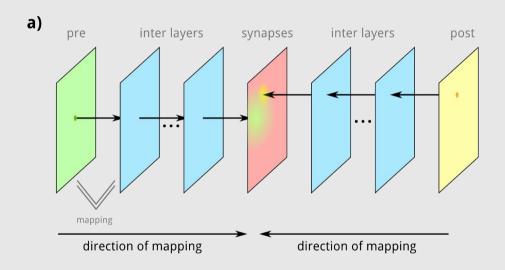


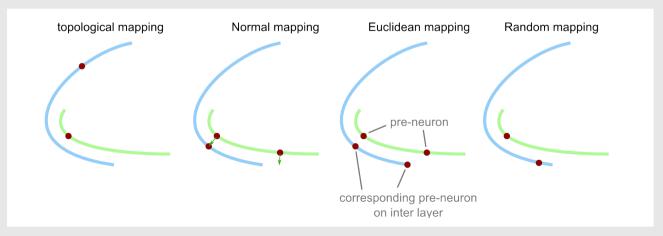




Main ideas

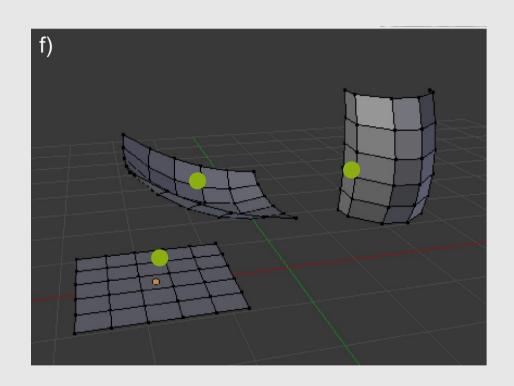
3. Create mappings between layers





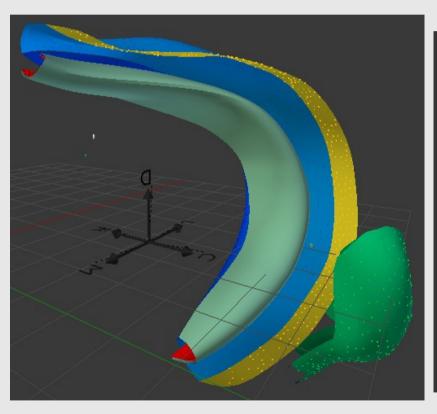


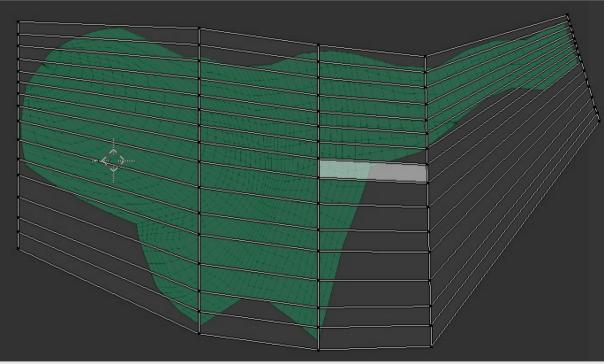
Topological mapping





UV mapping

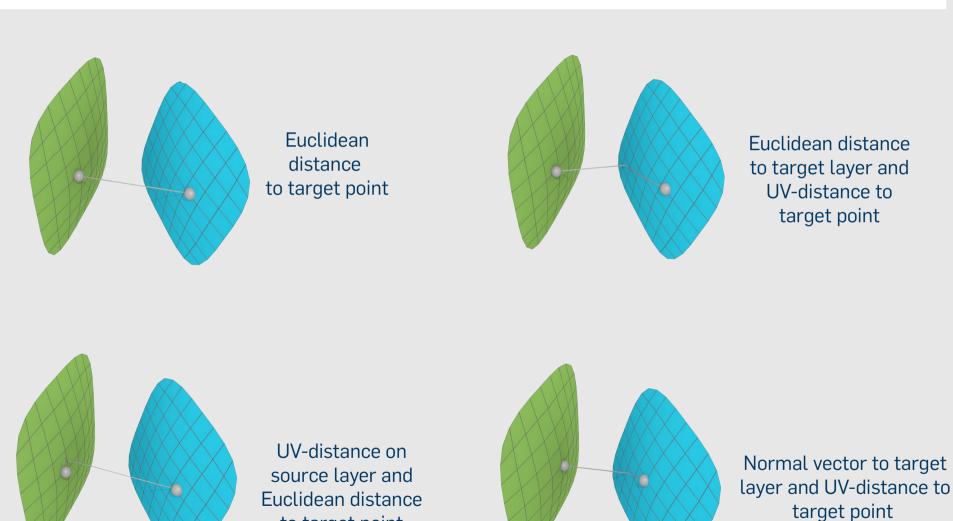






to target point

Distance calculation





Network calculation and export

- PAM calculates for a given set of neural layers and mappings connections and connection lengths between neurons
- data can be exported as Python binary (pickle-module) or as set of csv-files
- import for PyNEST available



Showcase



Advantages

- Mapping relations between neuronal layers on the mesoand macroscale level
- → PAM models can be easier compared with real neural networks
- lesion studies can be easier aligned with the model
- changes in the network can correspond to evolutionary changes of real networks
 - → size
 - number of neurons (=> connectivity)
 - → form
- comparatively low-dimensional representation of complex neural networks



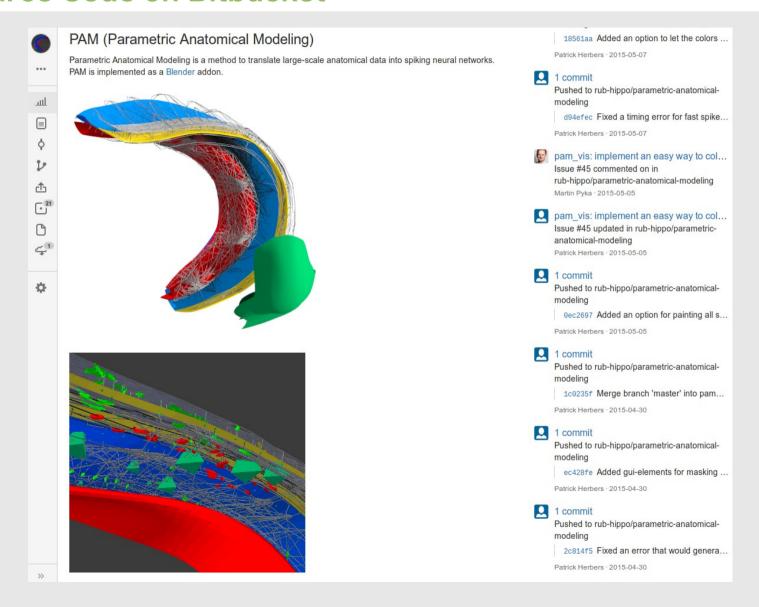
Connection to Experimentalists

 PAM could be used to document experimental findings (e.g. from tracer studies) in a formalized and parametric way

→ PAM could be helpful to let experimentalists and modelers work together on one model



Source Code on Bitbucket





Computational Constrains for more abstract models



Temporal Correlations between CA3 neurons



Temporal Correlations between CA3 neurons



Computational Constrains for more abstract models



Projection distances



Temporal Correlations between CA3 neurons



Computational Constrains for more abstract models



Comparative Computational Neuroscience

Pigeon and Rat Hippocampus



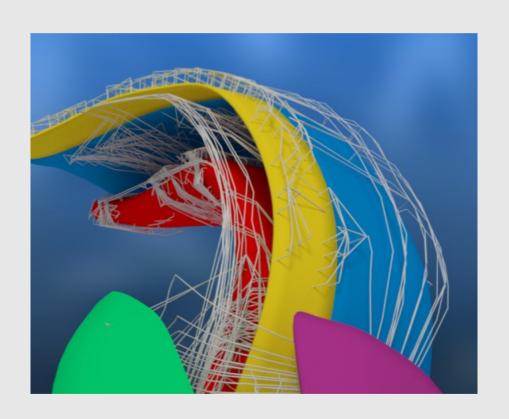
Comparative Computational Neuroscience

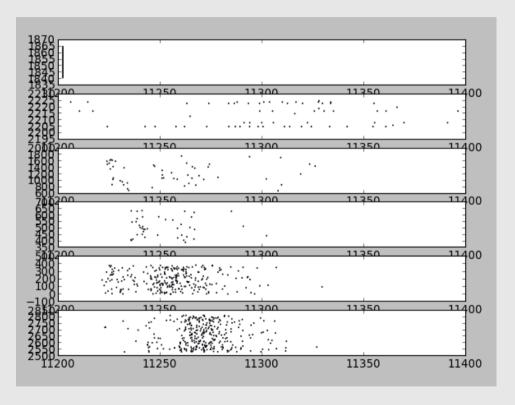
Pigeon and Rat Hippocampus



Temporal dynamics

How to understand, what is going on in the 3d hippocampus?







Temporal dynamics

Showcase



Temporal dynamics

Benefits

- compare spiking activity in the model with experimental data => theta traveling waves
- faster examination of the network in order to understand, why certain neurons spike
- colored spikes: understand why and when information are (not) combined
- in general: easier to compare with experimental data

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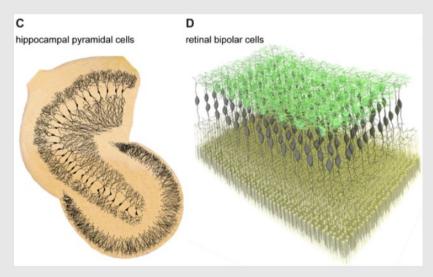


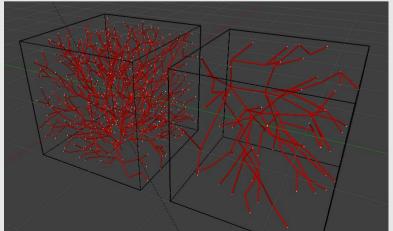
Ongoing projects

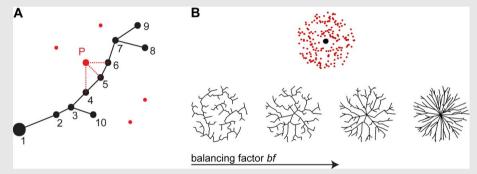
Dendritic morphology

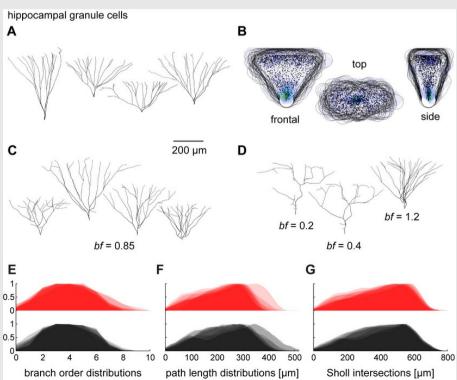
Implementation of the minimal spanning tree algorithm (TREE)

Toolbox, Cuntz et al., 2010)









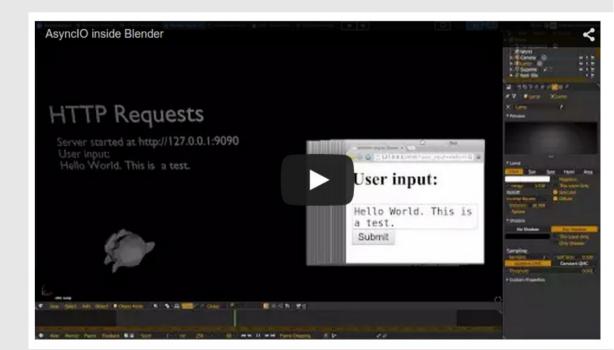


Planned projects

AsyncIO for Blender and parallel computing for HPC hardware

- Implementation of a standard event loop library for concurrency and asynchronous input/output
- A network simulation can then be started and analysed within Blender
- faster and more efficient interaction with a network

- Optimization of PAM for parallel computing
- use HPC for calculating networks of realistic sizes





Conclusion

- PAM allows to model spatial relations between different brain regions
 - Extends functionality of simulators like NEST, Brian, NEURON etc.
 - → Facilitates at one point in the future comparative computational neuroscience
- We need good data!
- NeuroML support is underway

Sourcecode is available on

https://bitbucket.org/rub-hippo/parametric-anatomical-modeling



Thank you for your attention!

