Topological explorations in neuroscience

Public Lecture

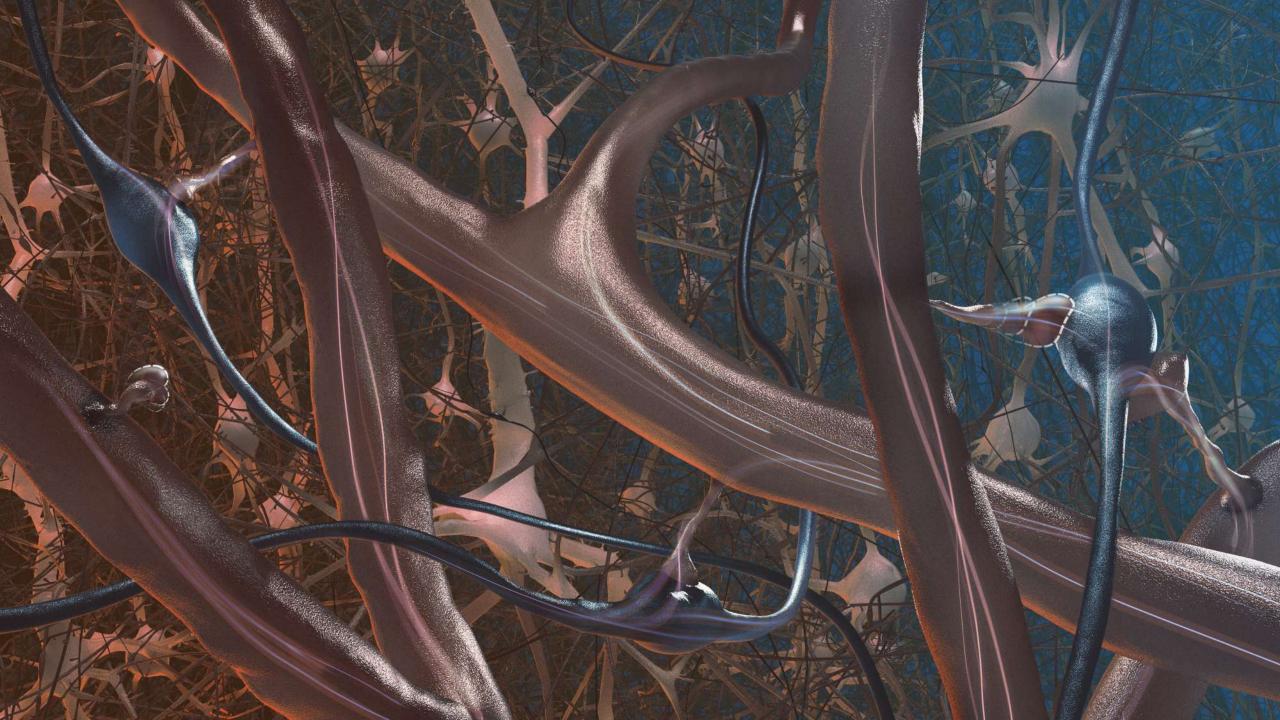
8th European Congress of Mathematicians

21 June 2021

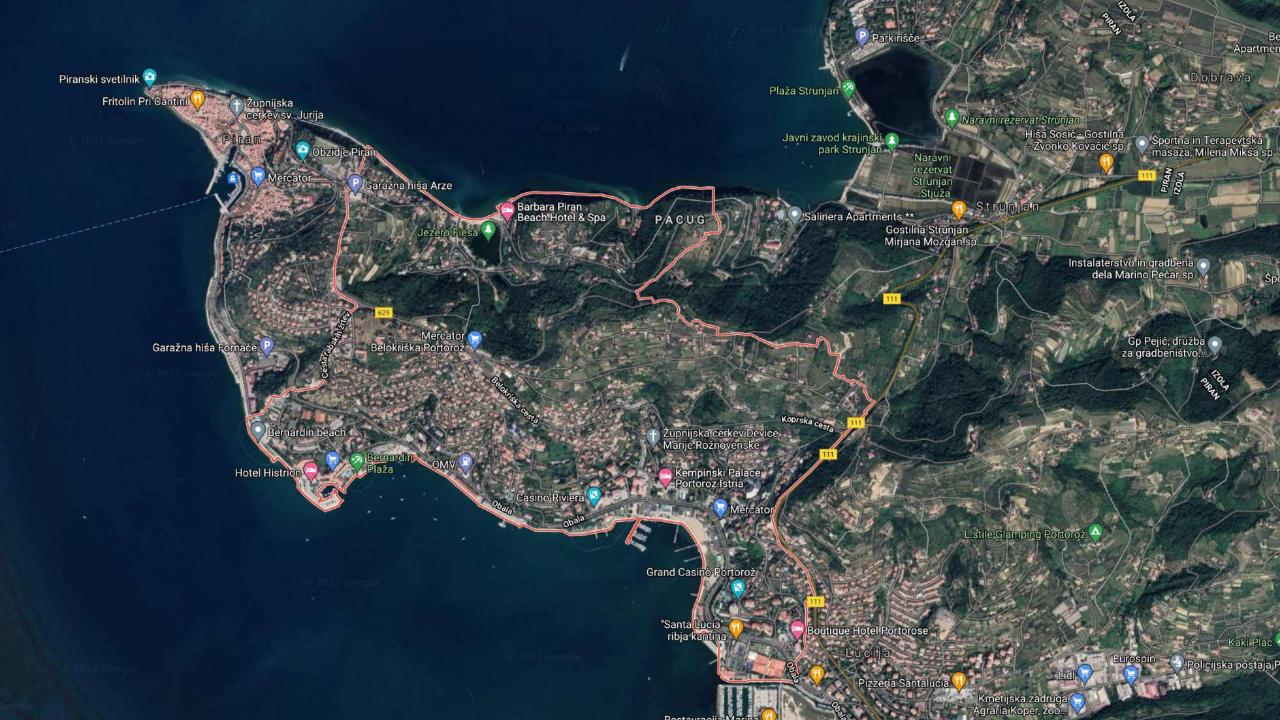
Blue Brain Project

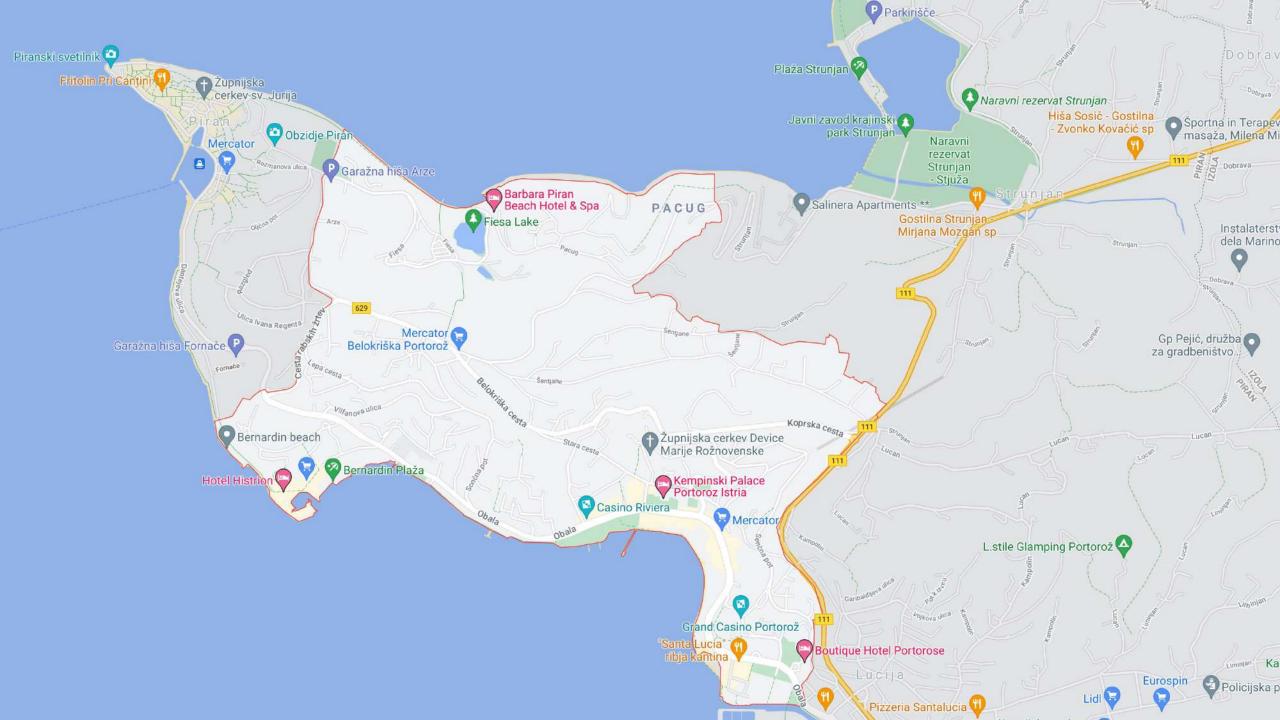


 $F_{p}(v) \otimes F_{q}(w)$ $(PoNoP) \square (QoNoQ) \xrightarrow{T_{N,N}} (P\otimes Q)o(NDN)o(((p; v_{\lambda}, -, v_{k}; p_{\lambda}, -p_{n})) (q; w_{\lambda}, -, v_{k}; p_{\lambda}, -p_{n})) \mapsto (poq; (v, w_{k}); j; (proqs)r, s)$ -> Bimod MÕN Fpoq (NDW) J⊗Q : FpN ---> FpN' b: Faw ----> Faw' $\rightarrow O_{p}$ Þq $a^{\#}: N \longrightarrow B_{o}N'_{o}B$ $f^{\#}: N \longrightarrow Q_{o}N'_{o}Q'$ $(P \otimes Q) \circ (V \square W) \circ (P \otimes Q) \circ (V \square W) \circ (P \otimes Q) \circ (P \otimes$ $\exists \mathcal{N} \Box \mathcal{M} \xrightarrow{a^{\#} \Box \mathcal{J}^{\#}} (\mathcal{P} \circ \mathcal{N} \circ \mathcal{P}) \Box (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{\mathsf{T}_{\mathcal{N}',\mathcal{N}'}} (\mathcal{P} \otimes \mathcal{Q}) \circ (\mathcal{P} \otimes \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q})} (\mathcal{P} \otimes \mathcal{Q}) \circ (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{Q})} (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{Q})} (\mathcal{P} \otimes \mathcal{Q}) \circ (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{Q})} (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{Q})} (\mathcal{Q} \circ \mathcal{N} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal{Q})} (\mathcal{Q} \circ \mathcal{Q}) \xrightarrow{(\mathcal{Q} \circ \mathcal$ $\frac{1}{\sqrt{2}} \left(\frac{\mathcal{P} \otimes \mathcal{Q}}{\mathcal{P} \otimes \mathcal{Q}} \right) \circ \left(\frac{\mathcal{P} \otimes \mathcal{Q}}{\mathcal{P} \otimes \mathcal{Q}} \right) \left(\frac{\mathcal{P} \otimes \mathcal{Q}}{\mathcal{P} \otimes \mathcal{Q}} \right) \circ \left(\frac{\mathcal{P} \otimes \mathcal{Q}}{\mathcal{P} \otimes \mathcal{Q}} \right) \right) \circ \left(\frac{\mathcal{P} \otimes \mathcal{Q}}{\mathcal{P} \otimes \mathcal{Q}} \right)$ $(T_{\mathcal{N}'\mathcal{W}'} \circ (\mathfrak{a}^{\#} \mathsf{D}\mathfrak{b}^{\#}))^{f} : F_{\mathcal{P}\otimes Q}(\mathcal{N} \mathsf{D}\mathcal{W}) \longrightarrow F_{\mathcal{P}\otimes Q}(\mathcal{N'} \mathsf{D}\mathcal{W'})$ \bigcirc $\sum_{g(g) \in Q \cup g \in L_Q(Q)} (\alpha' \otimes b') \circ (\alpha \otimes b) \stackrel{?}{=} (\alpha' \circ \alpha) \otimes (b' \circ b)$ $(\mathcal{P}\otimes\mathcal{Q})^{\circ_2}$ $(\mathcal{N}'\mathcal{D}\mathcal{W}')$ $(\mathcal{P}\otimes\mathcal{Q})^{\circ_2}$? (Q) Trimit (P&Q) (N' BW') (P&Q) L(P)&L(Q) P&Q $\mathcal{N} \xrightarrow{a^{\#}} F_{\mathcal{P}} \mathcal{N} \xrightarrow{a'} F_{\mathcal{P}} \mathcal{N}''$, Ah / LPOQ(POQ)









What is the Blue Brain Project??

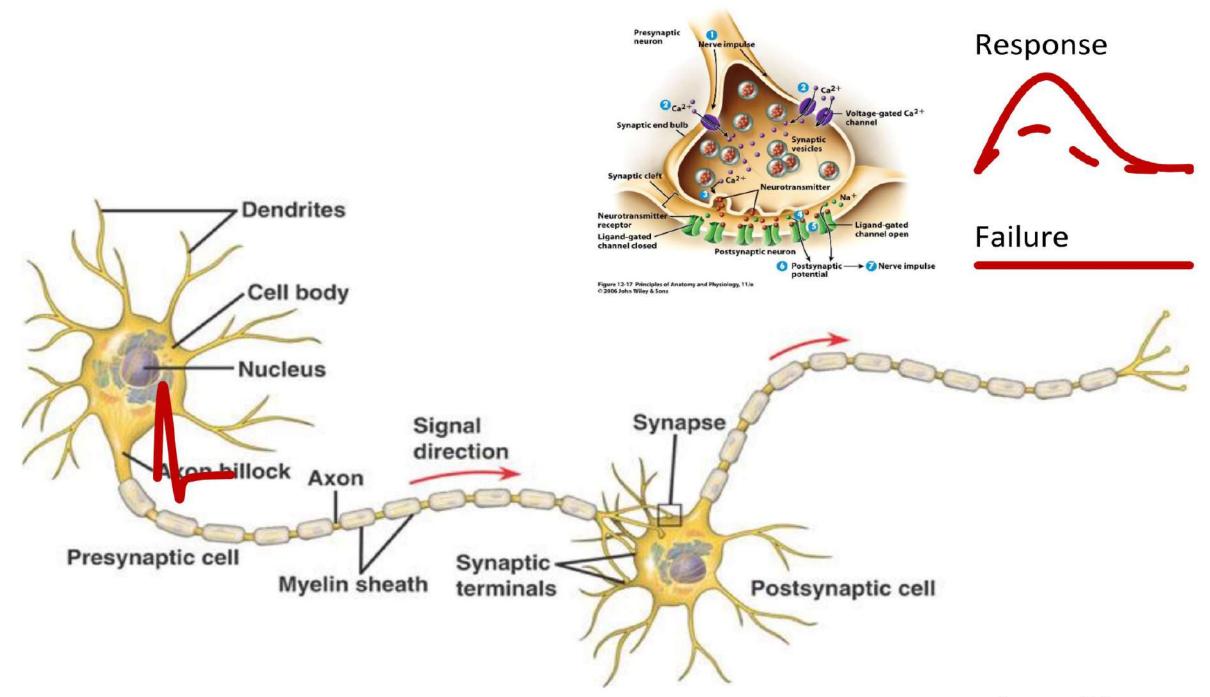


A.p

Rat



Human



From pmgbiology.com







Neuronal distribution in the rat somatosensory neocortex

Circuit: cxs1_v5.r0 Size: 31'346 neurons Target: mc2_Column Visualisation: 1'000 neurons Coloring: per layer 1 to 6 and synaptic class

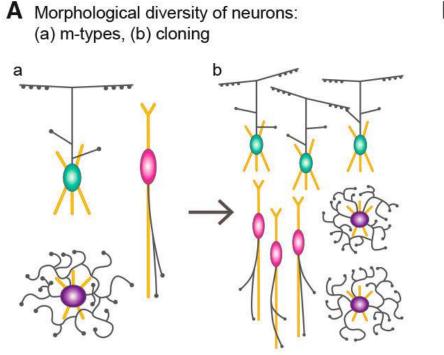
Vis. eng. & design: Nicolas Antille Scientific owners: Henry et al. Cell, 20 Excitatory samples

80% of neural cells are excitatory Inhibitory samples

-20% of neural cells are inhibitory

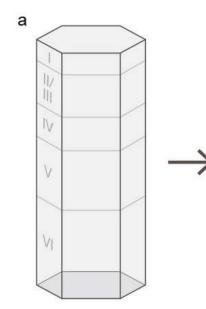
Blue Brain

Workflow: anatomy

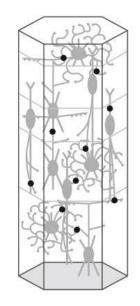


B Microcircuit anatomy: (a) Microcircuit dimensions, (b) m-type distribution, and morphology selection

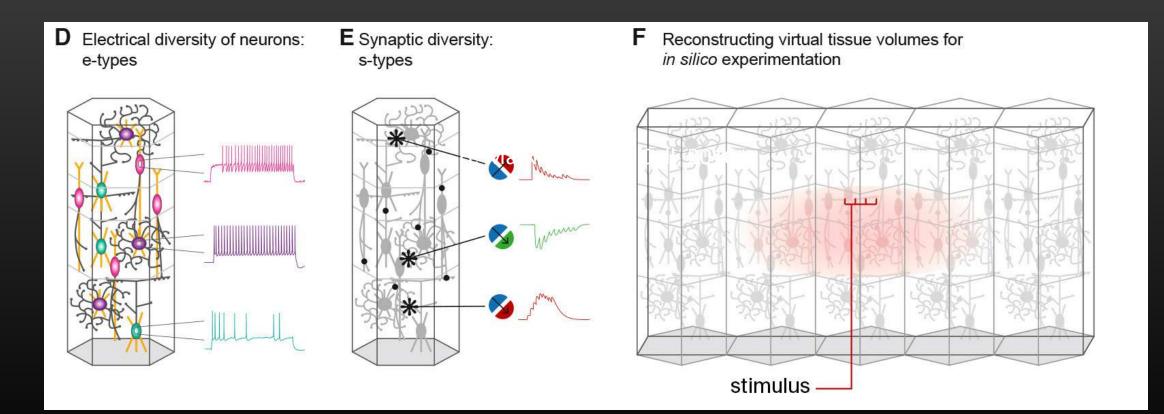
b



C Reconstructing microcircuit connectivity



Workflow: physiology



H. Markram et al., Cell, 2015

Study the emergent structural and functional properties of the microcircuit.

Study the emergent structural and functional properties of the microcircuit.

Study neurological disorders and neuroprostheses in silico.

Study the emergent structural and functional properties of the microcircuit.

Study neurological disorders and neuroprostheses in silico.

Reduce the need for animal testing in laboratory experiments.

Topological analysis of the microcircuit

Reimann et al. Frontiers in Computational Neuroscience, 2017.



Topology is...

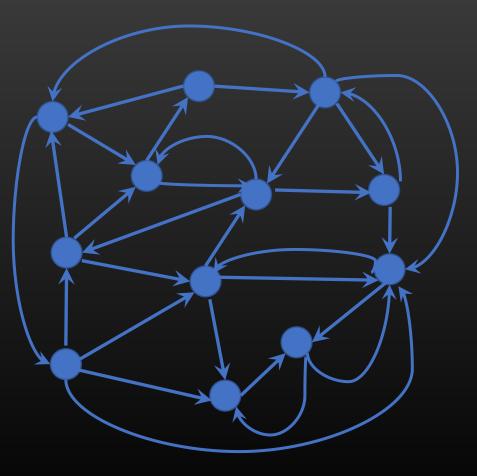
• the mathematics of shape;



Wikipedia, no license

Topology is...

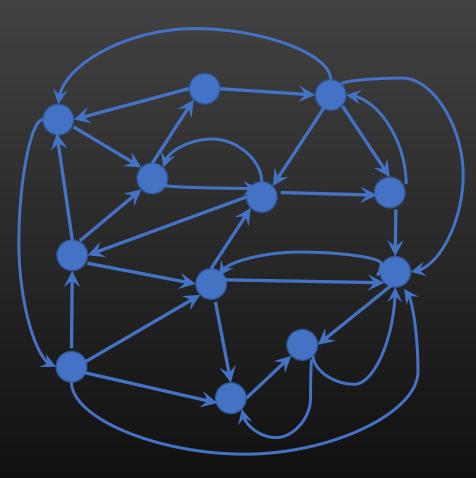
- the mathematics of shape;
- the mathematics of connectivity;



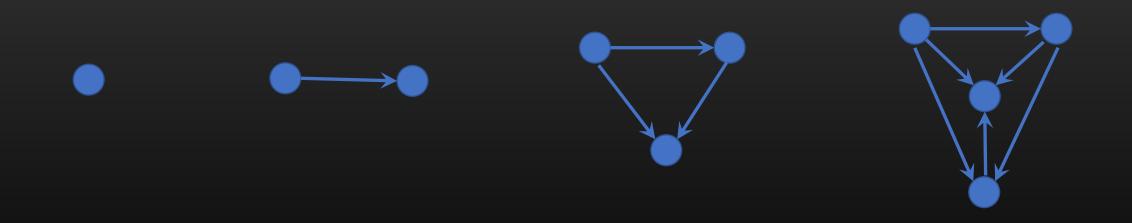
Topology is...

- the mathematics of shape;
- the mathematics of connectivity;
- the mathematics of emergence of global structure from local constraints.





• Analyze the huge network of directed connections among neurons in terms of much smaller significant subnetworks.

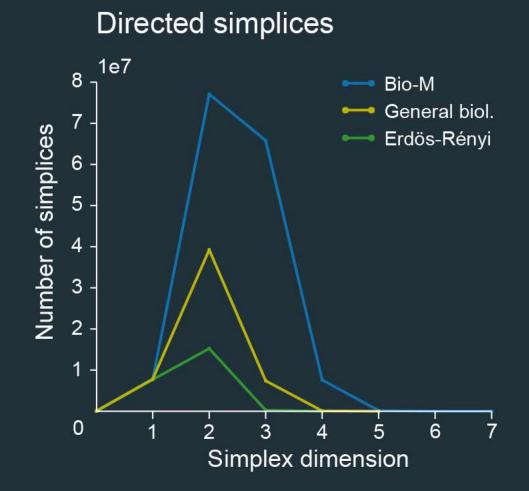


Directed simplices of dimensions 0, 1, 2, and 3

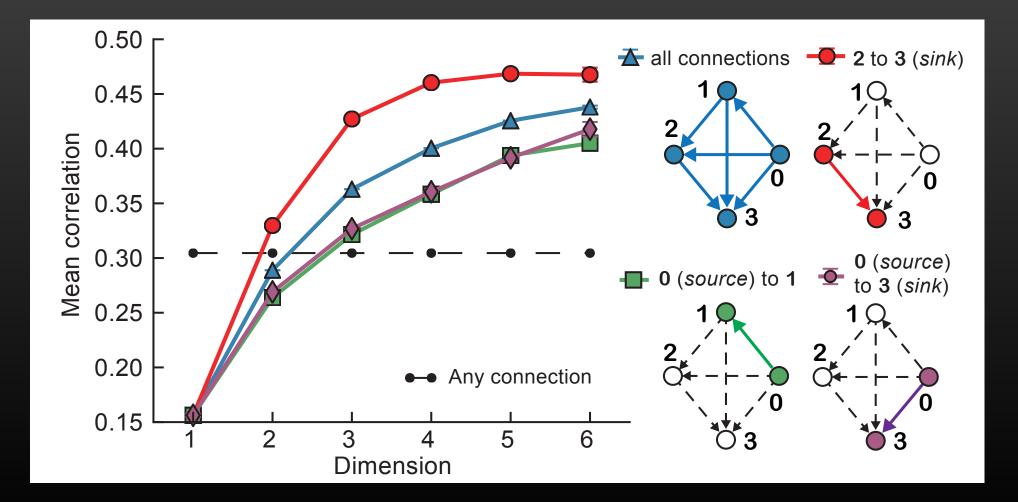
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- Analyze the huge network of directed connections among neurons in terms of much smaller significant subnetworks.
- The numbers of different types of significant subnetworks provide important local information about the whole network.
- Quantify how the significant subnetworks overlap in the larger network to obtain important global information.

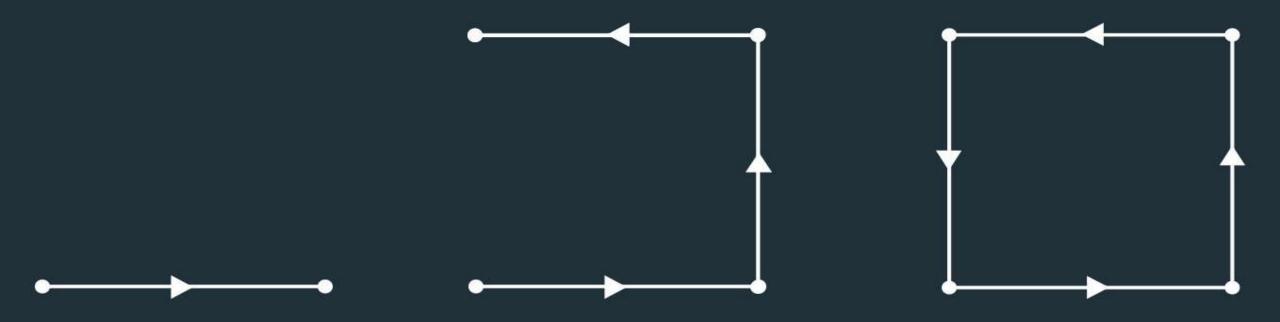
Measuring structure



The functional importance of simplices



The idea of a cavity

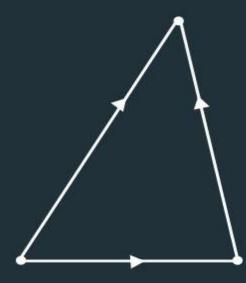


1 simplex

3 simplices...

a cavity made of 4 simplices

Higher dimensions for a cavity





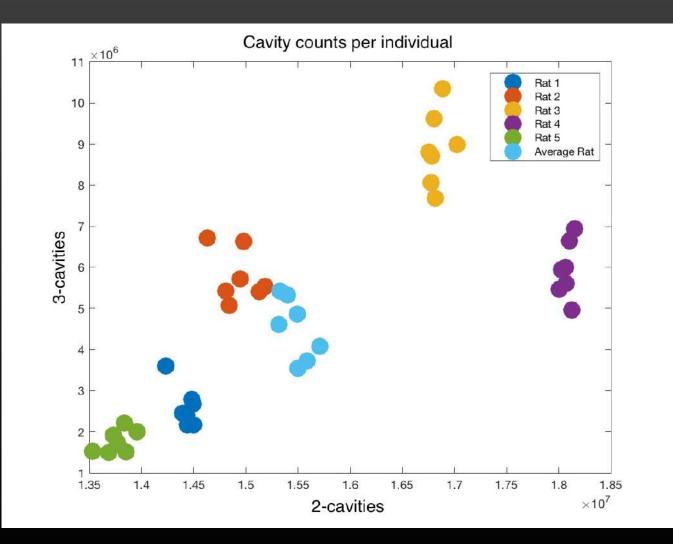


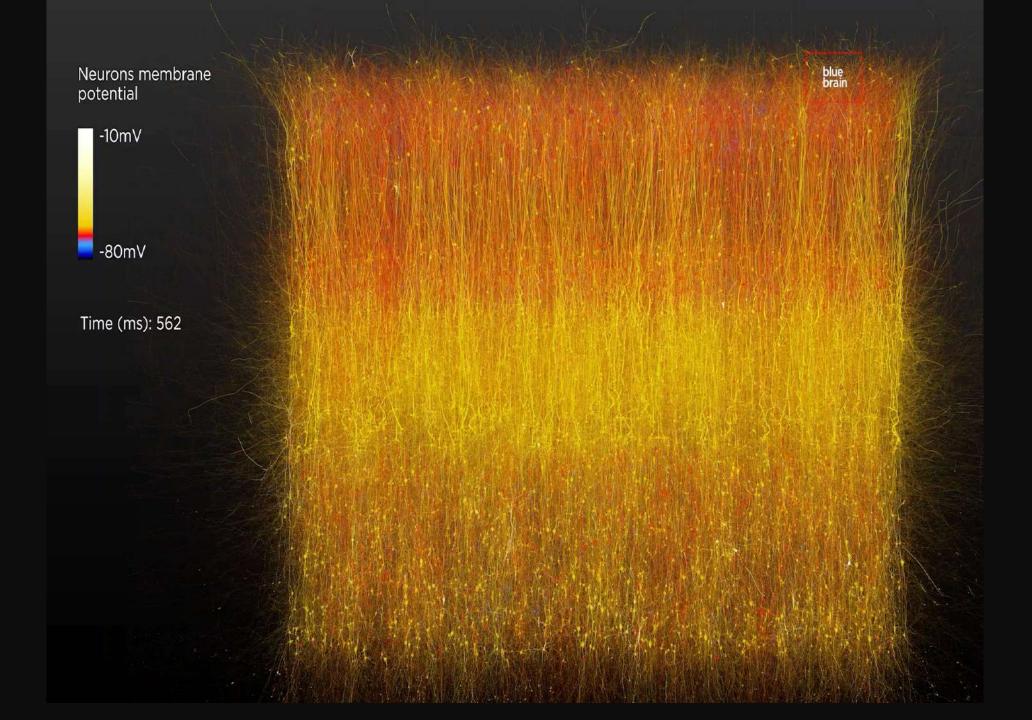
1 simplex

3 simplices...

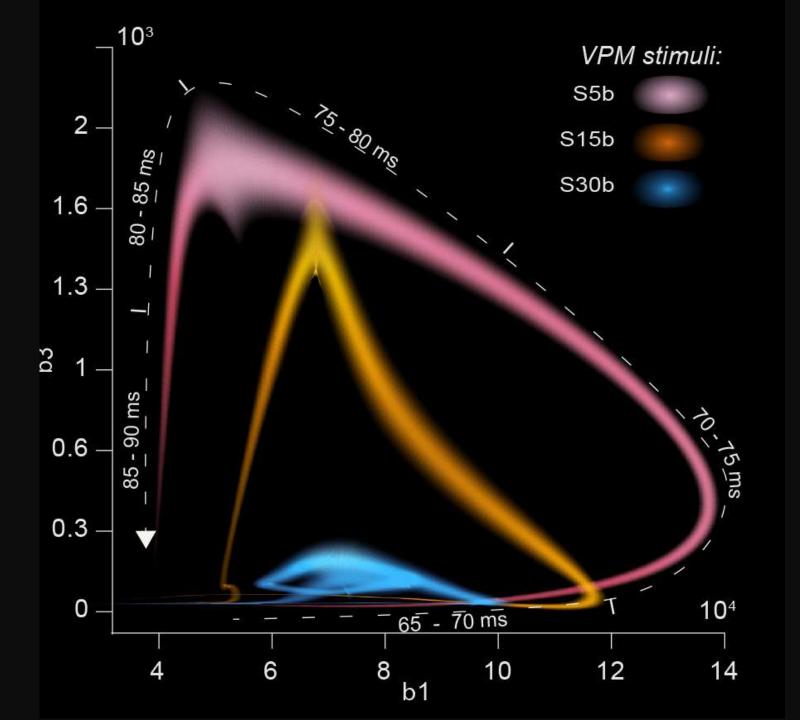
a cavity made of 8 simplices

Topology faithfully reflects biology





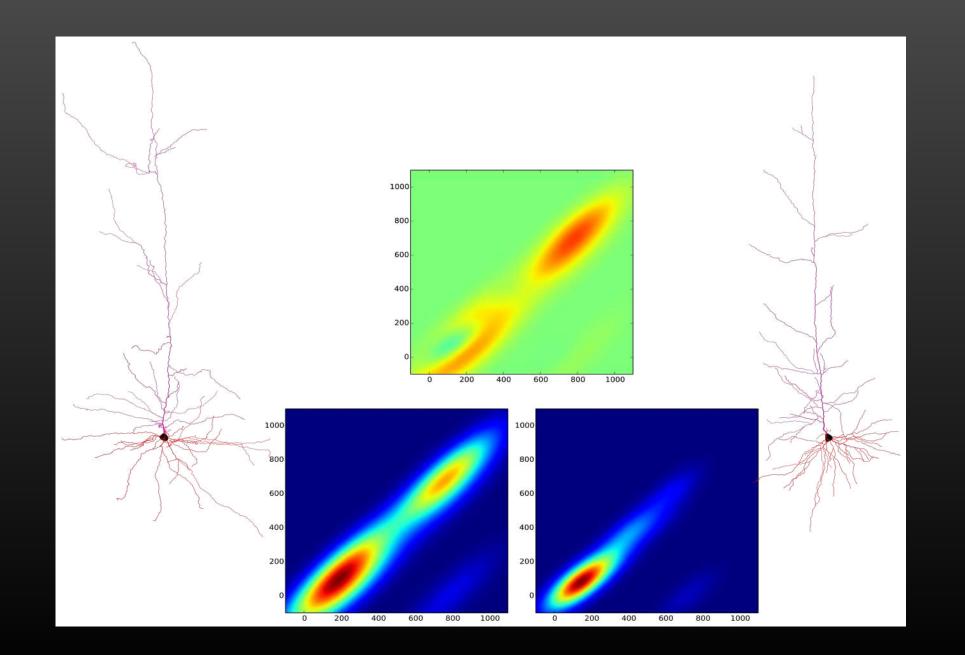


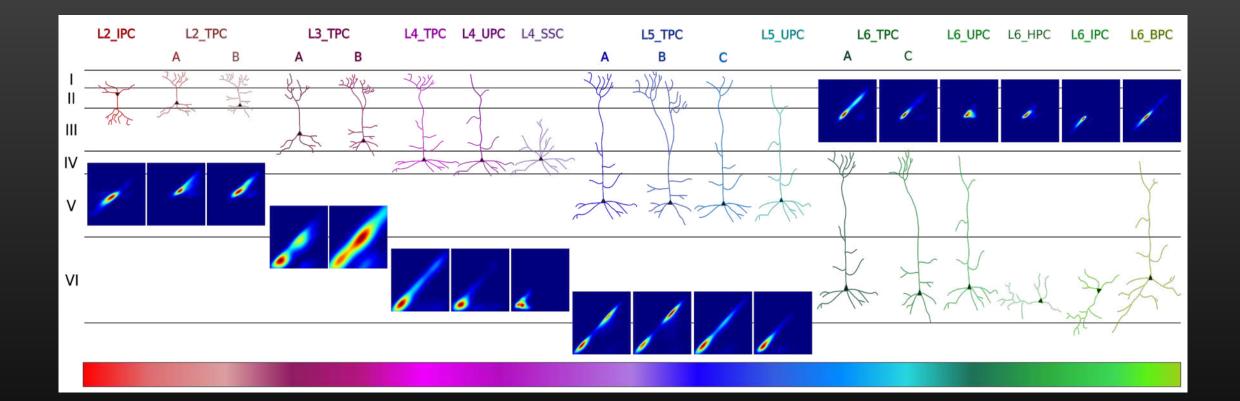


Classification of neuron morphologies

Y. Deitcher et al, Cerebral Cortex, 2017.L. Kanari et al, Neuroinformatics, 2018.L. Kanari et al, Cerebral Cortex, 2019.







Collaborators

- Stefania Ebli, Daniela Egas Santander, Adélie Garin, Celia Hacker, Nicolas Ninin, Martina Scolamiero (now KTH), Gard Spreemann, Katharine Turner (now ANU), and Dimitri Zaganidis (Laboratory for Topology and Neuroscience, EPFL)
- Nicolas Antille, Giuseppe Chindemi, Cyrille Favreau, Lida Kanari, Henry Markram, Taylor Newton, Max Nolte, and Michael Reimann (Blue Brain Project, EPFL)

- Dejan Govc, Ran Levi, Daniel Lütgehetmann, and Jason Smith (Aberdeen)
- Pawel Dlotko (Swansea)
- Rodrigo Perin (Laboratory of Neural Microcircuitry, EPFL)

Thank you!