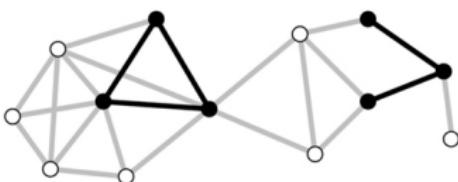


Modelling the nonrandom and dynamic structure of local cortical circuits

Felix Z. Hoffmann

Slides: bit.ly/xbcn18



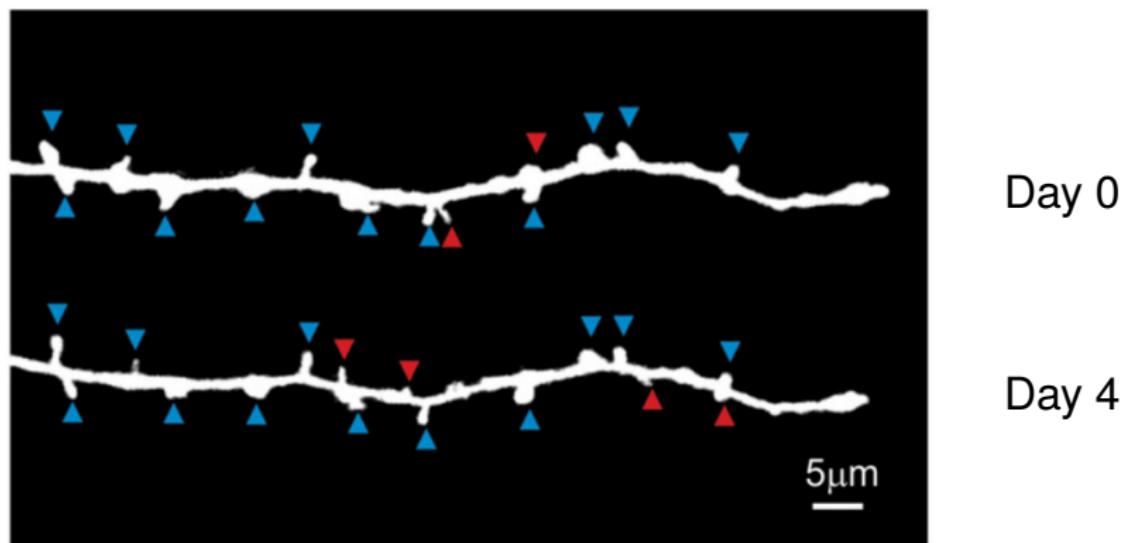
Bernstein Conference 2018 Satellite Workshop:
Adaptivity and Inhomogeneity in Neuronal Networks



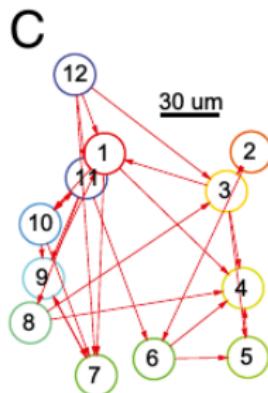
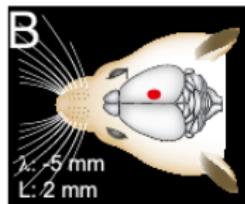
FIAS Frankfurt Institute
for Advanced Studies



The dynamic connectome

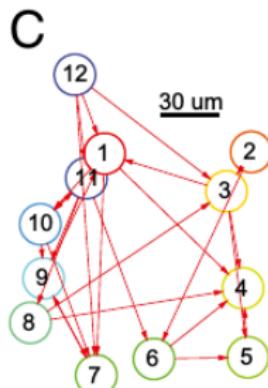
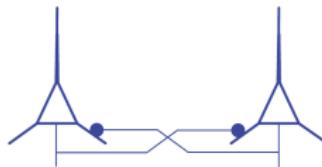
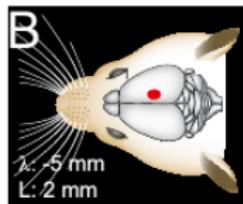


Robust nonrandom connectivity patterns



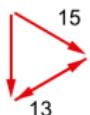
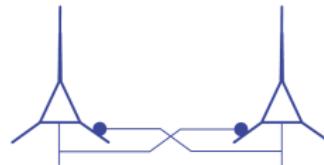
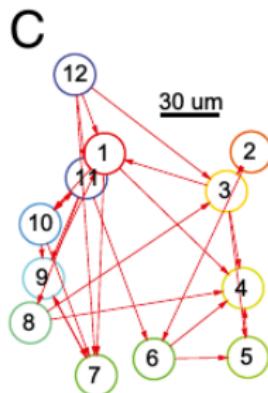
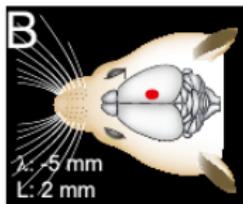
Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

Robust nonrandom connectivity patterns



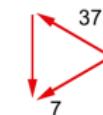
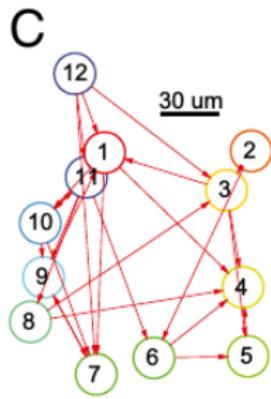
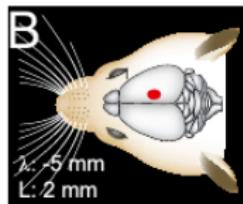
Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

Robust nonrandom connectivity patterns



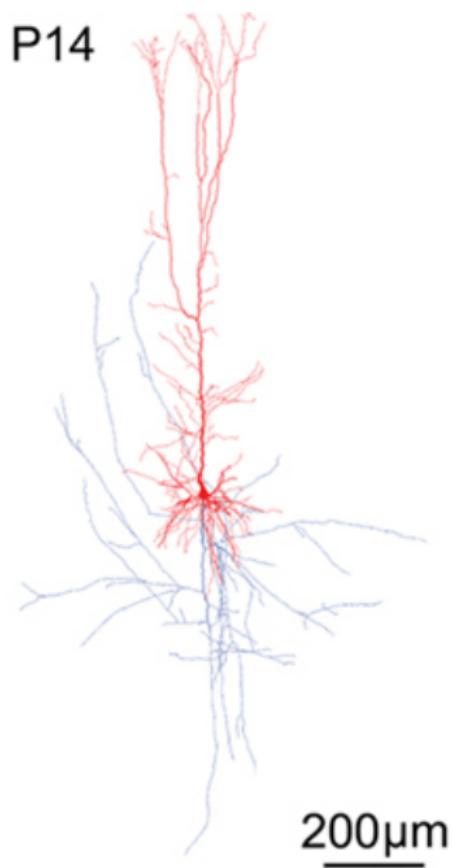
Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

Robust nonrandom connectivity patterns

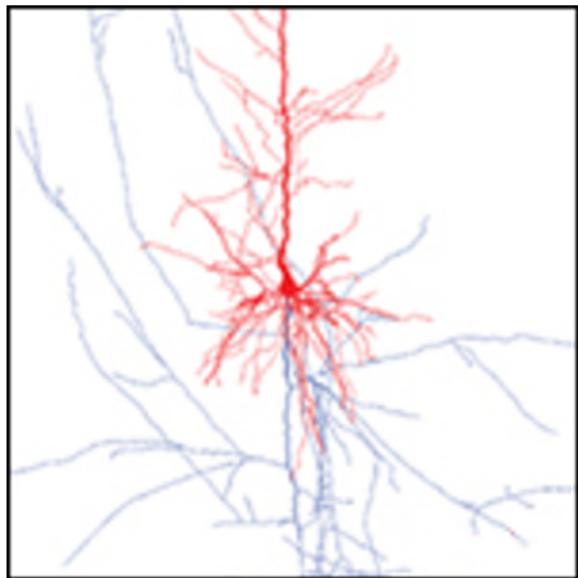


Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

Anisotropy in stereotypical axon morphology

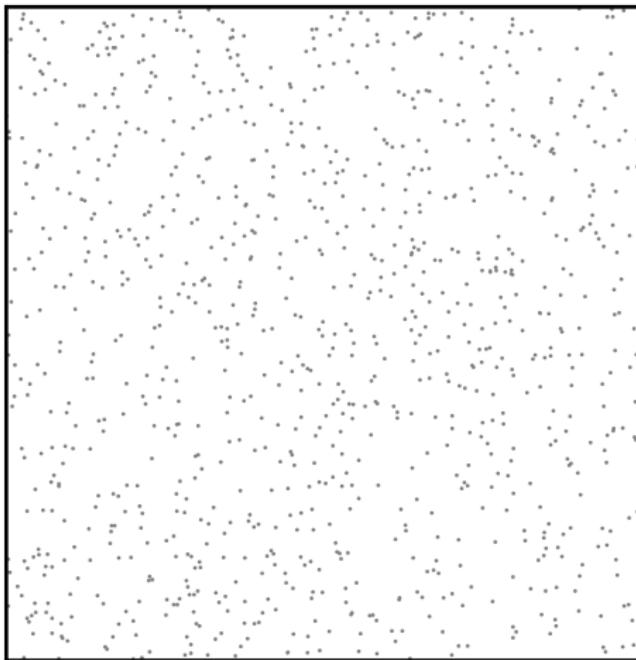


Anisotropy in stereotypical axon morphology



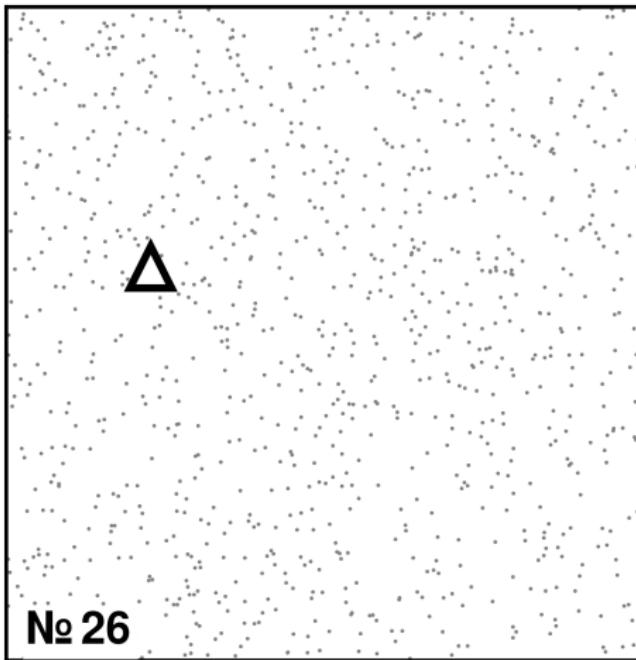
$200\mu\text{m}$

Anisotropic network model



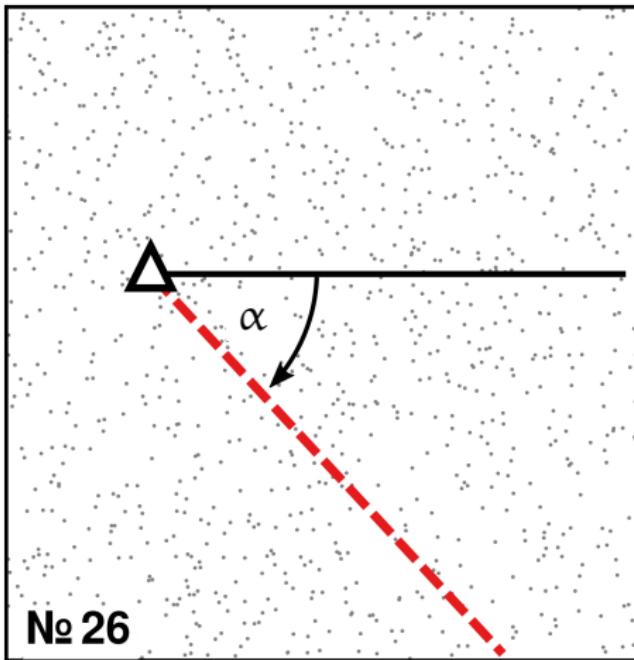
$N = 1000$

Anisotropic network model



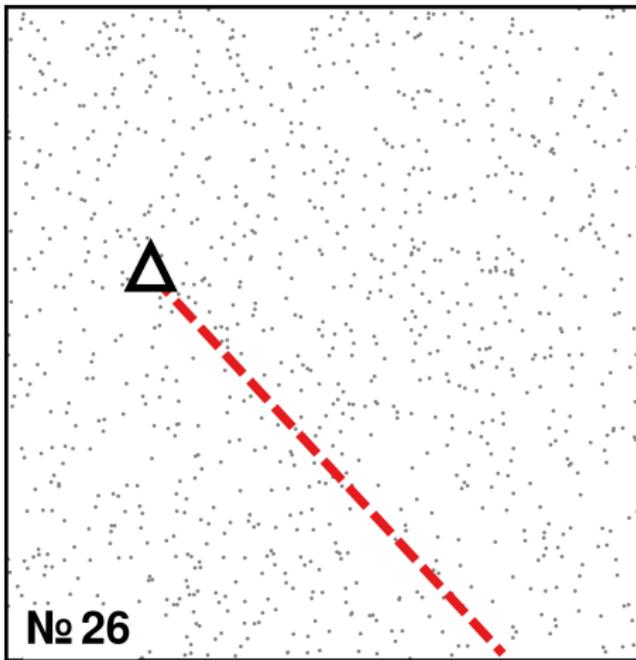
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Anisotropic network model



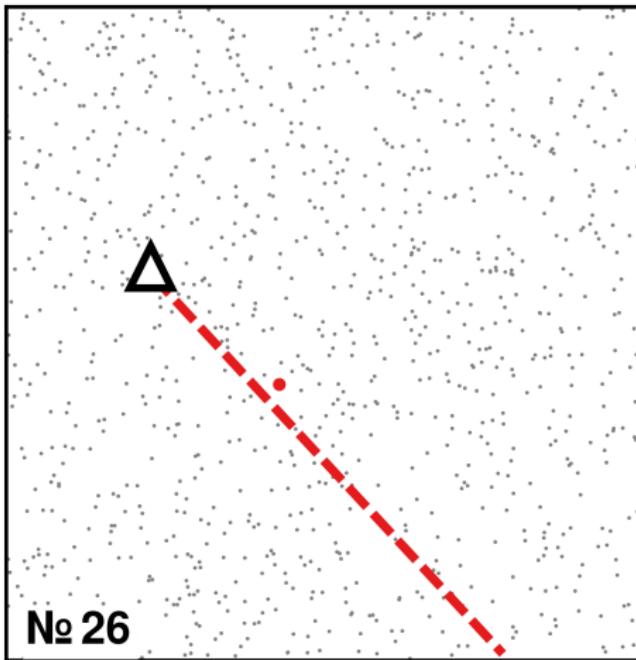
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Anisotropic network model



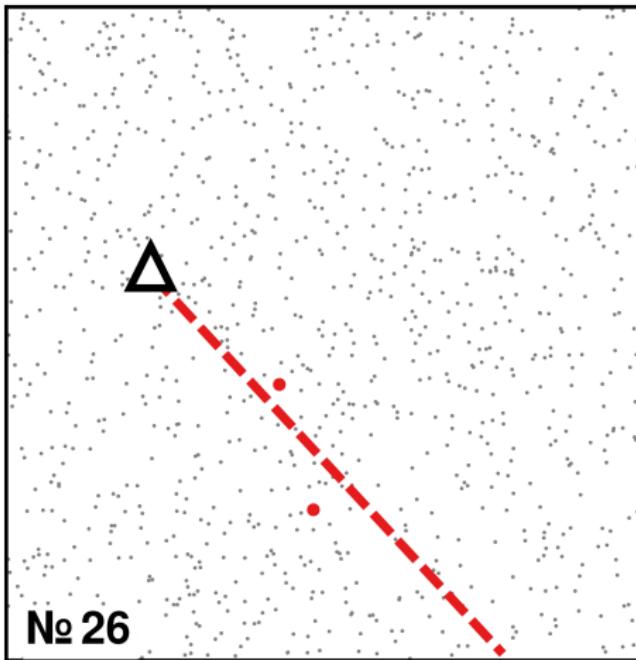
$N = 1000$

Anisotropic network model



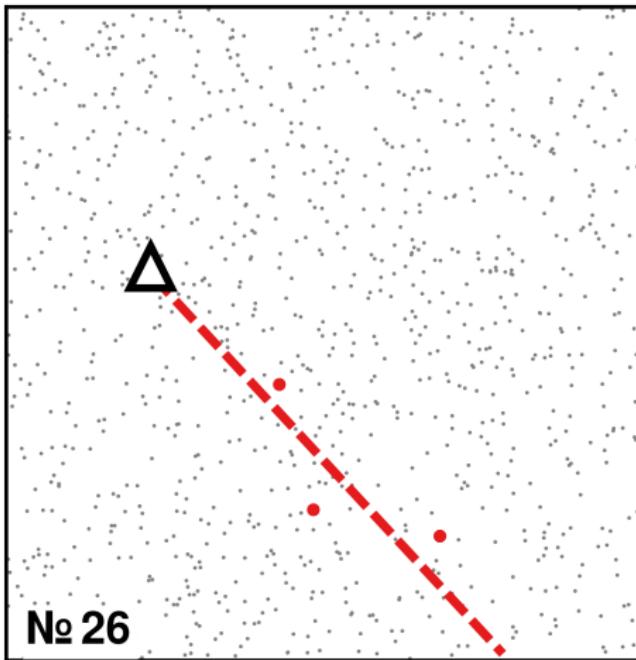
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Anisotropic network model



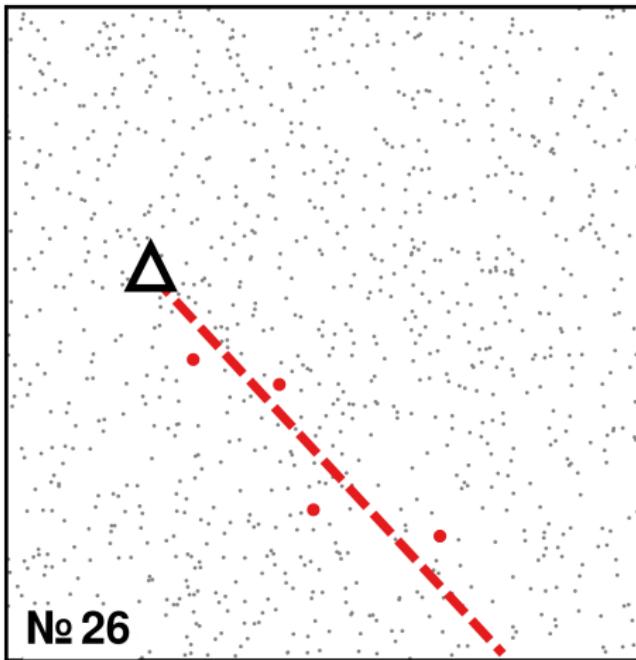
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Anisotropic network model



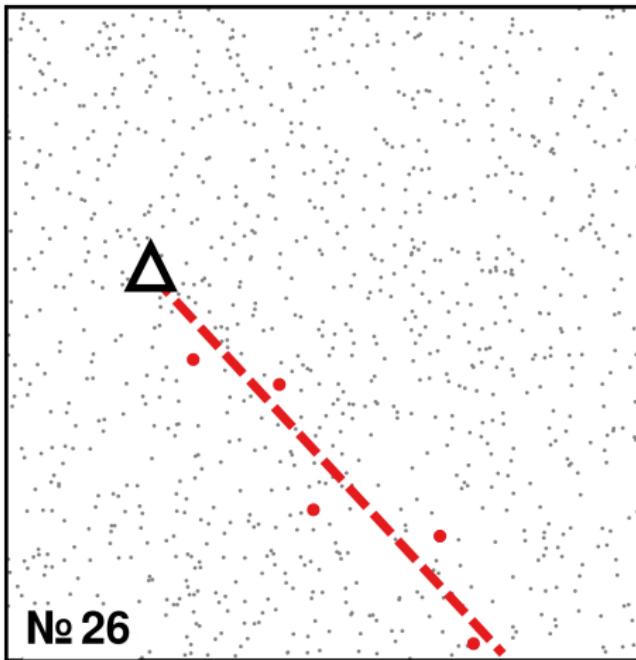
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Anisotropic network model



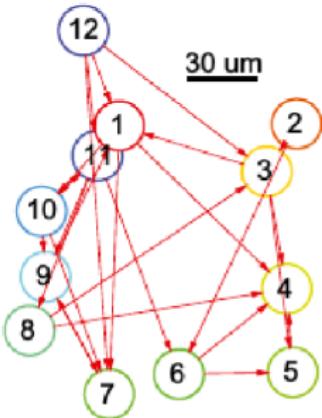
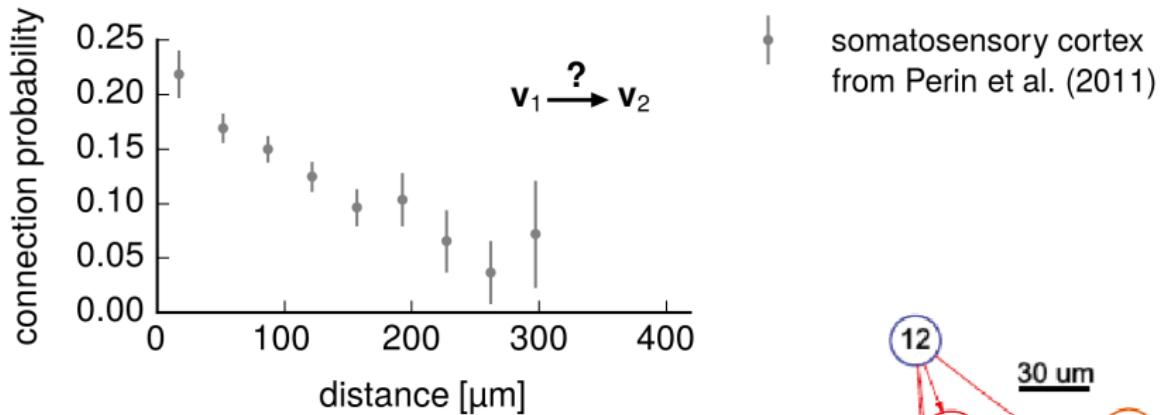
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Anisotropic network model

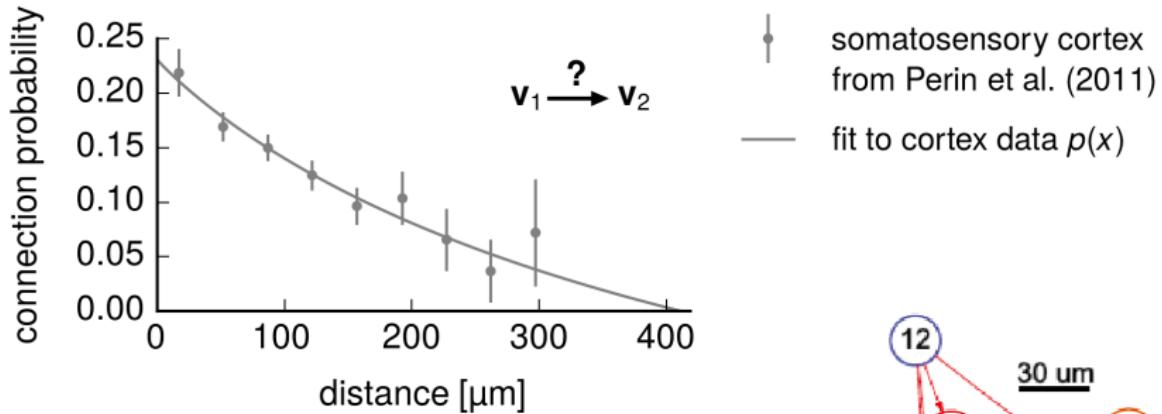


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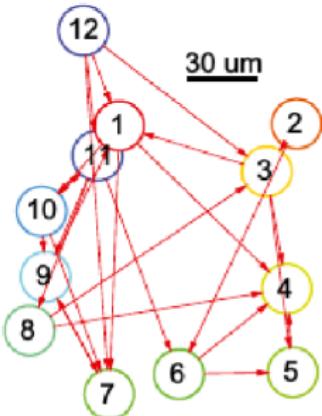
Anisotropic network model – Distance-dependency



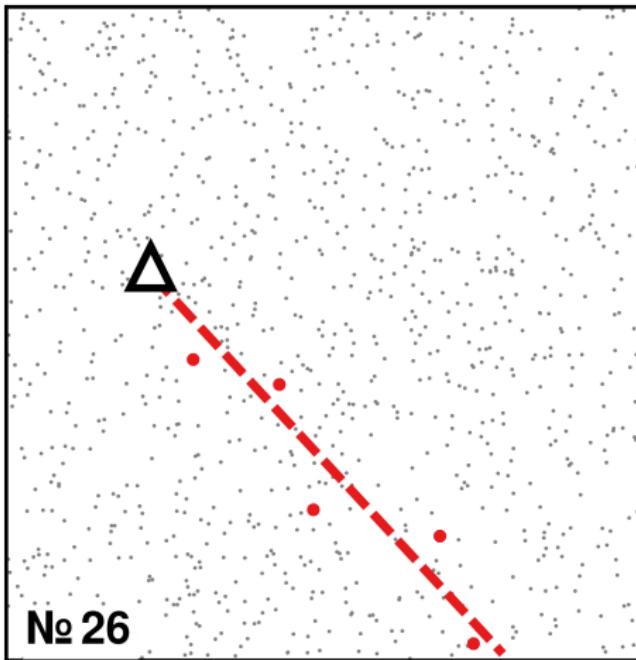
Anisotropic network model – Distance-dependency



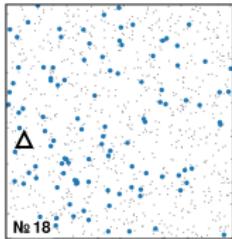
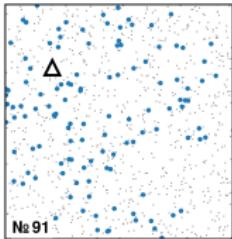
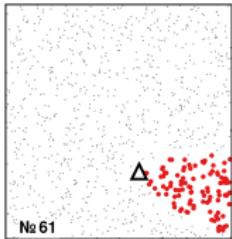
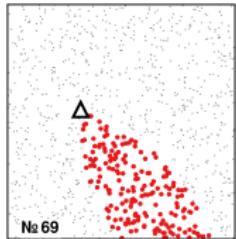
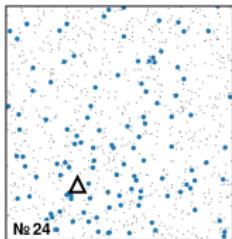
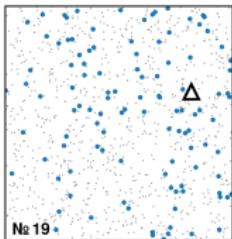
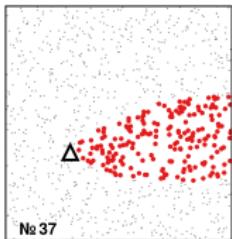
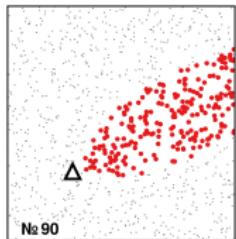
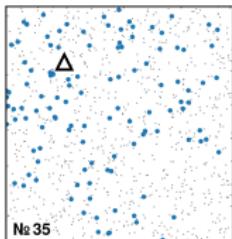
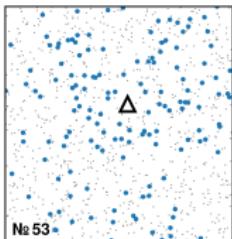
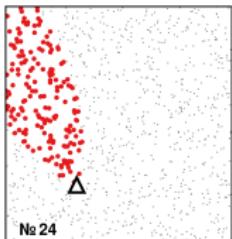
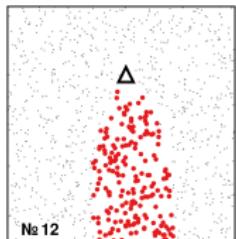
● somatosensory cortex
from Perin et al. (2011)
— fit to cortex data $p(x)$



Anisotropic network model



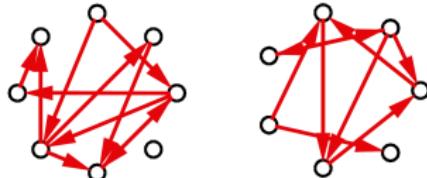
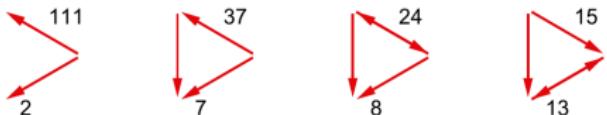
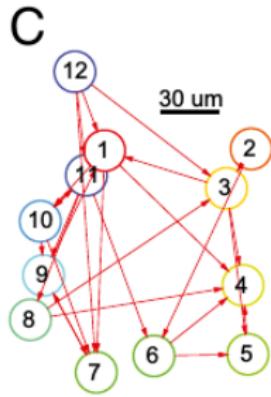
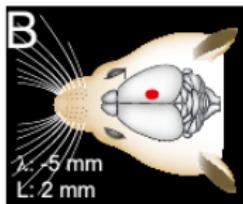
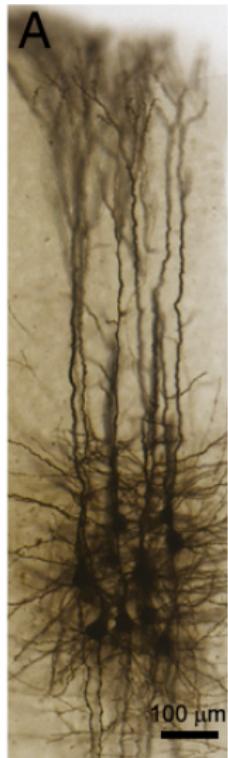
Anisotropic network model



targets

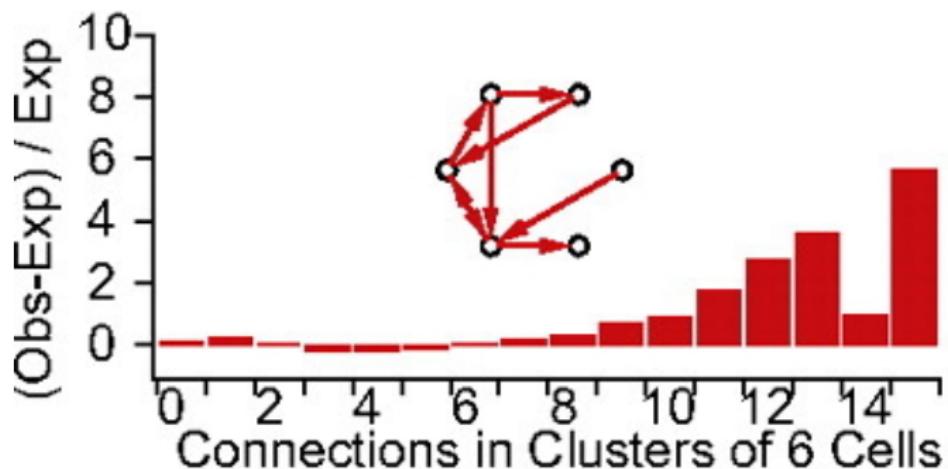
inputs

Robust nonrandom connectivity patterns

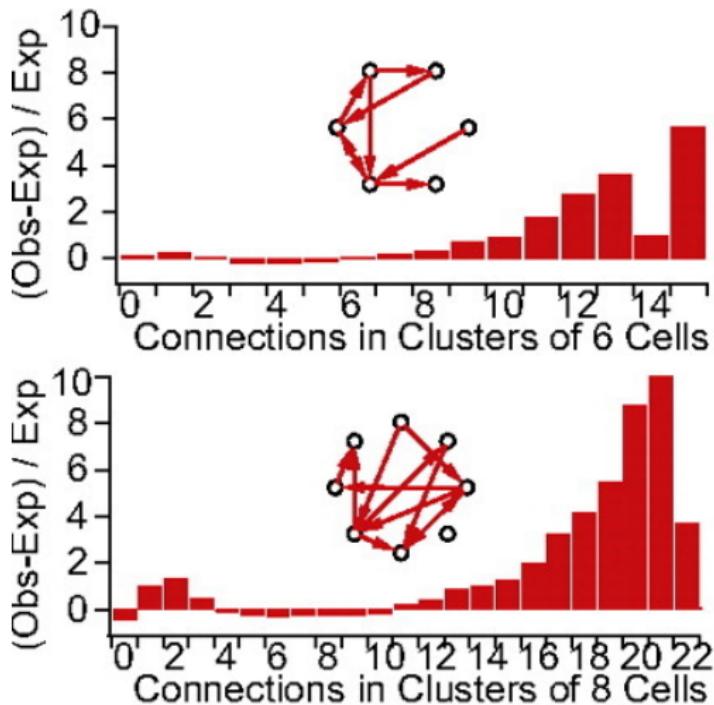


Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

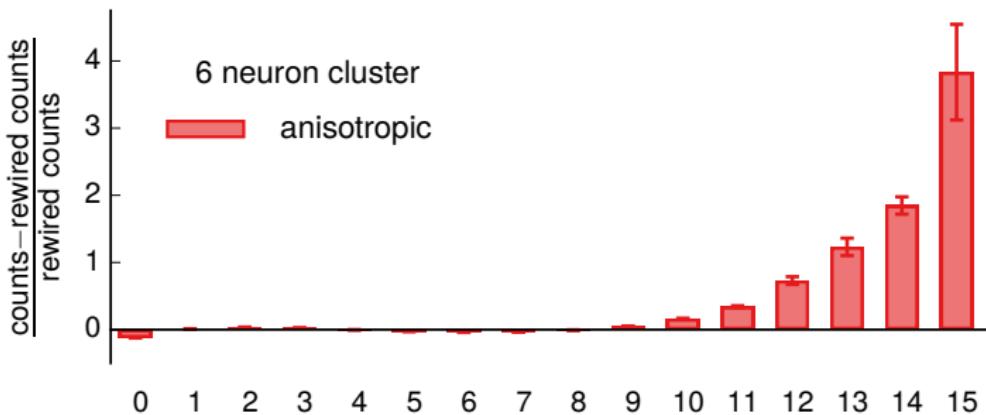
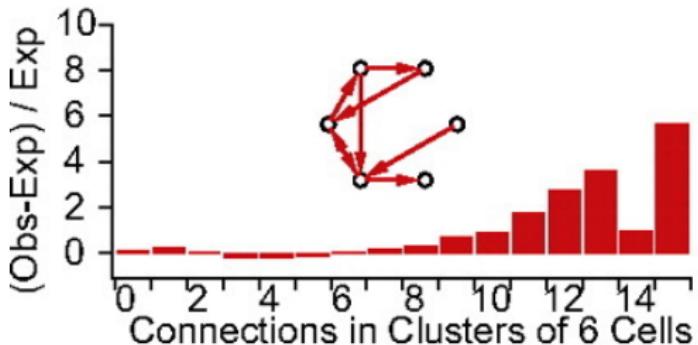
Connection counts in neuron clusters



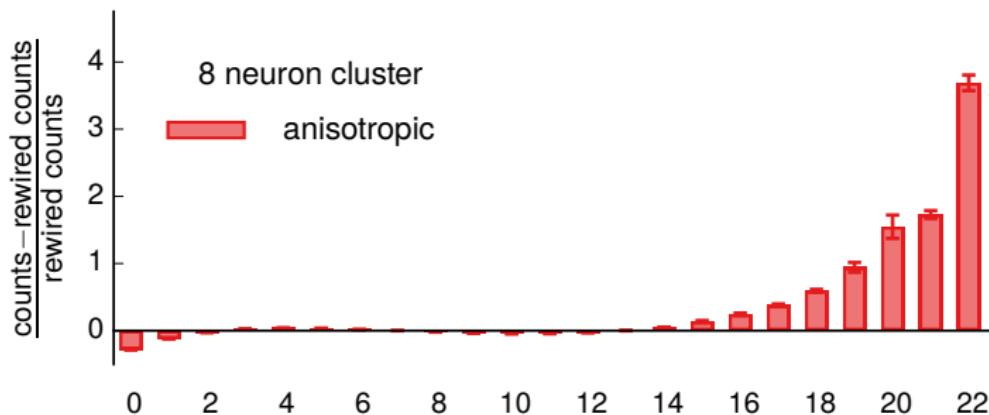
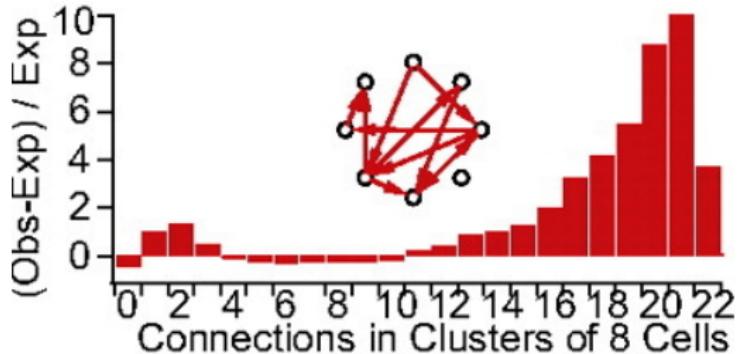
Connection counts in neuron clusters



Connection counts in neuron clusters

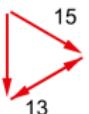
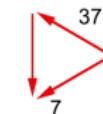
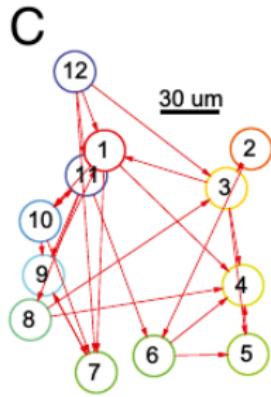
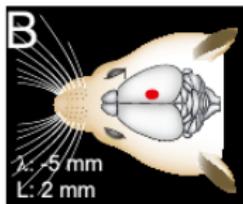


Connection counts in neuron clusters



Results

Robust nonrandom connectivity patterns



Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

Overrepresentation of reciprocal connections

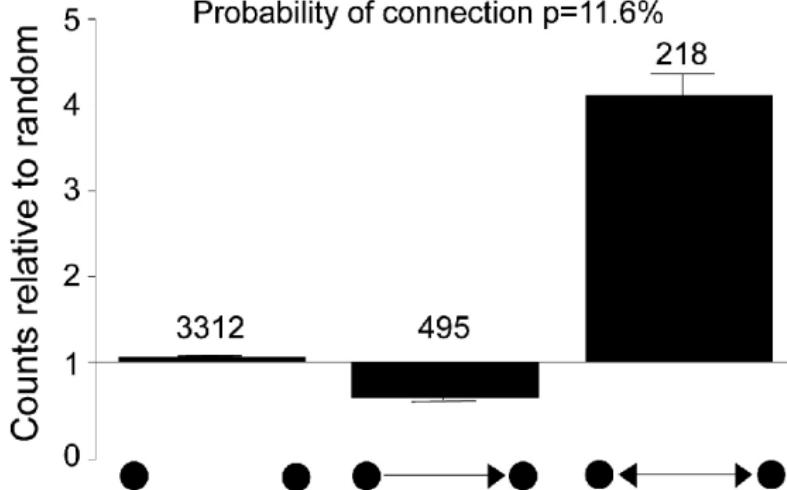
A

Null hypothesis assumes independent connection probabilities

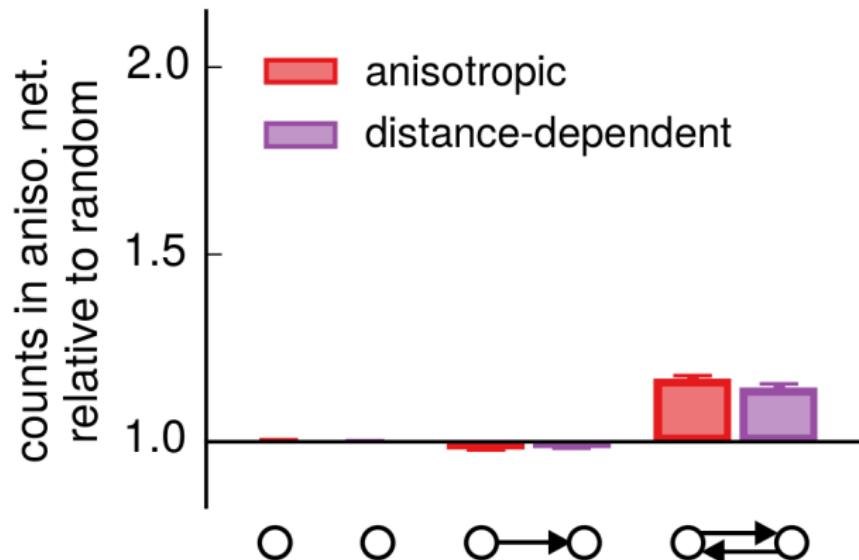


B

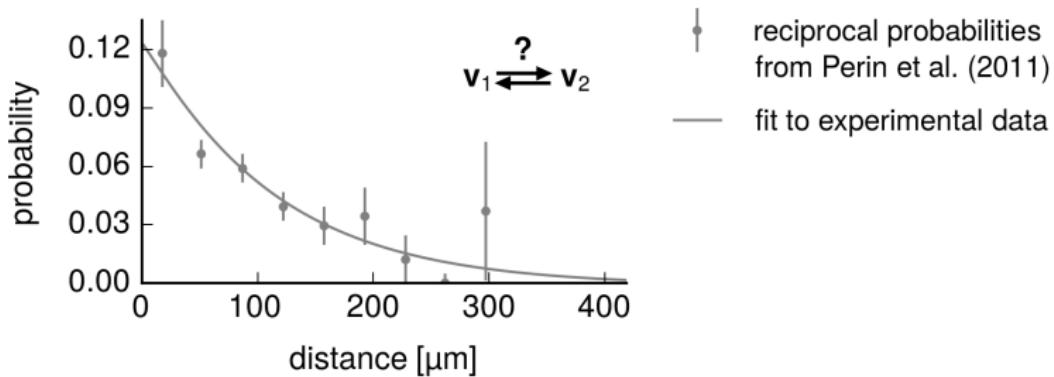
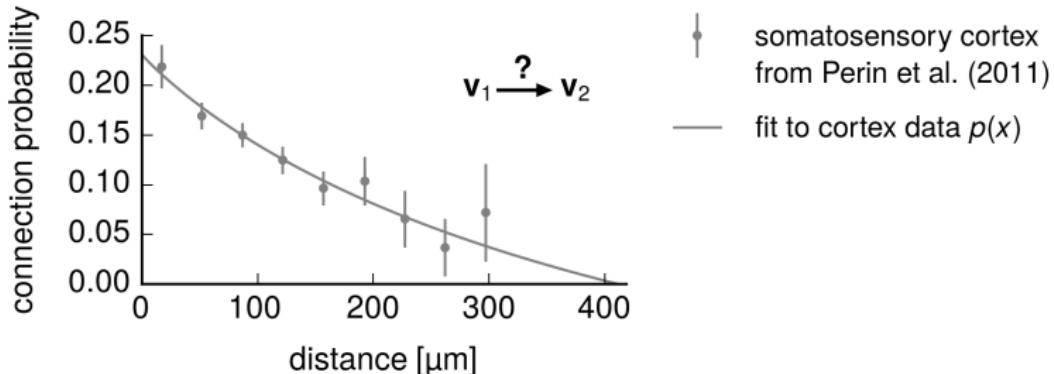
Probability of connection $p=11.6\%$



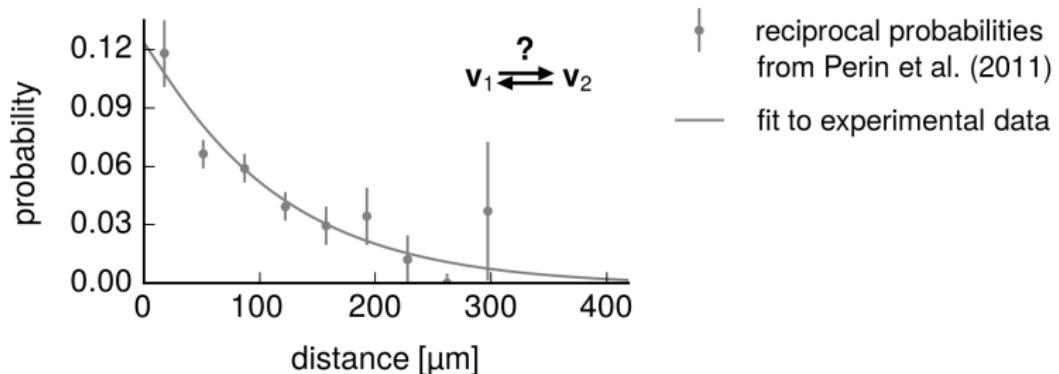
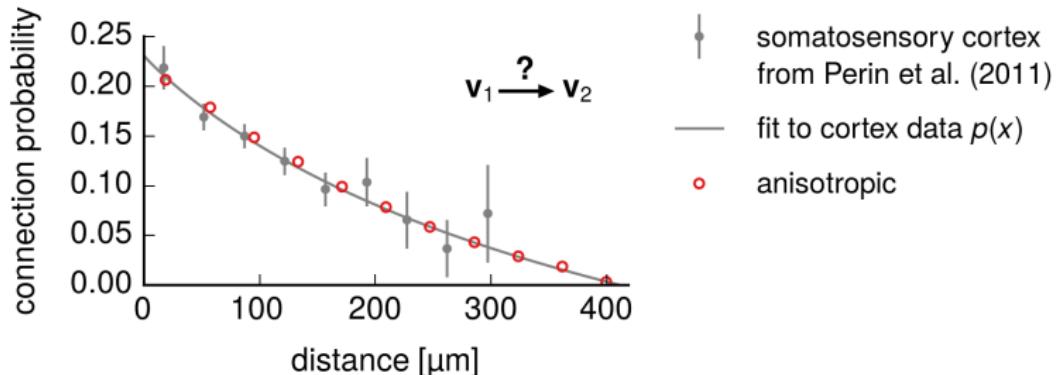
Overrepresentation of reciprocal connections



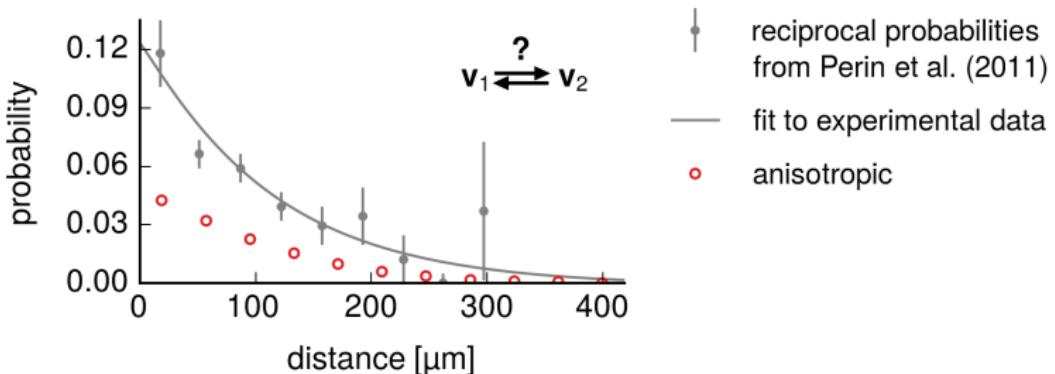
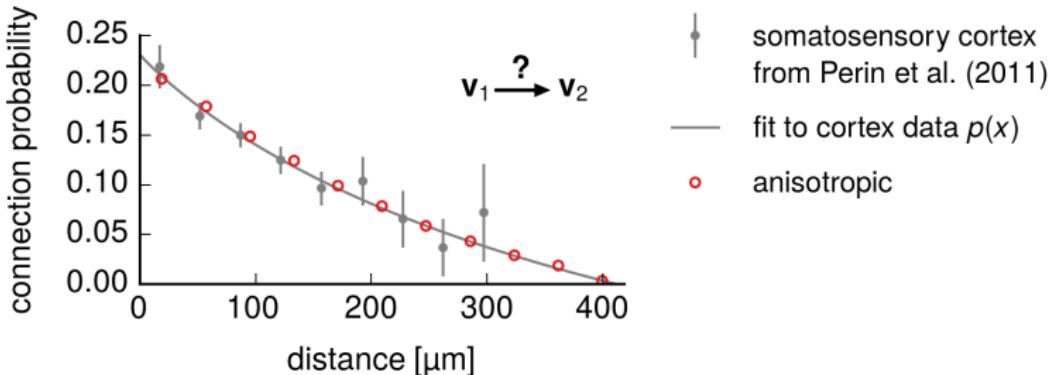
Reciprocity – Distance-dependent?



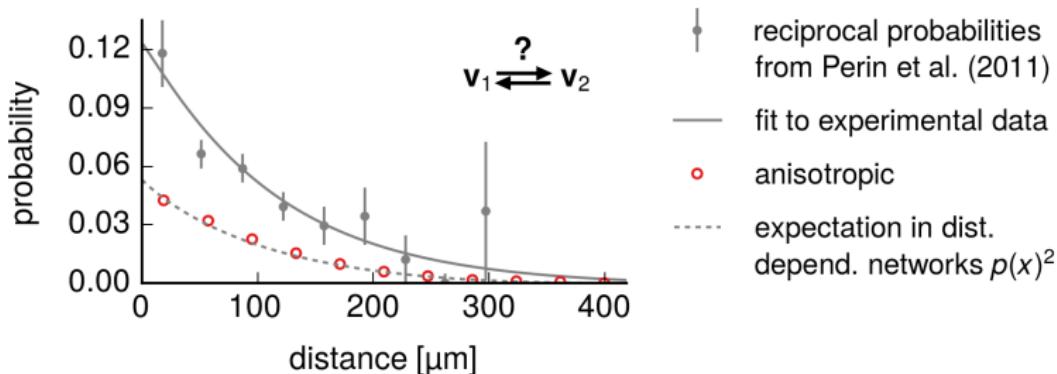
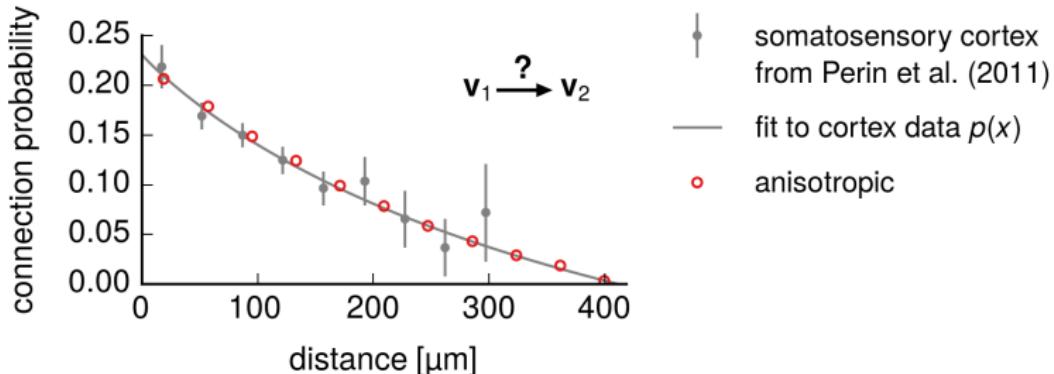
Reciprocity – Distance-dependent?



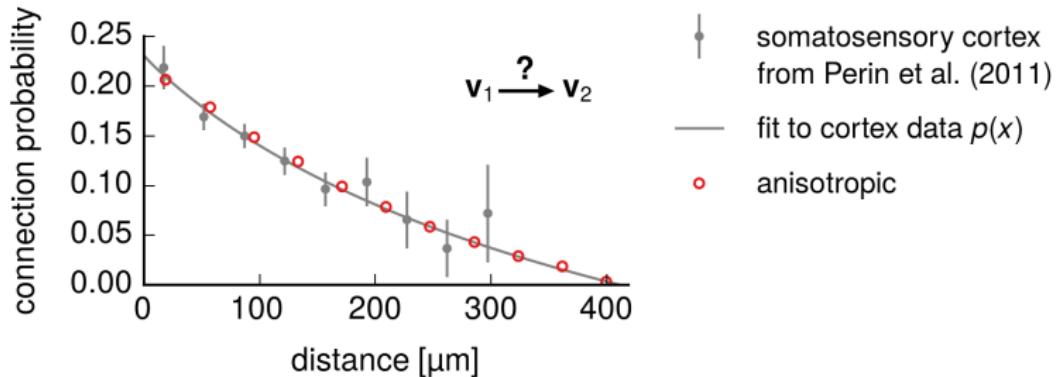
Reciprocity – Distance-dependent?



Reciprocity – Distance-dependent?

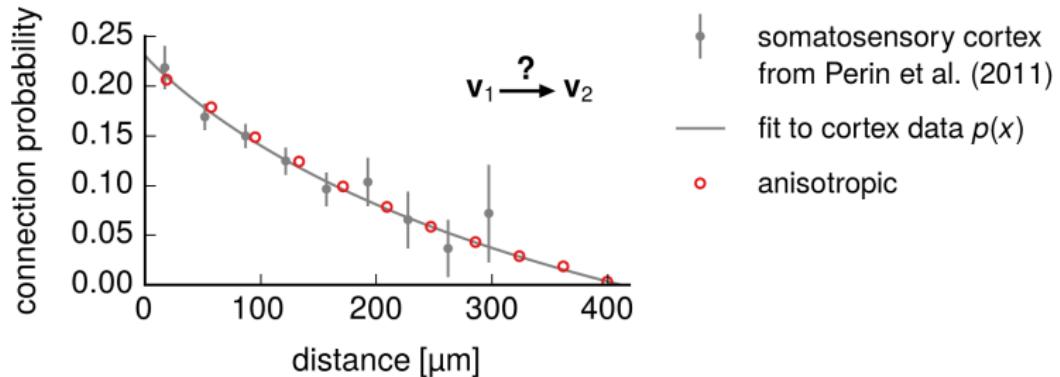


Reciprocity – Distance-dependent?



Other sources for the overrepresentation of bidirectional connections?

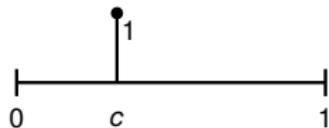
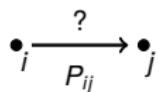
Reciprocity – Distance-dependent?



Other sources for the overrepresentation of bidirectional connections?

⇒ Hoffmann, FZ and Triesch, J (2017). Nonrandom Network Connectivity Comes in Pairs. *Network Neuroscience*

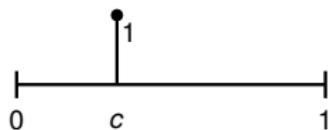
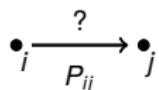
Standard random network model



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

Standard random network model



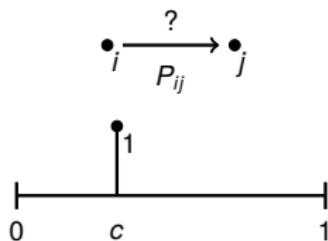
Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

Overall connection probability

$$\mu = P_{ij} = c$$

Standard random network model



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

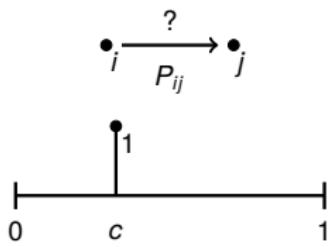
Overall connection probability

$$\mu = P_{ij} = c$$

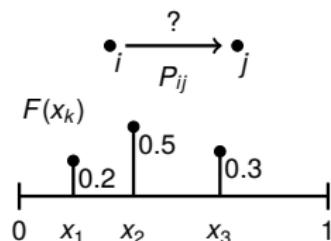
Bidirectional connection

$$P_{\text{bidir}} = P_{ij}P_{ji} = c^2$$

Standard random network model



Varying connection probabilities



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

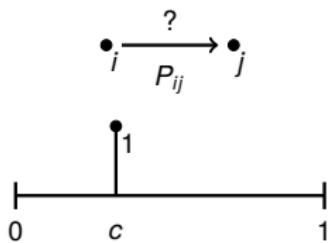
Overall connection probability

$$\mu = P_{ij} = c$$

Bidirectional connection

$$P_{\text{bidir}} = P_{ij}P_{ji} = c^2$$

Standard random network model



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

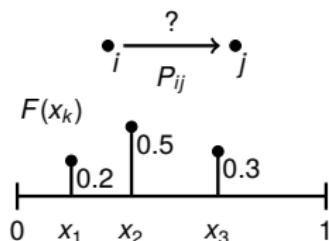
Overall connection probability

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Bidirectional connection

$$P_{\text{bidir}} = P_{ij}P_{ji} = c^2$$

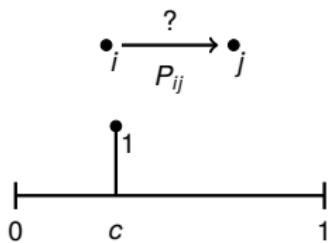
Varying connection probabilities



Probability of connection a random variable P_{ij} ,

$$\mathbf{Prob}(P_{ij} = x_k) = F(x_k)$$

Standard random network model



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

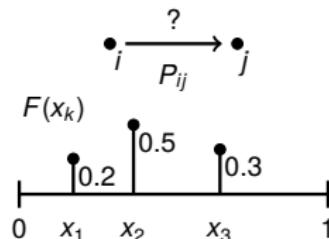
Overall connection probability

$$\mu = P_{ij} = c$$

Bidirectional connection

$$P_{\text{bidir}} = P_{ij}P_{ji} = c^2$$

Varying connection probabilities



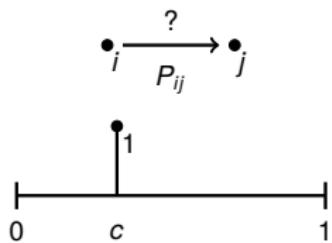
Probability of connection a random variable P_{ij} ,

$$\mathbf{Prob}(P_{ij} = x_k) = F(x_k)$$

Overall connection probability

$$\mu = \sum_{k=1}^m F(x_k)x_k$$

Standard random network model



Probability of connection a constant P_{ij} ,

$$P_{ij} = c$$

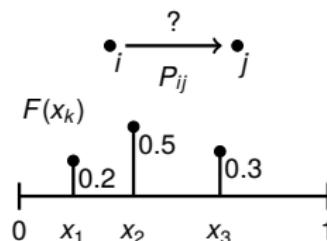
Overall connection probability

$$\mu = P_{ij} = c$$

Bidirectional connection

$$P_{\text{bidir}} = P_{ij}P_{ji} = c^2$$

Varying connection probabilities



Probability of connection a random variable P_{ij} ,

$$\text{Prob}(P_{ij} = x_k) = F(x_k)$$

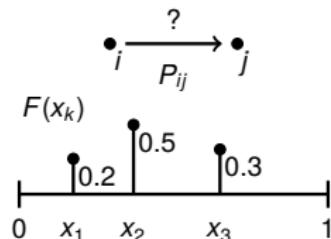
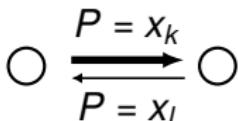
Overall connection probability

$$\mu = \sum_{k=1}^m F(x_k)x_k$$

Bidirectional connection

$$P_{\text{bidir}} = ?$$

Varying connection probabilities



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Overall connection probability

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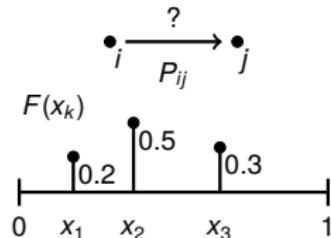
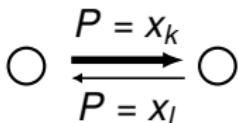
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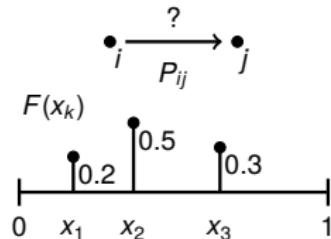
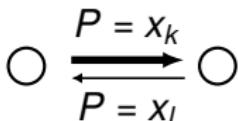
Overall connection probability

$$\mu = \sum_{k=1}^m F(x_k)x_k$$

Bidirectional connection

$$P_{\text{bidir}} = \sum_{k=1}^m \sum_{l=1}^m F(x_k)x_k F(x_l|x_k)x_l$$

Varying connection probabilities



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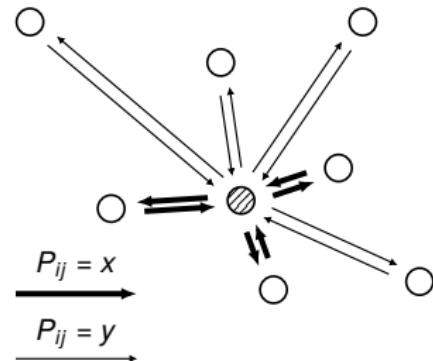
Overall connection probability

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Bidirectional connection

$$P_{\text{bidir}} = \sum_{k=1}^m \sum_{l=1}^m F(x_k)x_k F(x_l|x_k)x_l$$

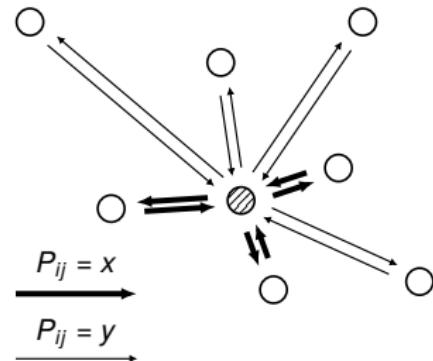
$$F(x_l|x_k) = \begin{cases} 1 & \text{if } l = k \\ 0 & \text{otherwise.} \end{cases}$$



$$P_{\text{bidir}} = \sum_{k=1}^m \sum_{l=1}^m F(x_k) x_k F(x_l | x_k) x_l$$

$$= \sum_{k=1}^m F(x_k) x_k^2.$$

$$F(x_l | x_k) = \begin{cases} 1 & \text{if } l = k \\ 0 & \text{otherwise.} \end{cases}$$

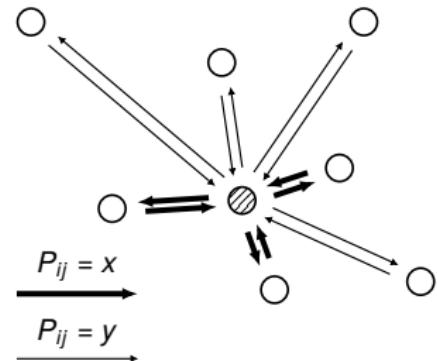


$$\begin{aligned}
 P_{\text{bidir}} &= \sum_{k=1}^m \sum_{l=1}^m F(x_k) x_k F(x_l | x_k) x_l \\
 &= \sum_{k=1}^m F(x_k) x_k^2.
 \end{aligned}$$

$$F(x_l | x_k) = \begin{cases} 1 & \text{if } l = k \\ 0 & \text{otherwise.} \end{cases}$$

Relative overrepresentation ρ is the fraction

$$\rho = \frac{P_{\text{bidir}}}{\mu^2} = \frac{\sum_{k=1}^m F(x_k) x_k^2}{\left(\sum_{k=1}^m F(x_k) x_k\right)^2}.$$



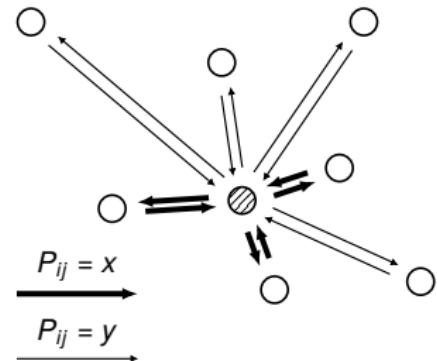
$$P_{\text{bidir}} = \sum_{k=1}^m \sum_{l=1}^m F(x_k) x_k F(x_l | x_k) x_l$$

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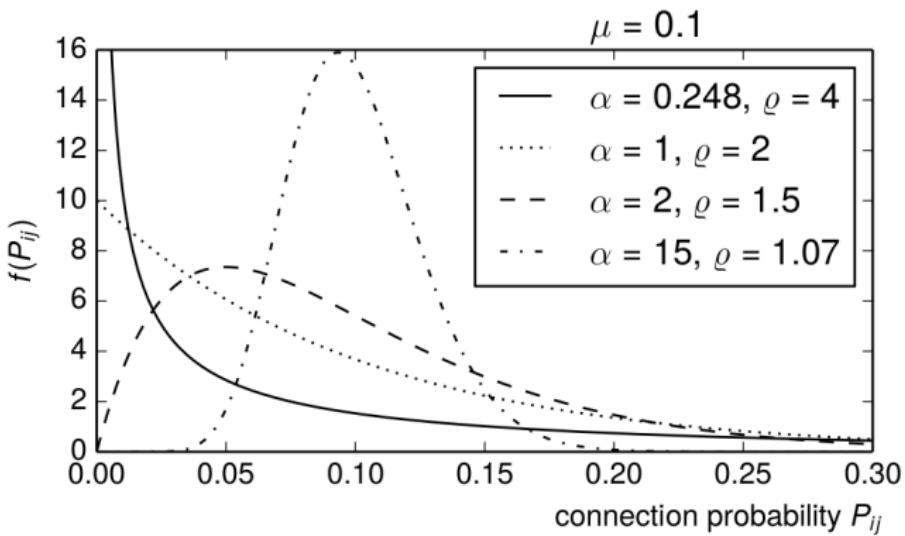
Relative overrepresentation ρ is the fraction

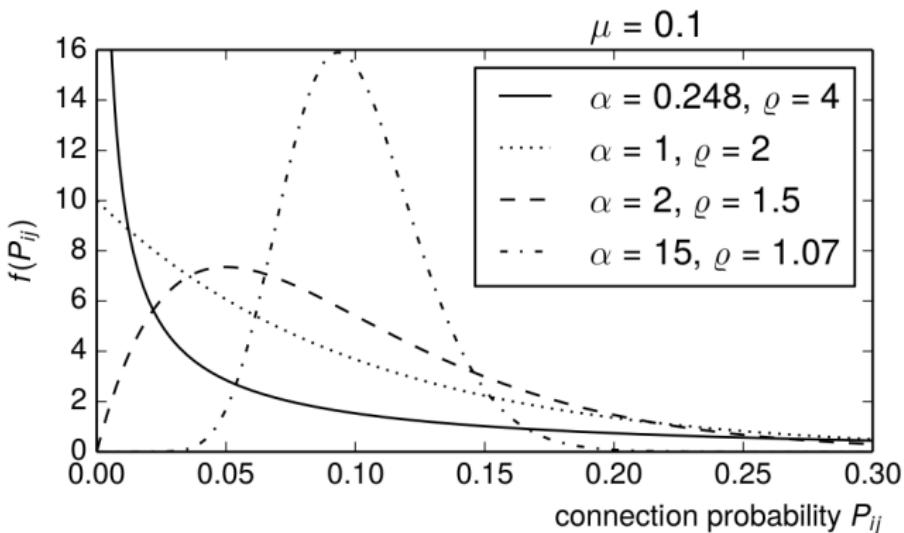
$$\rho = \frac{P_{\text{bidir}}}{\mu^2} = \frac{\sum_{k=1}^m F(x_k) x_k^2}{\left(\sum_{k=1}^m F(x_k) x_k\right)^2}.$$



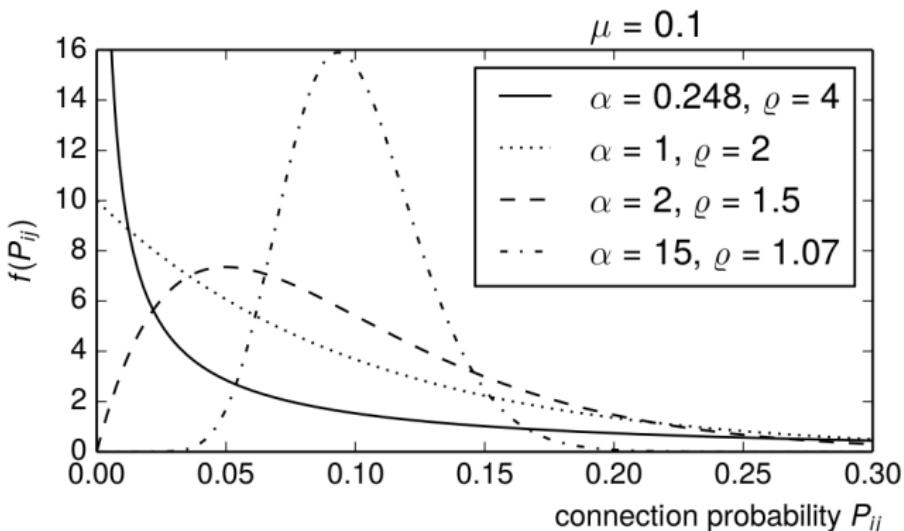
By Jensen's inequality,

$$\left(\sum_{k=1}^m F(x_k) x_k \right)^2 \leq \sum_{k=1}^m F(x_k) x_k^2 \quad \text{and thus} \quad \rho \geq 1.$$



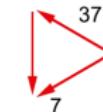
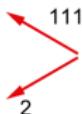
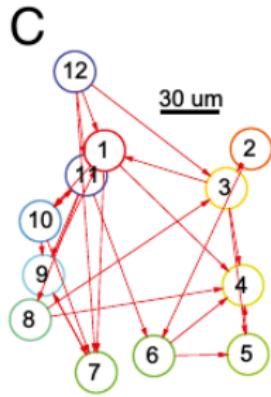
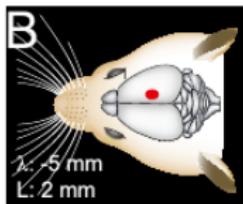


- multiple neuron properties together can cause strong overrepresentation



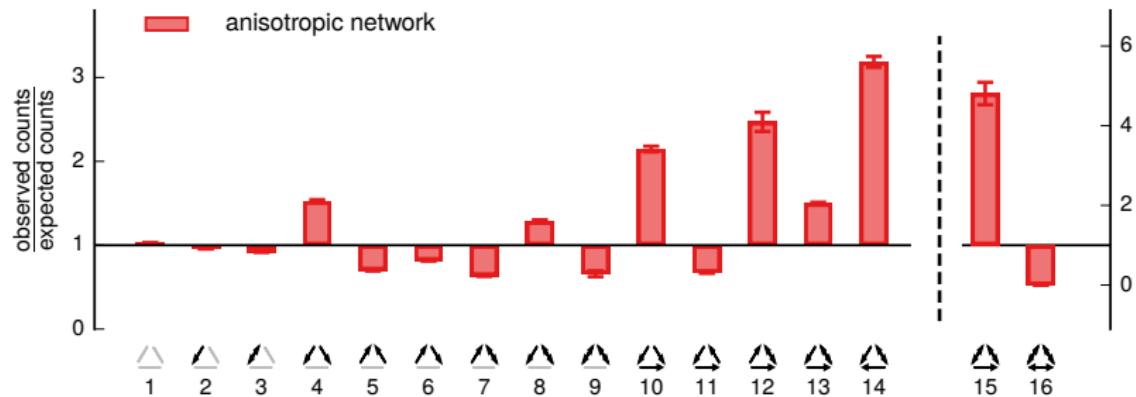
- multiple neuron properties together can cause strong overrepresentation
- example: higher connection probability in functionally related cells (Lee et al. 2016)

Robust nonrandom connectivity patterns



Perin et al. 2011; Song et al. 2005; Markram et al. 1997; Miner and Triesch 2016; Gal et al. 2017; Vegué et al. 2017

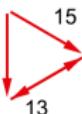
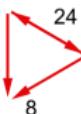
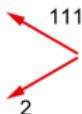
Triplet motif statistics in anisotropic networks



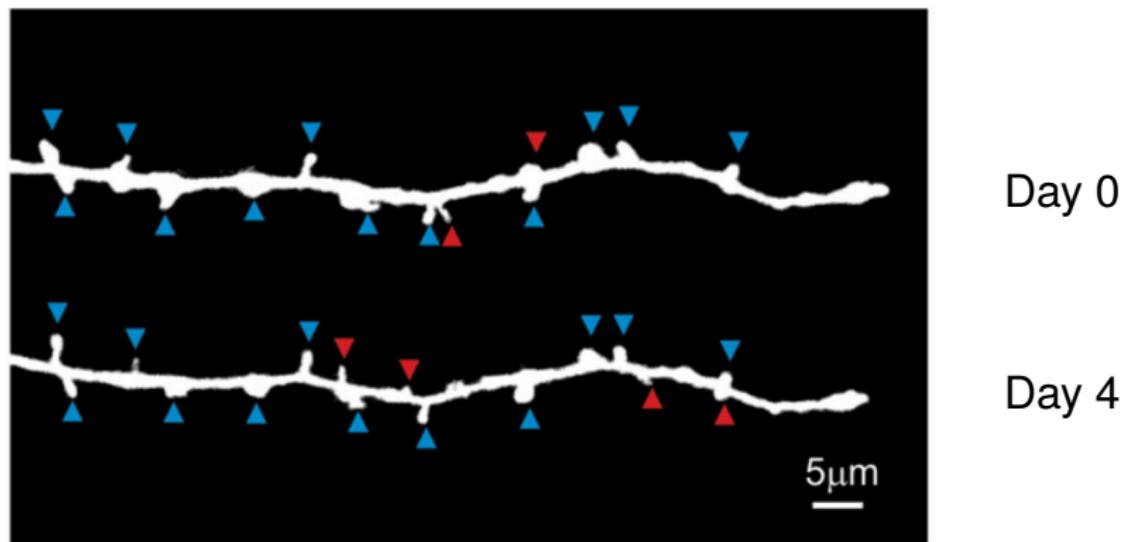
⇒ More about this: Poster W77, Wednesday!

Nonrandom connectivity from anisotropy

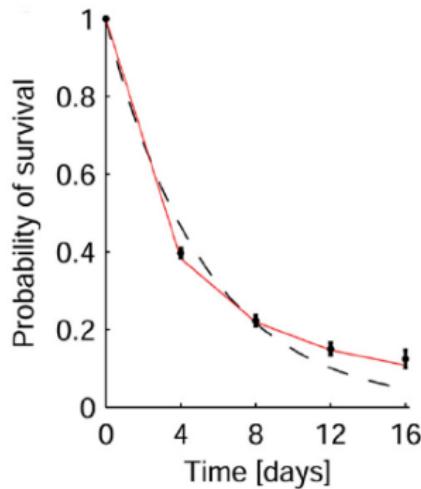
P14



The dynamic connectome

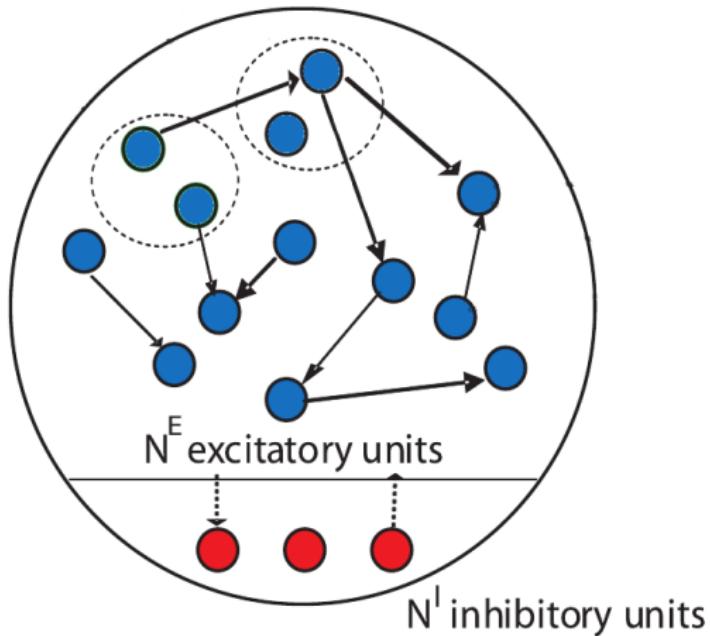


The dynamic connectome

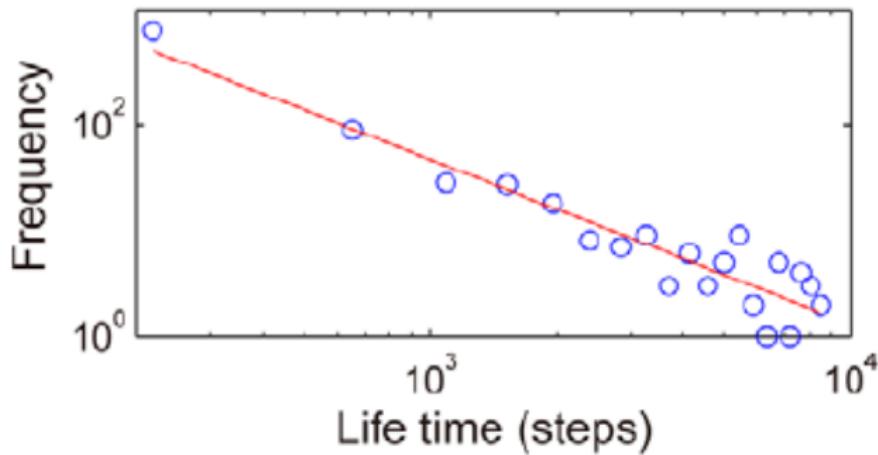


Survival probability of $\rho(t) = (t + 1)^{-\gamma}$ with $\gamma \approx 1.4$,
equivalently lifetime distribution of $f(t) = \gamma(t + 1)^{-(\gamma+1)}$

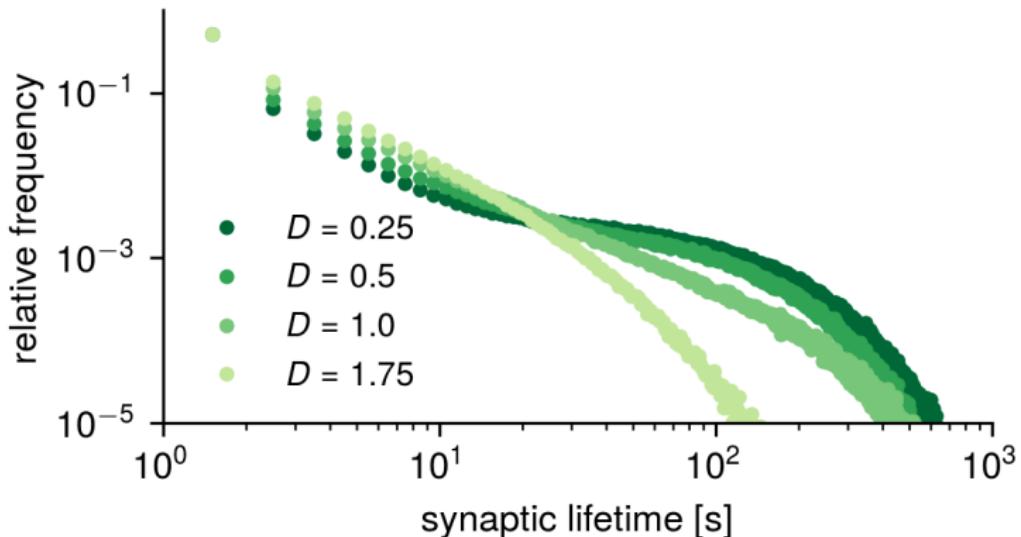
Synapse dynamics in self-organizing recurrent networks



Synapse dynamics in self-organizing recurrent networks



Lifetimes in the self-organizing recurrent network



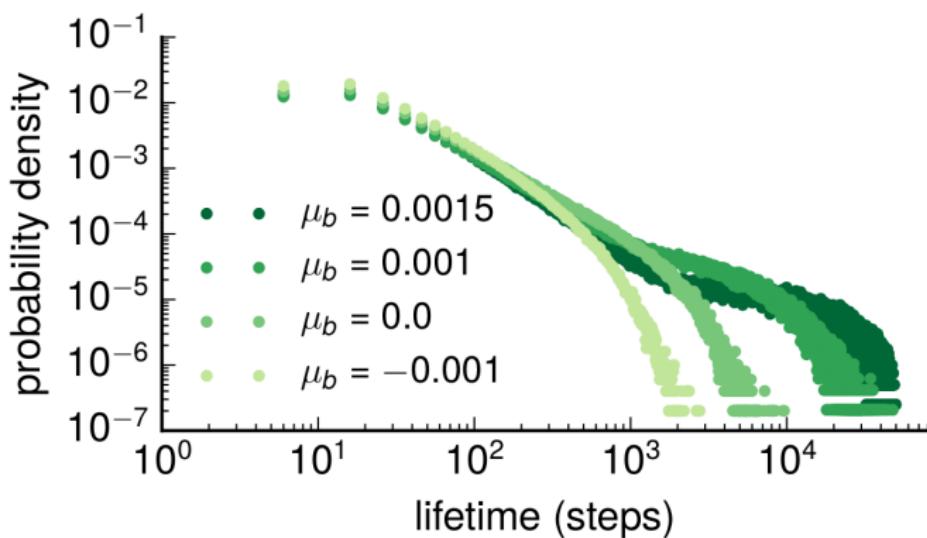
D = LTD–LTP balance

Lifetimes modelled by a stochastic process

Kesten process (Kesten 1973; Statman et al. 2014)

$$X_{n+1} = a_n X_n + b_n$$

as a model for synapse dynamics



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Stefano Cardanobile*

*Florence Kleberg
Triesch lab*



FIAS Frankfurt Institute
for Advanced Studies



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