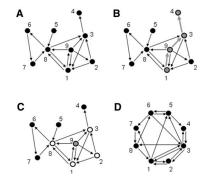
CS 4884: Small World of Brain Networks



T. M. Murali

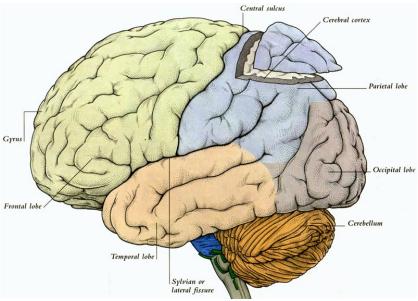
February 13, 2020

Motivation

- The Watts-Strogatz paper set off a storm of research.
- It has 41,500 citations. Even in 2004, it had more than 2,100 citations.
- The C. elegans neuronal network is small-world.

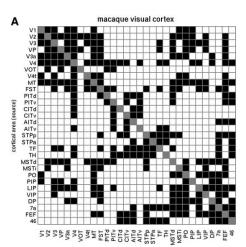
Do mammalian brain networks have the small world property?

Visual and Cerebral Cortices

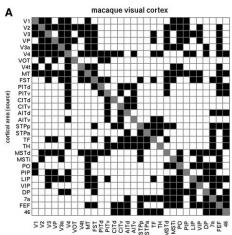


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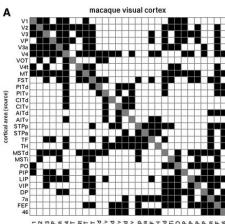




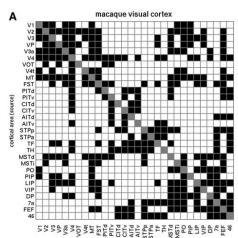
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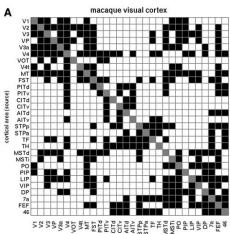


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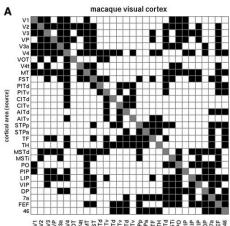
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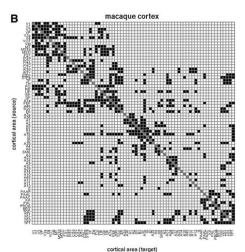
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 Corresponds to an unweighted, directed graph.
- n = 30, m = 311. (The authors use N for the number of nodes and K for the number of edges.)

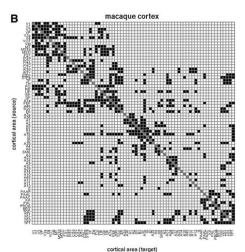
Datasets: Macaque Cerebral Cortex



• n = 71, m = 746.

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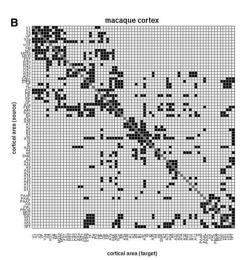
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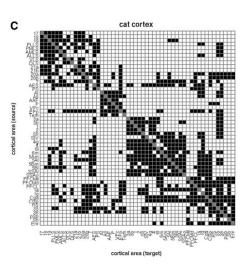
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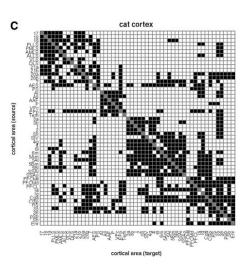
- n = 71, m = 746.
- What is the relation between this graph and the one for the macaque visual cortex? Most of the edges in the previous graph are in this one.

Datasets: Cat Cerebral Cortex



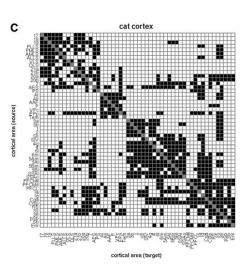
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Datasets: Cat Cerebral Cortex



- n = 52, m = 820.
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Datasets: Cat Cerebral Cortex



- n = 52, m = 820.
- What approximation did the authors make? Discarded density information.
- We will ignore density-based connectivity data sets.

Reference Cases

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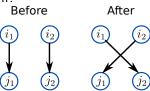
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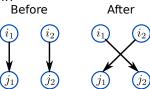
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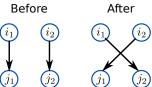
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• Degree-preserving lattice matrix:

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 Degree-preserving lattice matrix: Very poorly specified. Will need to read the code to understand.

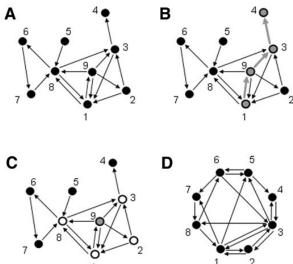
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 - ▶ Clustering coefficient, c(v), called "cluster index" and denoted by $\gamma(v)$ in this paper.

Computing I(G) and c(G) for Directed Graphs



• Appropriately generalise definitions for undirected graphs.

Scaling I(G) and c(G)

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• What values will a small world network have for these quantities? Small $I_{scl}(G)$ and large $c_{scl}(G)$.

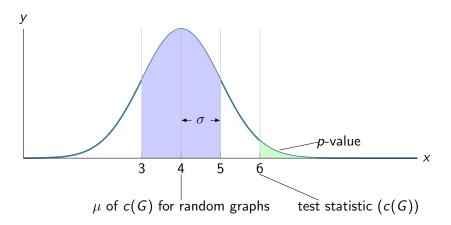
Results for I(G) and c(G)

Table 1. Path Length (λ, λ_{scl}) and Cluster Index (γ, γ_{scl}) for Large-Scale Connection Matrices of Cortico-Cortical Pathways

Topology	λ	γ	λ_{scl}	γ_{scl}
MVC	1.7256	0.5313	0.2188	0.5645
R _{30,311}	1.6680 (0.0038)*	0.3616 (0.0048) *		
L _{30,311}	1.9313 (0.0018)*	0.6622 (0.0000) *		
Rio _{30,311}	1.6880 (0.0033)*	0.4305 (0.0059) *		
Lio _{30,311}	1.8190 (0.0391)	0.6214 (0.0243)		
MC	2.3769	0.4614	0.1927	0.6117
R _{71,746}	2.0310 (0.0051)*	0.1497 (0.0030) *		
L _{71,746}	3.8262 (0.0099)*	0.6593 (0.0002) *		
Rio _{71,746}	2.1159 (0.0133)*	0.2409 (0.0047) *		
Lio _{71,746}	2.8901 (0.1173)*	0.8992 (0.0211) *		
CC	1.8114	0.5514	0.2498	0.6292
R _{52,820}	1.7014 (0.0013)*	0.3103 (0.0026) *		
L _{52,820}	2.1418 (0.0024)*	0.6933 (0.0000) *		
Rio _{52,820}	1.7217 (0.0037)*	0.4023 (0.0030) *		
Lio _{52,820}	1.8570 (0.0283)	0.5893 (0.0172)		

Measures for reference cases represent means and standard deviations (in brackets) for 10 exemplars. Topologies: MVC = macaque visual cortex (Fig. 1A), MC = macaque cortex (Fig. 1B), CC = cat cortex (Fig. 1C), $R_{\rm N,K}$ = random, $L_{\rm N,K}$ = lattice, Rio_{N,K}, Lio_{N,K} = random, lattice matrices with in-degree and out-degree distribution preserved. Statistical significance for all comparisons between cortical matrices and random, lattice, Rio, or Lio matrices marked by "*" are ν =0.001, the remaining comparisons are ν =<0.05.

Statistical Significance



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- These differences are not due to the degree distributions but due to some other intrinsic properties of the connectomes.
- Caveats:
 - p-values are likely to be underestimated. They should be estimated from empirical distributions built from many more random samples.
 - ▶ No indication of correction for testing multiple hypotheses.
 - ▶ No p-value associated with the scaled values of I(G) and c(G).

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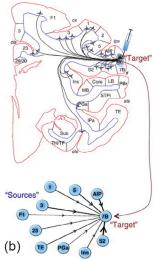
• What should $\sigma(G)$ be for a network with the small world property? $c(G) > c_{\text{random}}$ and $I(G) > I_{\text{random}}$, so $\sigma(G) \approx 1$ or $\sigma(G) > 1$.

Is Small-Worldness Universal in Connectomes?



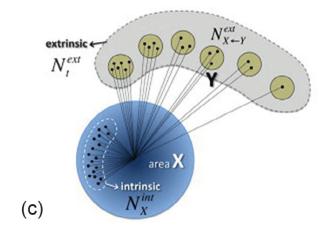
- How does small-worldness emerge in connectomes?
- Do all connectomes have the small-world property?

Structural Connectivity at the Mesoscale: Macaque



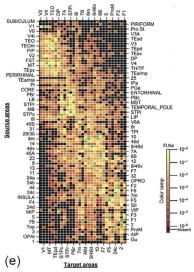
Use retrograde tract tracing. Determine edges coming into node representing area of injection from "labelled" nodes representing neurons that the tracer reaches.

Structural Connectivity at the Mesoscale: Macaque



Injection is at X: $w(Y,X) = \frac{\text{number of neurons labelled in } Y}{\text{total number of labelled neurons}}$

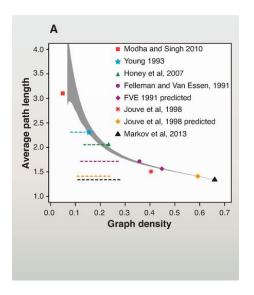
Structural Connectivity at the Mesoscale: Macaque



Example of connectivity matrix.

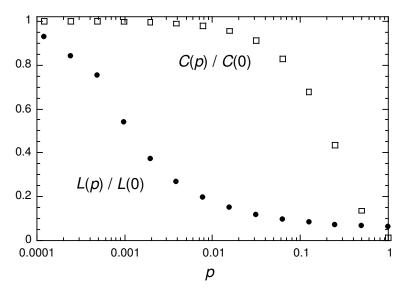
Edge weights range over six orders of magnitude.

Arguments Against Universality



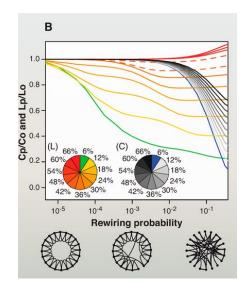
Markov et al., Cortical high-density counterstream architectures. Science, 2013.

Arguments Against Universality



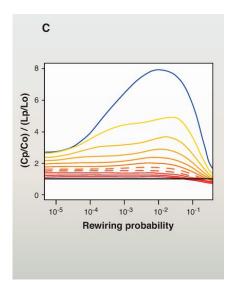
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Table 1. Small-World Metrics.a

	Ma	Macaque		Mouse	
	Binary	Weighted	Binary	Weighted	
Γ	1.21±0.014	1.59 ± 0.007	1.31±0.004	1.76 ± 0.009	
Λ	1.00 ± 0.000	$\textbf{1.27} \pm \textbf{0.057}$	$\textbf{1.00} \pm \textbf{0.000}$	$\textbf{1.47} \pm \textbf{0.021}$	
σ	1.21 ± 0.014	$\textbf{1.25} \pm \textbf{0.071}$	1.31 ± 0.004	1.20 ± 0.019	
ф	N/A	0.574 ± 0.041	N/A	$\boldsymbol{0.800 \pm 0.002}$	

^aFor the macaque and mouse connectomes, we show the mean and standard deviation of the normalized clustering coefficient (Γ) , the normalized path length (Λ) , the small-world index (σ) , and the small-world propensity (ϕ) for both binary and weighted graphs.

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