

Reproducibility of graph metrics of human brain structural networks

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Neuroinformatics with the Insight ToolKit

ABSTRACT

Recent interest in the human connectome has led to the application of graph theoretical analysis to human brain structural networks, in particular white matter connectivity inferred from diffusion imaging and fiber tractography. While these methods have been used to study a variety of patient populations, there has been less examination of the reproducibility of these methods. These graph metrics typically derive from fiber tractography, however a number of tractography algorithms exist and many of these are known to be sensitive to user-selected parameters. The methods used to derive a connectivity matrix from fiber tractography output also influence the resulting graph metrics. Here we examine how these algorithm and parameter choices influence the reproducibility of proposed graph metrics.

Keywords: Structure Tractography Connectivity Brain Network Reproducibility

1 INTRODUCTION

Test retest of functional graph metrics via MEG Deuker et al. (2009)
Test retest of functional graph metrics via fMRI Telesford et al. (2010)
Test retest of structural graph metrics via DTI Owen et al. (2013)
Test retest of structural graph metrics via DTI and DSI with multiple labeling schemes Bassett et al. (2011)
Intra and inter subject variability of structural graph metrics via DTI for binary and weighted networks Cheng et al. (2012)
Correlations between pairs of regions using a variety of structural measures Irimia and Van Horn (2012)

Novel contributions

1. Public data and fully open source
2. In-depth examination of deterministic tractography parameters
3. Probabilistic tractography extensions
4. In-depth analysis of streamline-to-matrix conversion
5. Provides plug-and-play framework for evaluation of new methods
6. Easy to extend to functional study (BOLD and ASL)

Table 1. Descriptions and references for graph metrics examined in this study.

Node metrics	Description	Reference
Degree	Number of connections for a node	Watts and Strogatz (1998) Watts and Strogatz (1998) Latora and Marchiori (2001)
Clustering coefficient	Local neighborhood connectivity	
Path length	Average shortest path to all other nodes	
Global efficiency	“Closeness” to all other nodes	
Local efficiency	“Closeness” to local nodes	
Whole-graph metrics		
Small-world	Degree to which high-degree nodes preferentially inter-connect	Watts and Strogatz (1998)
Synchronizability		Motter et al. (2005)
Assortativity		Newman (2002)
Hierarchy		Ravasz and Barabási (2003)
Cost efficiency		Achard and Bullmore (2007)
Rich-club coefficient		Colizza et al. (2006)
Network similarity measures		
Network overlap	Percentage of common edges in constant density networks	van Wijk et al. (2010)
Edge overlap		?

Table 2. Formulas for node metrics.

Degree	$K_i = \sum_{j=1}^n A_{ij}$
Clustering coefficient	$C_i = 2 * e_i / K_i (K_i - 1)$
Path length	$L = 1/N(N-1) \sum_{ij \in n, i \neq j} d_{ij}$
Global efficiency	$E_{glob} = E(G) = 1/N(N-1) \sum_{i \neq j \in G} 1/d_{ij}$
Local efficiency	$E_{loc} = 1/N \sum_{i \in n} E(G_i)$

2 MATERIAL & METHODS

28 Science goes here.
29

2.1 NODE METRICS

30 For more details on the node metrics see Rubinov and Sporns (2010).

2.2 WHOLE-GRAPH METRICS

31 More formulas go here.

3 RESULTS

32 Overview of what we found

3.1 TRACTOGRAPHY

33 Algorithms, parameters

3.2 MATRIX DERIVATION

34 Turning streamlines into nice $N \times N$ matrices

4 DISCUSSION

4.1 DATA SHARING

DISCLOSURE/CONFLICT-OF-INTEREST STATEMENT

35 The authors declare that the research was conducted in the absence of any commercial or financial
36 relationships that could be construed as a potential conflict of interest.

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SUPPLEMENTAL DATA

39 Maybe need this, maybe not

REFERENCES

- 40 Deuker, L., Bullmore, E. T., Smith, M., Christensen, S., Nathan, P. J., Rockstroh, B., et al. (2009)
- 41 Reproducibility of graph metrics of human brain functional networks. *Neuroimage* 47 1460–1468.
- 42 Telesford, Q. K., Morgan, A. R., Hayasaka, S., Simpson, S. L., Barret, W., Kraft, R. A., et al. (2010)
- 43 Reproducibility of graph metrics in fmri networks. *Frontiers in neuroinformatics* 4.
- 44 Owen, J. P., Ziv, E., Bukshpun, P., Pojman, N., Wakahiro, M., Berman, J. I., et al. (2013) Test-
- 45 retest reliability of computational network measurements derived from the structural connectome of
- 46 the human brain. *Brain Connect* 3 160–176. doi:10.1089/brain.2012.0121.
- 47 Bassett, D. S., Brown, J. A., Deshpande, V., Carlson, J. M., and Grafton, S. T. (2011) Conserved and
- 48 variable architecture of human white matter connectivity. *Neuroimage* 54 1262–1279. doi:10.1016/j.
- 49 neuroimage.2010.09.006.
- 50 Cheng, H., Wang, Y., Sheng, J., Kronenberger, W. G., Mathews, V. P., Hummer, T. A., et al.
- 51 (2012) Characteristics and variability of structural networks derived from diffusion tensor imaging.
- 52 *Neuroimage* 61 1153–1164. doi:10.1016/j.neuroimage.2012.03.036.
- 53 Irimia, A. and Van Horn, J. D. (2012) The structural, connectomic and network covariance of the human
- 54 brain. *Neuroimage* 66C 489–499. doi:10.1016/j.neuroimage.2012.10.066.
- 55 Watts, D. J. and Strogatz, S. H. (1998) Collective dynamics of 'small-world' networks. *Nature* 393
- 56 440–442. doi:10.1038/30918.
- 57 Latora, V. and Marchiori, M. (2001) Efficient behavior of small-world networks. *Physical review letters*
- 58 87 198701.
- 59 Motter, A. E., Zhou, C., and Kurths, J. (2005) Enhancing complex-network synchronization. *EPL*
- 60 (*Europhysics Letters*) 69 334.
- 61 Newman, M. E. (2002) Assortative mixing in networks. *Physical review letters* 89 208701.
- 62 Ravasz, E. and Barabási, A.-L. (2003) Hierarchical organization in complex networks. *Physical Review*
- 63 *E* 67 026112.
- 64 Achard, S. and Bullmore, E. (2007) Efficiency and cost of economical brain functional networks. *PLoS*
- 65 *Comput Biol* 3 e17. doi:10.1371/journal.pcbi.0030017.
- 66 Colizza, V., Flammini, A., Serrano, M. A., and Vespignani, A. (2006) Detecting rich-club ordering in
- 67 complex networks. *Nature physics* 2 110–115.
- 68 van Wijk, B. C. M., Stam, C. J., and Daffertshofer, A. (2010) Comparing brain networks of different size
- 69 and connectivity density using graph theory. *PLoS One* 5 e13701. doi:10.1371/journal.pone.0013701.

- 70 Rubinov, M. and Sporns, O. (2010) Complex network measures of brain connectivity: uses and
71 interpretations. *Neuroimage* 52 1059–1069. doi:10.1016/j.neuroimage.2009.10.003.