

Reproducibility of graph metrics of human brain structural networks

Jeffrey T. Duda^{1,*}, Philip A. Cook¹ and James C. Gee¹

¹Penn Image Computing and Science Laboratory, University of Pennsylvania, Department of Radiology, Philadelphia, PA, USA

Correspondence*:

Jeffrey T. Duda

Penn Image Computing and Science Laboratory, University of Pennsylvania, Department of Radiology, 3600 Market Street, Suite 370, Philadelphia, PA, USA, jtduda@seas.upenn.edu

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 and/or alternate data sets
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 Provide an overview of what we are examining here

2.1 NODE METRICS

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

2.2 WHOLE-GRAPH METRICS

30 More formulas go here.

2.3 NETWORK SIMILARITY

31 Use these to examine methods that extract sub-networks such as rich-club and constant density networks.

2.4 TRACTOGRAPHY

32 Algorithms, parameters

2.5 MATRIX DERIVATION

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

2.6 NEUROIMAGING DATA

34 The Multi-Modal MRI Reproducibility Resource ? provides a test-retest data set consisting of 21 subjects
35 with two time points each for which T1-weighted anatomical images and diffusion tensor images were
36 acquired. Other image types, not examined here, were also acquired making this data useful for future
37 examinations of structure and function. A population averaged template and manually defined cortical
38 labels for one time point for each subject are available from the Mindboggle project.

2.7 IMAGE ANALYSIS

39 Minimal processing

- 40 1. Intra-subject registration for label transfer
- 41 2. B0-T1 registration
- 42 3. DTI reconstruction
- 43 4. MTR ?

44 Optional processing

- 45 1. Registration to template (for additional label sets)

46 Future processing

- 47 1. Cortical thickness
- 48 2. BOLD / ASL / etc

3 RESULTS

49 Overview of what we found

4 DISCUSSION

4.1 DATA SHARING

DISCLOSURE/CONFLICT-OF-INTEREST STATEMENT

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

ACKNOWLEDGEMENT

52 Shoutouts to our peeps

53 *Funding:* Shoutout to our peep\$

SUPPLEMENTAL DATA

54 Maybe need this, maybe not

REFERENCES

- Deuker, L., Bullmore, E. T., Smith, M., Christensen, S., Nathan, P. J., Rockstroh, B., et al. (2009) Reproducibility of graph metrics of human brain functional networks. *Neuroimage* 47 1460–1468.
- Telesford, Q. K., Morgan, A. R., Hayasaka, S., Simpson, S. L., Barret, W., Kraft, R. A., et al. (2010) Reproducibility of graph metrics in fmri networks. *Frontiers in neuroinformatics* 4.
- Owen, J. P., Ziv, E., Bukshpun, P., Pojman, N., Wakahiro, M., Berman, J. I., et al. (2013) Test-retest reliability of computational network measurements derived from the structural connectome of the human brain. *Brain Connect* 3 160–176. doi:10.1089/brain.2012.0121.
- Bassett, D. S., Brown, J. A., Deshpande, V., Carlson, J. M., and Grafton, S. T. (2011) Conserved and variable architecture of human white matter connectivity. *Neuroimage* 54 1262–1279. doi:10.1016/j.neuroimage.2010.09.006.
- Cheng, H., Wang, Y., Sheng, J., Kronenberger, W. G., Mathews, V. P., Hummer, T. A., et al. (2012) Characteristics and variability of structural networks derived from diffusion tensor imaging. *Neuroimage* 61 1153–1164. doi:10.1016/j.neuroimage.2012.03.036.
- Irimia, A. and Van Horn, J. D. (2012) The structural, connectomic and network covariance of the human brain. *Neuroimage* 66C 489–499. doi:10.1016/j.neuroimage.2012.10.066.
- Watts, D. J. and Strogatz, S. H. (1998) Collective dynamics of 'small-world' networks. *Nature* 393 440–442. doi:10.1038/30918.
- Latora, V. and Marchiori, M. (2001) Efficient behavior of small-world networks. *Physical review letters* 87 198701.
- Motter, A. E., Zhou, C., and Kurths, J. (2005) Enhancing complex-network synchronization. *EPL (Europhysics Letters)* 69 334.
- Newman, M. E. (2002) Assortative mixing in networks. *Physical review letters* 89 208701.
- Ravasz, E. and Barabási, A.-L. (2003) Hierarchical organization in complex networks. *Physical Review E* 67 026112.
- Achard, S. and Bullmore, E. (2007) Efficiency and cost of economical brain functional networks. *PLoS Comput Biol* 3 e17. doi:10.1371/journal.pcbi.0030017.
- Colizza, V., Flammini, A., Serrano, M. A., and Vespignani, A. (2006) Detecting rich-club ordering in complex networks. *Nature physics* 2 110–115.
- van Wijk, B. C. M., Stam, C. J., and Daffertshofer, A. (2010) Comparing brain networks of different size and connectivity density using graph theory. *PLoS One* 5 e13701. doi:10.1371/journal.pone.0013701.
- Rubinov, M. and Sporns, O. (2010) Complex network measures of brain connectivity: uses and interpretations. *Neuroimage* 52 1059–1069. doi:10.1016/j.neuroimage.2009.10.003.