

Supplementary Information for

Adolescent development of multiscale structural wiring and functional interactions in the human connectome.

Bo-yong Park, Casey Paquola, Richard A. I. Bethlehem, Oualid Benkarim, Neuroscience in Psychiatry Network (NSPN) Consortium, Bratislav Mišić, Jonathan Smallwood, Edward T. Bullmore, Boris C. Bernhardt

†A complete list of investigators from the Neuroscience in Psychiatry Network (NSPN) Consortium can be found in the Supporting Information.

Bo-yong Park
Email: boyong.park@inha.ac.kr
Boris C. Bernhardt
Email: boris.bernhardt@mcgill.ca

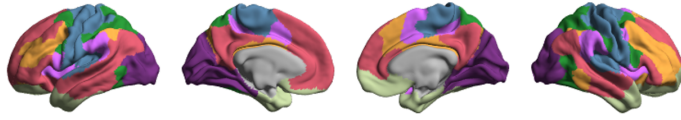
This PDF file includes:

Figures S1 to S12

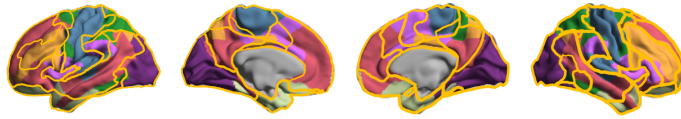
Other supplementary materials for this manuscript include the following:

Neuroscience in Psychiatry Network (NSPN) Consortium author list

A. Yeo parcellation



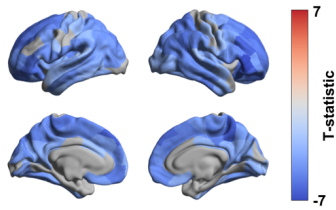
B. Desikan-Killiany subparcellation



- Visual
- Somatomotor
- Dorsal attention
- Ventral attention
- Limbic
- Frontoparietal
- Default mode

Fig. S1. Remapping of the Yeo-7 parcellation to the subparcellation of the Desikan-Killiany atlas used for this study. (A) Intrinsic functional networks based on Yeo, Krienen, et al. (1). **(B)** Parcels of the 200 node Desikan-Killiany subparcellation were assigned to each of these networks, and the boundaries of the Yeo 7-networks are shown with lines.

A. Age-effects on cortical thickness



B. Correlation between multiscale cortical structural differentiation and cortical thickness

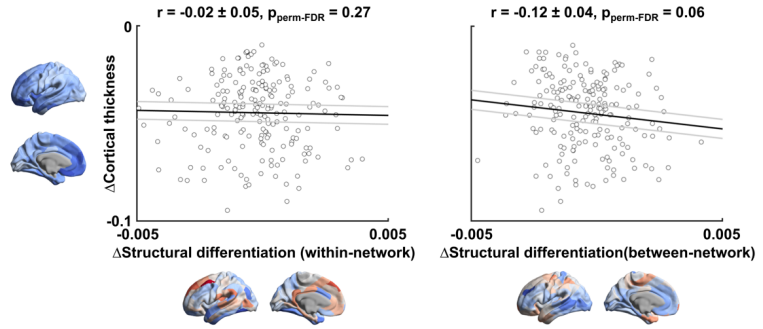


Fig. S2. Cortical thickness effects. (A) The t-statistics of the identified regions that showed significant age-related changes in cortical thickness. **(B)** Linear correlations between time-related changes in cortical thickness and within/between-network structural differentiation.

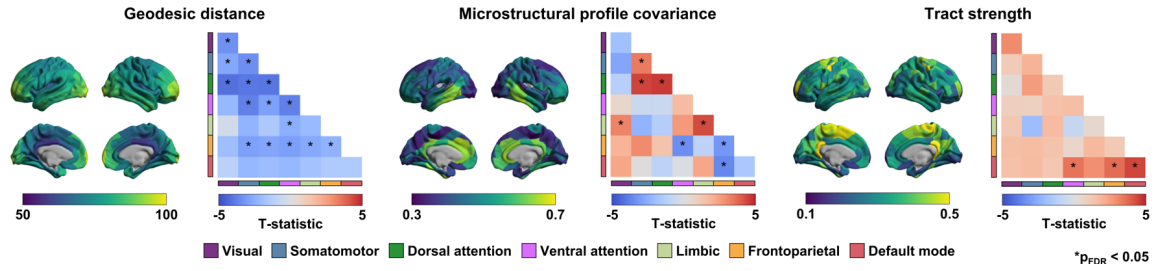


Fig. S3. Age-effects on each cortical wiring feature. The spatial maps of geodesic distance, microstructural profile covariance, and tract strength are shown on the brain surface. The t-statistics of age-related changes on each cortical feature within- and between-networks, with significant ($p_{FDR} < 0.05$) results marked with asterisks. *Abbreviation:* FDR, false discovery rate.

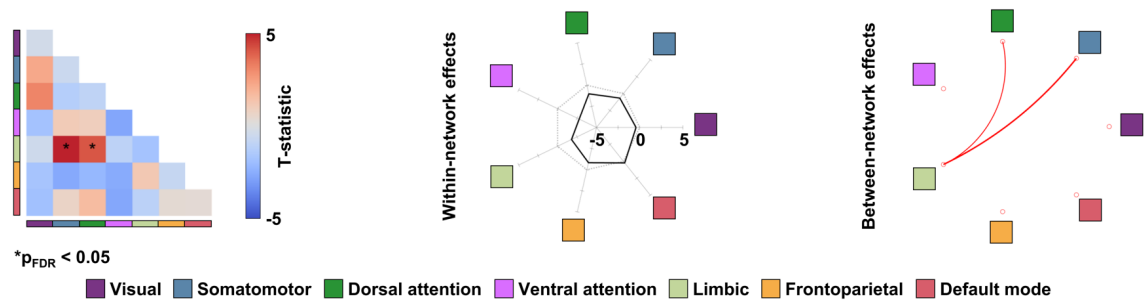


Fig. S4. Age-effects on multiscale cortical structural differentiation after controlling for mean structural differentiation. The t-statistics of age-effects are reported in the matrix, and within- and between-network effects are represented with radar and circular plots, respectively. For details, see *Fig. 1*.

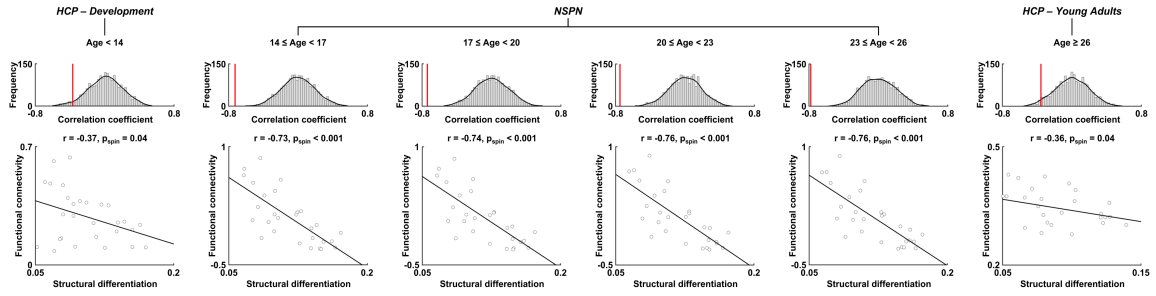
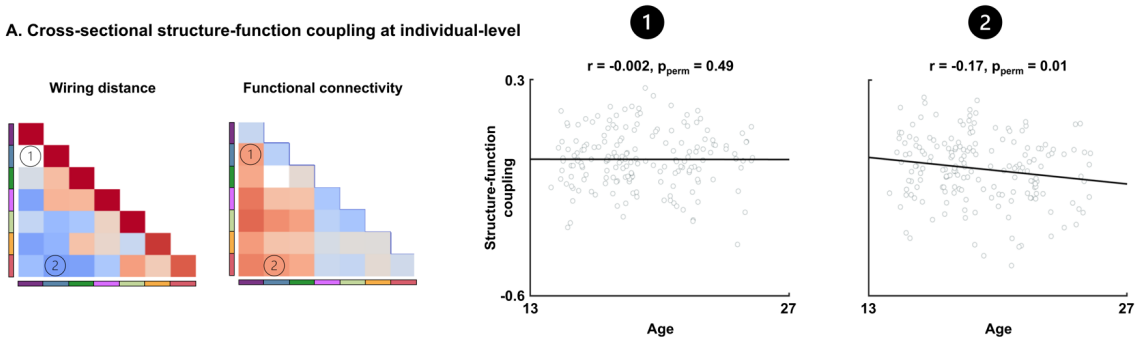


Fig. S5. Association between functional connectivity and structural differentiation during development. Cross-sectional structure-function coupling between functional connectivity and structural differentiation at different age bins. The histogram indicates a null distribution of correlation coefficients generated based on 1,000 spin tests, and the actual r -value is represented with a red bar.

A. Cross-sectional structure-function coupling at individual-level



B. Longitudinal structure-function coupling at individual-level

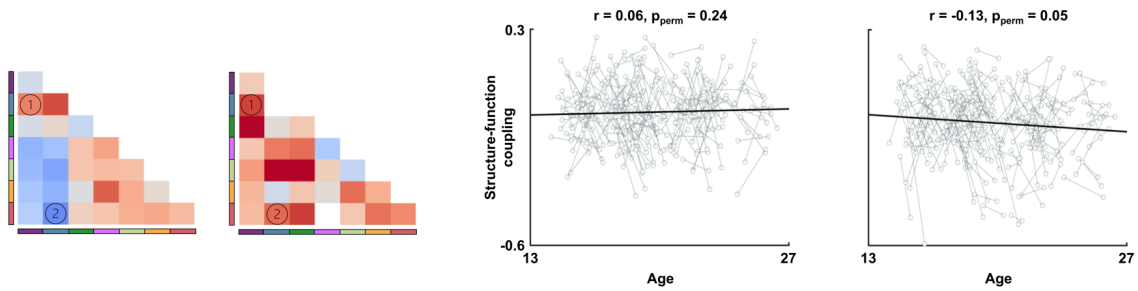


Fig. S6. Inter-individual differences in structure-function coupling. (A) The scatter plots show cross-sectional structure-function coupling between the connections that showed significant age-effects on both structural differentiation and functional connectivity, which are marked with 1 and 2. **(B)** We reported the results of longitudinal structure-function coupling.

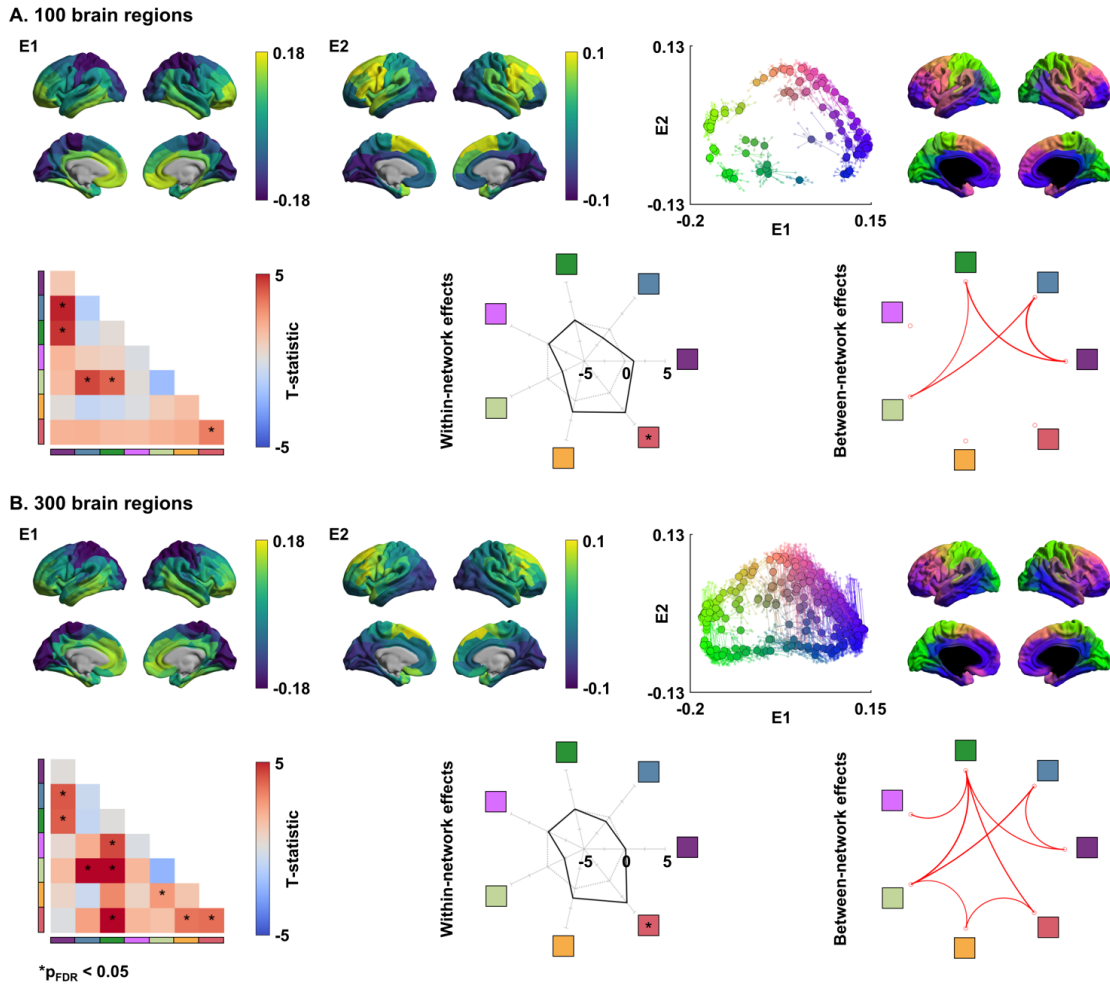


Fig. S7. Structural eigenvectors and age-effects on multiscale cortical structural differentiation using different parcellation scales. (A) Results using 100 and **(B)** 300 parcellations. Two eigenvectors (E1, E2) estimated from the cortical wiring features (top) and t-statistics of age-effects within- and between-networks (bottom) are reported. For details, see *Fig. 1*.

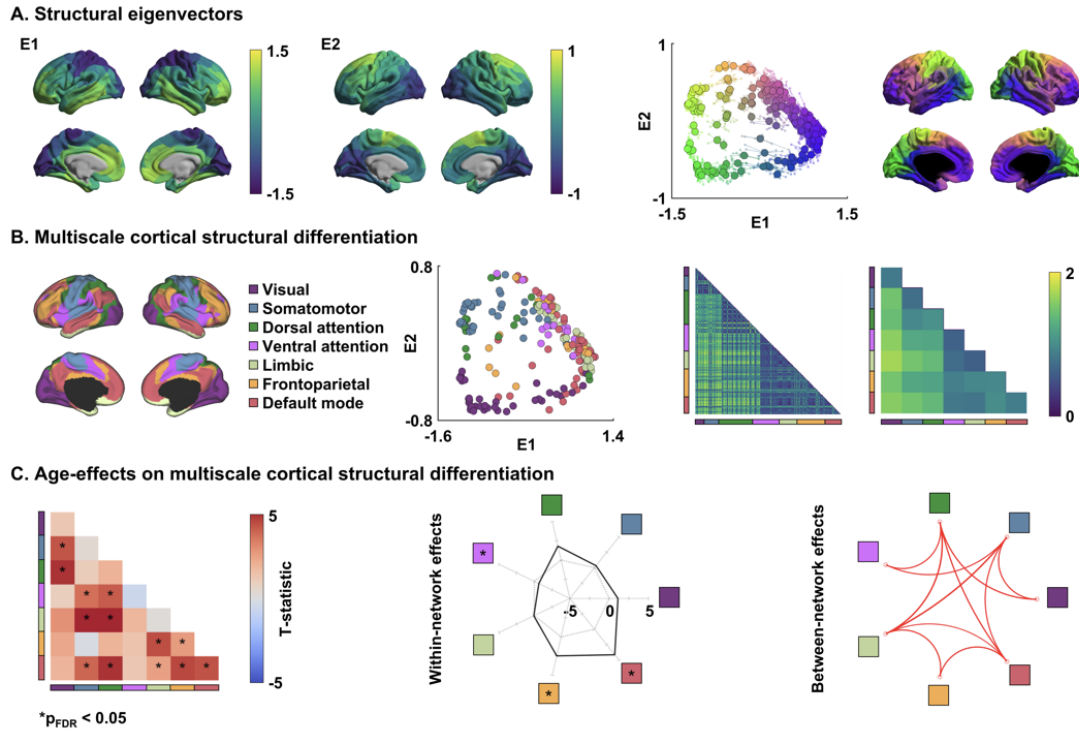


Fig. S8. Structural eigenvectors derived using principal component analysis and age-effects on multiscale cortical structural differentiation. (A) Two eigenvectors (E1, E2) estimated from the cortical wiring features. **(B)** The structural differentiation summarized based on functional communities. **(C)** The t-statistics of age-effects on structural differentiation within- and between-networks. For details, see *Fig. 1*.

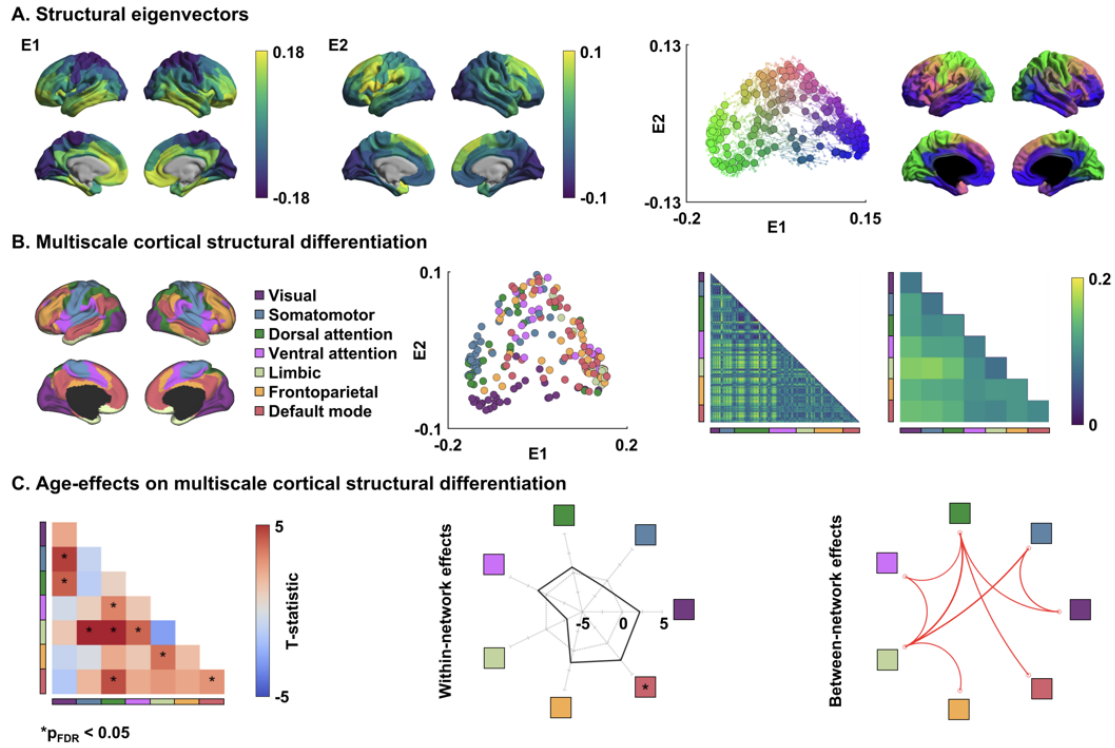
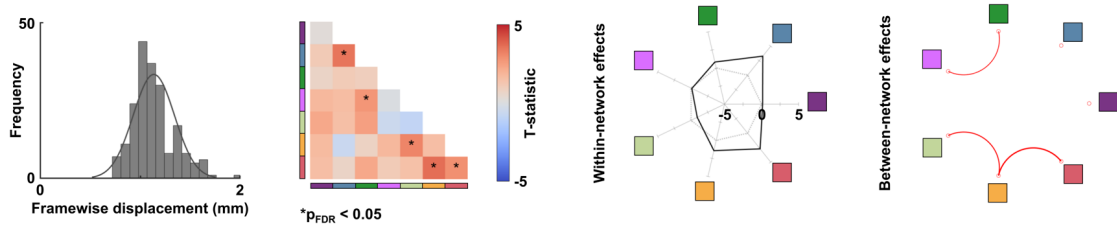


Fig. S9. Structural eigenvectors and age-effects on multiscale cortical structural differentiation using Schaefer 200 parcellation. (A) Two eigenvectors (E1, E2) estimated from the cortical wiring features. (B) The structural differentiation summarized based on functional communities. (C) The t-statistics of age-effects on structural differentiation within- and between-networks. For details, see Fig. 1.

A. Age-effects on multiscale cortical structural differentiation of low head motion participants



B. Age-effects on functional connectivity of low head motion participants

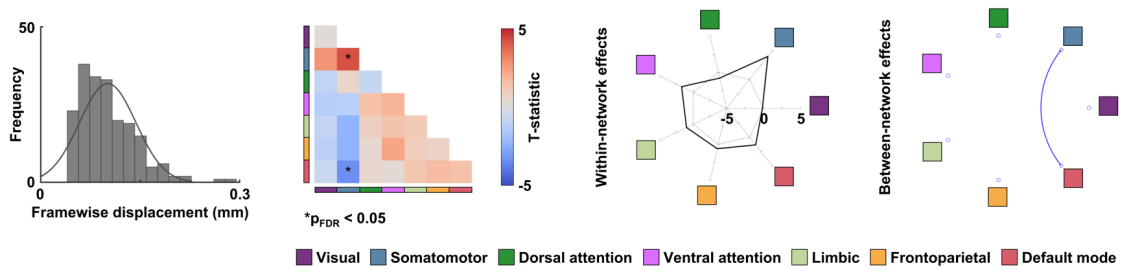


Fig. S10. Age-effects on structural and functional measures based on participants with low head motion. (A) Age-effects on structural differentiation and (B) functional connectivity. The histograms indicate frequency of framewise displacement. For details, see Fig. 1 and 2.

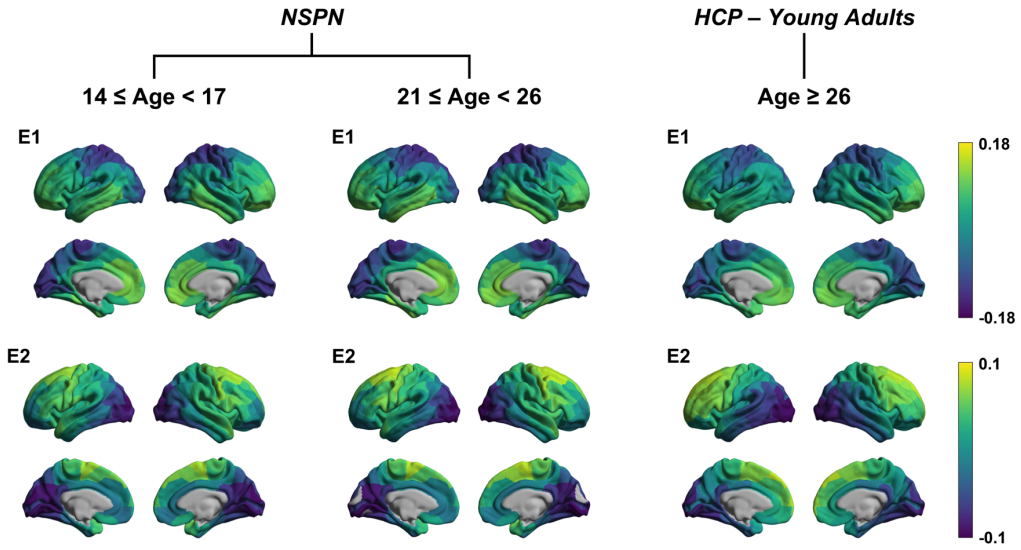
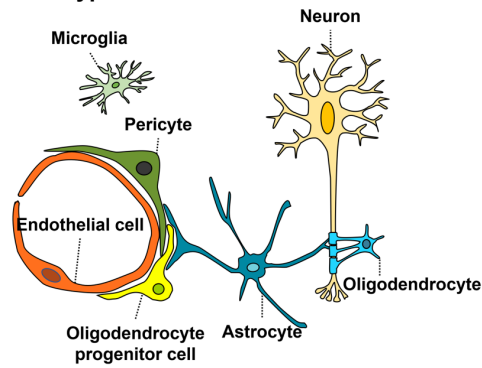


Fig. S11. Structural eigenvectors of different age bins. We estimated two eigenvectors (E1, E2) from cortical wiring features using (A) younger (age < 17) and (B) older (age ≥ 21) adolescence from the *NSPN* dataset, and (C) those using the *HCP – Young Adults* dataset. Abbreviations: *NSPN*, the Neuroscience in Psychiatry Network; *HCP*, Human Connectome Project.

A. Schema of cell-types



B. Cell-type specific expression analysis

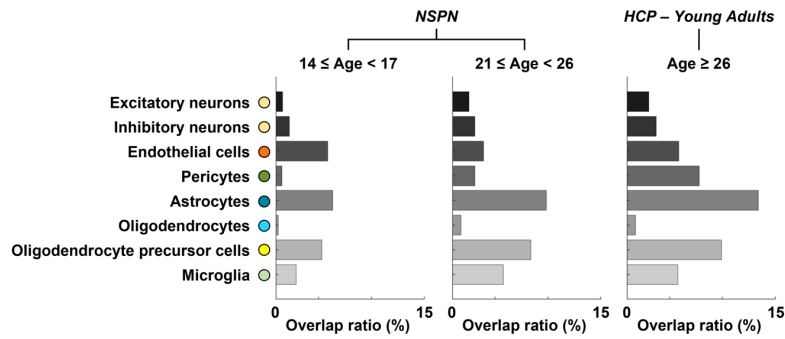


Fig. S12. Transcriptomic analysis. (A) Schema of different cell types and (B) overlap ratio between cell-type specific genes and the genes associated with multiscale cortical wiring of younger (age < 17) and older (age ≥ 21) adolescence from the NSPN dataset, and young healthy adults from the HCP – Young Adults database. Abbreviations: NSPN, the Neuroscience in Psychiatry Network; HCP, Human Connectome Project.

Neuroscience in Psychiatry Network (NSPN) Consortium author list

Principal investigators:

Edward Bullmore (CI from 01/01/2017)^{1,2,3}; Raymond Dolan^{4,5}; Ian Goodyer (CI until 01/01/2017)¹; Peter Fonagy⁶; Peter Jones¹

NSPN (funded) staff:

Michael Moutoussis^{4,5}; Tobias Hauser^{4,5}; Sharon Neufeld¹; Rafael Romero-Garcia^{1,2}; Michelle St Clair¹; Petra Vértes^{1,2}; Kirstie Whitaker^{1,2}; Becky Inkster¹; Gita Prabhu^{4,5}; Cinly Ooi¹; Umar Toseeb¹; Barry Widmer¹; Junaid Bhatti¹; Laura Willis¹; Ayesha Alrumaithi¹; Sarah Birt¹; Aislinn Bowler⁵; Kalia Cleridou⁵; Hina Dadabhoy⁵; Emma Davies¹; Ashlyn Firkins¹; Sian Granville⁵; Elizabeth Harding⁵; Alexandra Hopkins^{4,5}; Daniel Isaacs⁵; Janchai King⁵; Danae Kokorikou^{5,6}; Christina Maurice¹; Cleo McIntosh¹; Jessica Memarzia¹; Harriet Mills⁵; Ciara O'Donnell¹; Sara Pantaleone⁵; Jenny Scott¹; Beatrice Kiddle¹; Ela Polek¹

Affiliated scientists:

Pasco Fearon⁶; John Suckling¹; Anne-Laura van Harmelen¹; Rogier Kievit^{4,7}; Sam Chamberlain¹

¹*Department of Psychiatry, University of Cambridge, United Kingdom*

²*Behavioural and Clinical Neuroscience Institute, University of Cambridge, United Kingdom*

³*ImmunoPsychiatry, GlaxoSmithKline Research and Development, United Kingdom*

⁴*Max Planck University College London Centre for Computational Psychiatry and Ageing Research, University College London, UK*

⁵*Wellcome Centre for Human Neuroimaging, University College London, United Kingdom*

⁶*Research Department of Clinical, Educational and Health Psychology, University College London, United Kingdom*

⁷*Medical Research Council Cognition and Brain Sciences Unit, University of Cambridge, United Kingdom*

SI References

1. B. T. T. Yeo, *et al.*, The organization of the human cerebral cortex estimated by intrinsic functional connectivity. *J. Neurophysiol.* **106**, 1125–1165 (2011).