

Microstructural development of human primary and nonprimary auditory cortex during the perinatal period

Brian B. Monson,^{1,2,3} Einat Liebenthal,² Simon K. Warfield,³ Terrie E. Inder,² Jeffrey J. Neil³

¹Department of Speech and Hearing Science, College of Applied Health Sciences, University of Illinois at Urbana-Champaign
²Department of Pediatric Newborn Medicine, Brigham & Women's Hospital, Harvard Medical School; ³Boston Children's Hospital, Harvard Medical School



Introduction



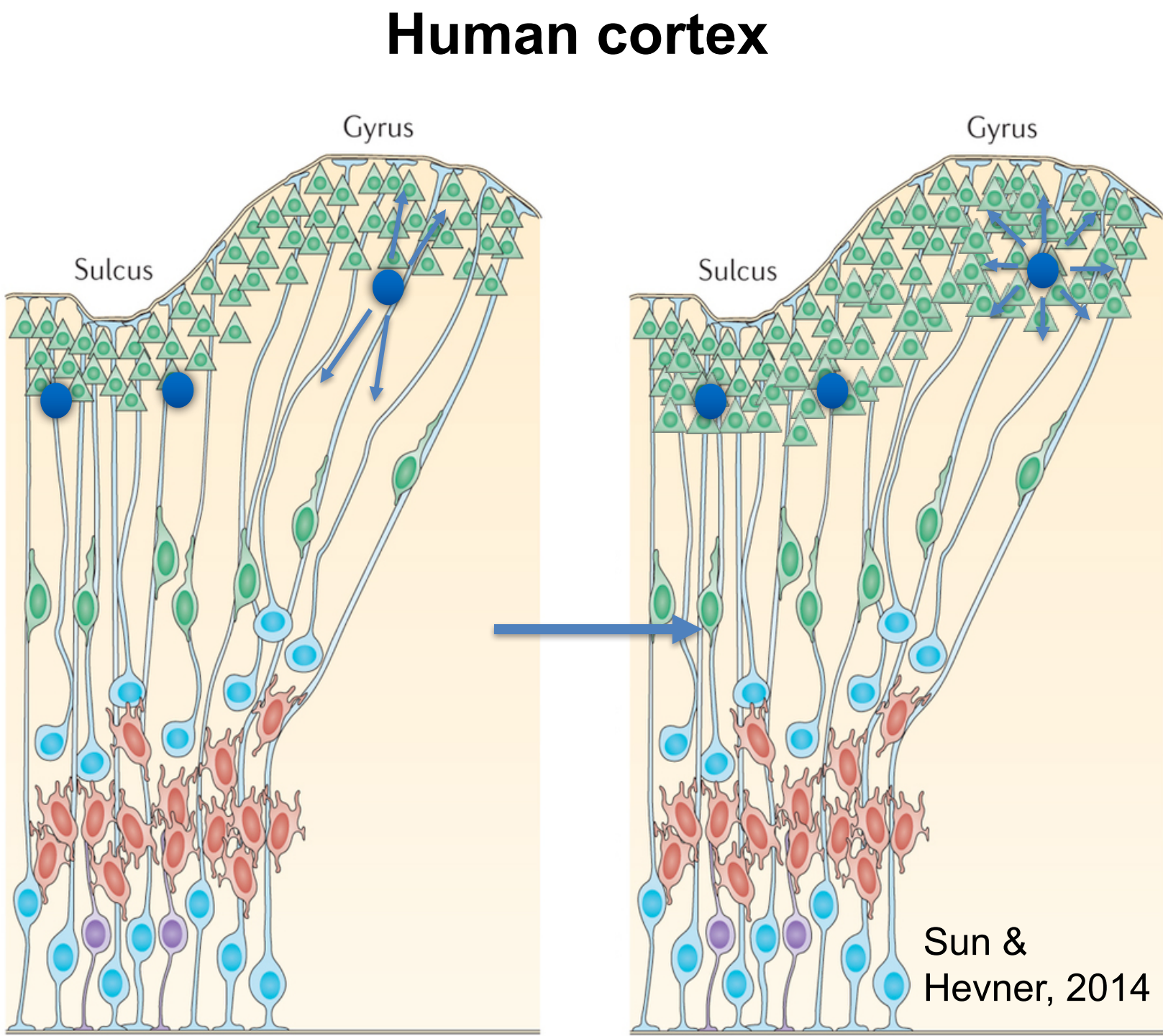
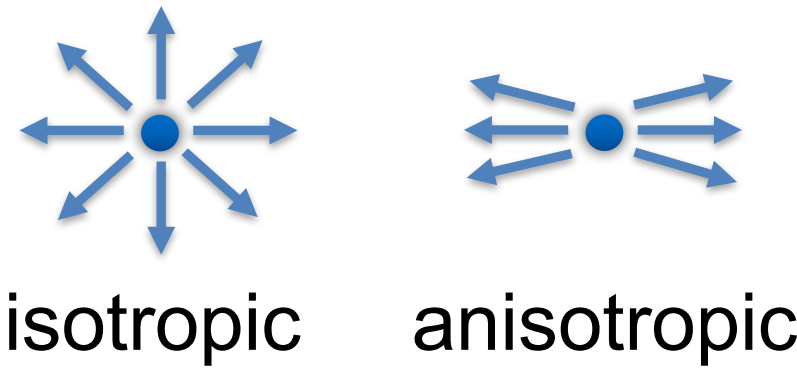
The human auditory system undergoes a period of rapid neurodevelopment *in utero*.
Postmortem histology has provided insight into the structural maturation of auditory cortex during this period. Neuroimaging can reveal structural maturational processes *in vivo*.

Aim

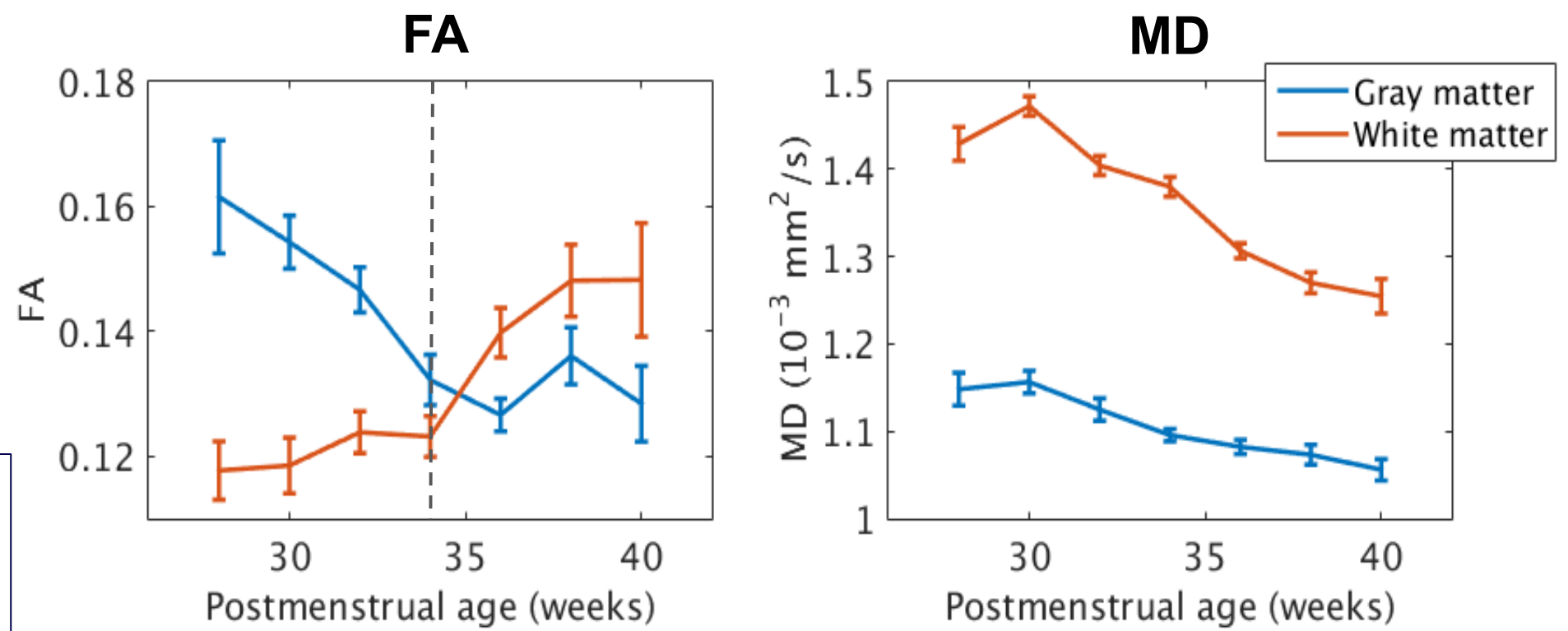
- To characterize the maturational timeline of auditory cortex during the perinatal period.
- To compare maturational timelines for primary (pAC) and nonprimary auditory cortical (nAC) regions.
- To examine the effect of preterm birth on auditory cortex maturation.

Diffusion MRI and Cortical Maturation

Diffusion MRI measures random motion of water molecules within tissue. Parameters obtained include: fractional anisotropy (FA) and mean diffusivity (MD).



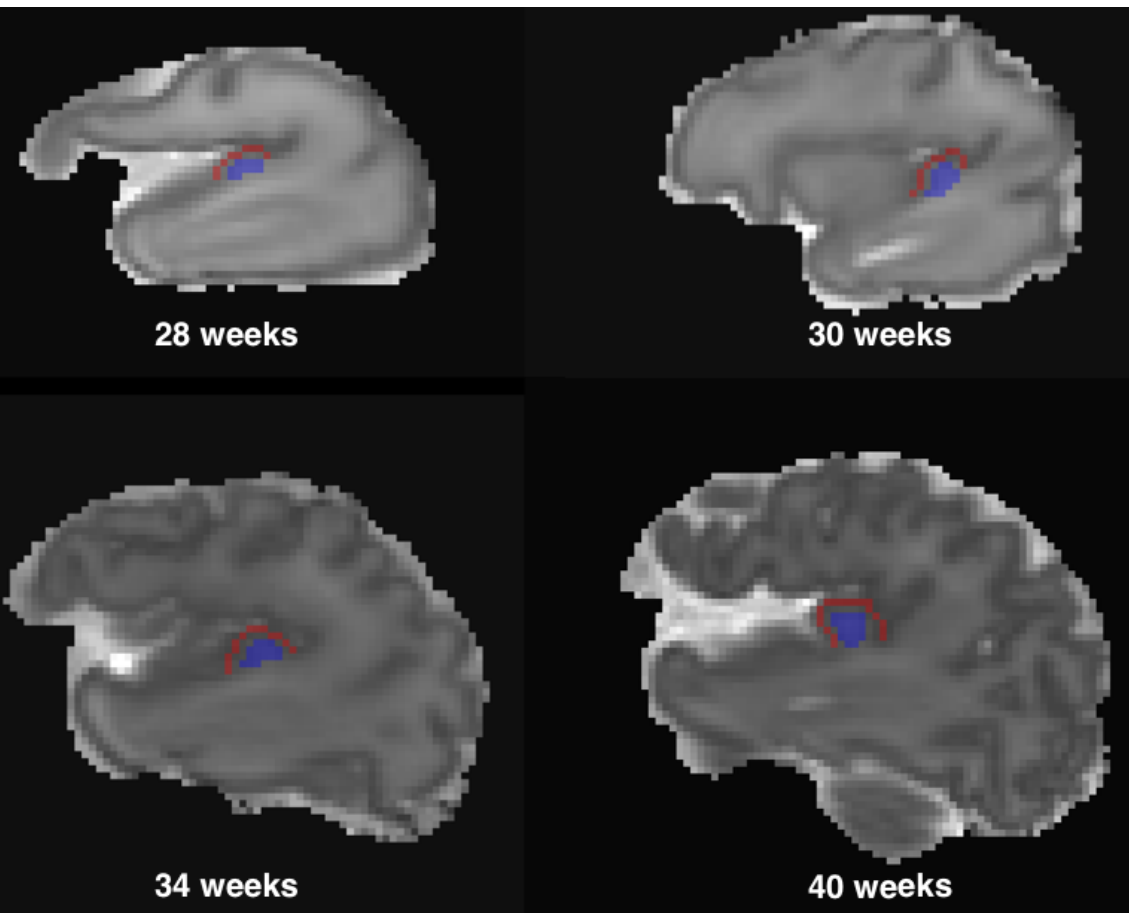
In developing *gray* matter, FA starts high (due to the radial structure of radial glia) and *decreases* with time; MD also *decreases* with time due to increasing neuron density and decreasing water content.
In developing *white* matter, FA starts low and *increases* with time due to myelination and precursors to myelination; MD *decreases* with time due to increasing cell density and decreasing water content.



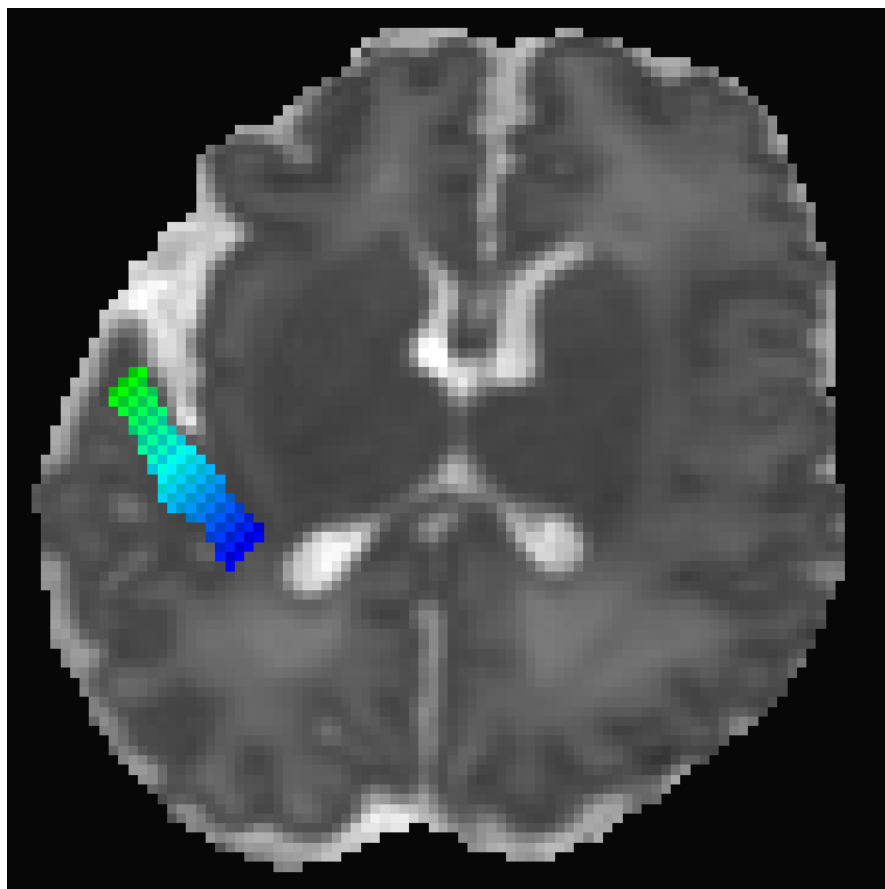
Acknowledgments: The authors wish to thank Daniel Polley, PhD, Margaret Kenna, MD, and Christopher Smyser, MD for helpful discussions regarding the data. We thank Abraham Brownell for assistance with the analysis.
Funding: This work was funded in part by NIH. Conference travel support provided by NIH/NIDCD.

Method

- Usable data collected for 90 preterm infants (birth gestational age <30 weeks) who underwent MR imaging up to 4 times during NICU stay, from 26 to 42 weeks PMA
- Multi-shell acquisition at 48 directions
- Maturation of gray and adjacent white matter along the axis of left hemisphere Heschl's gyrus measured *in vivo* using diffusion weighted image analysis

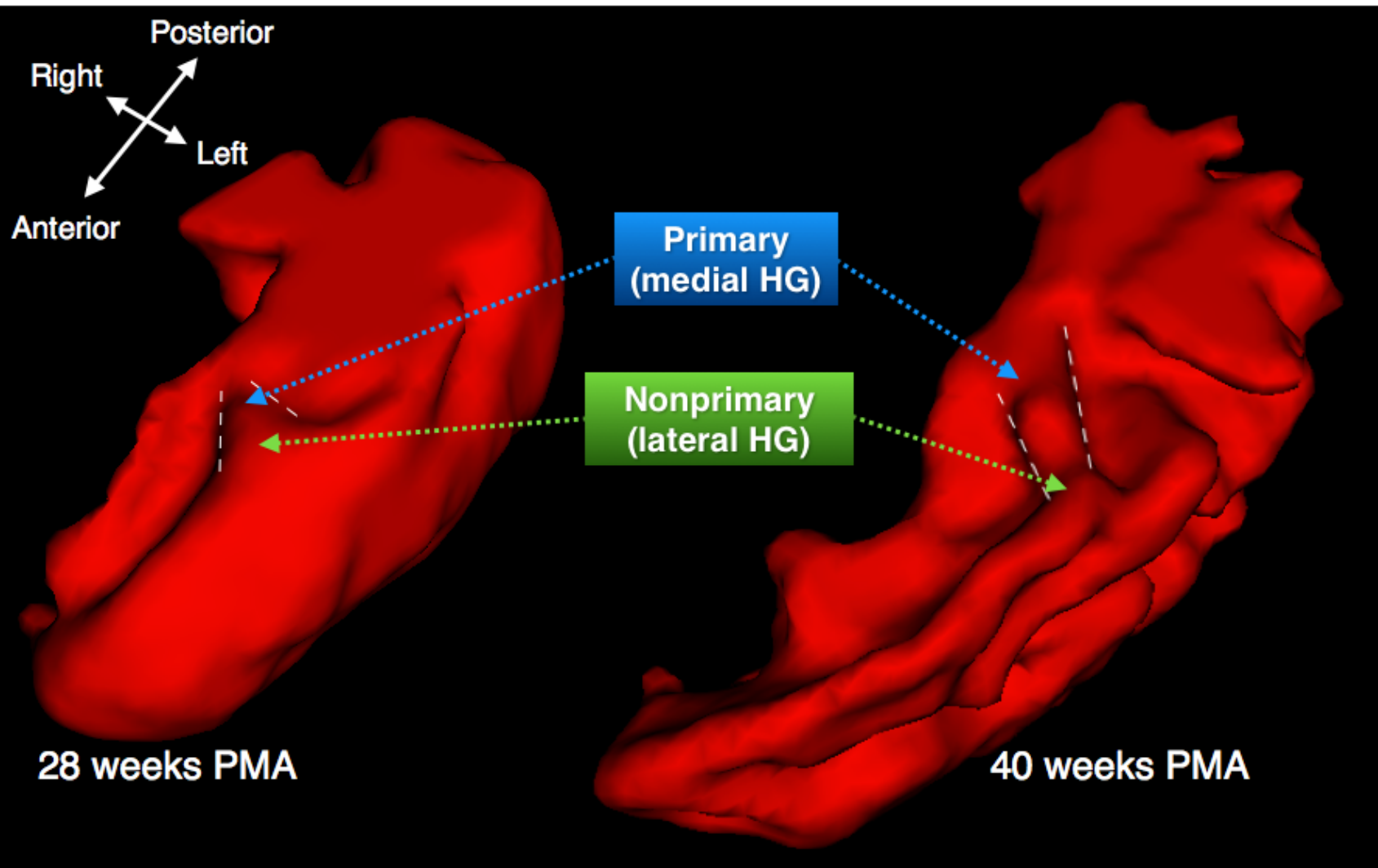


Regions of interest located in Heschl's gyrus at different ages

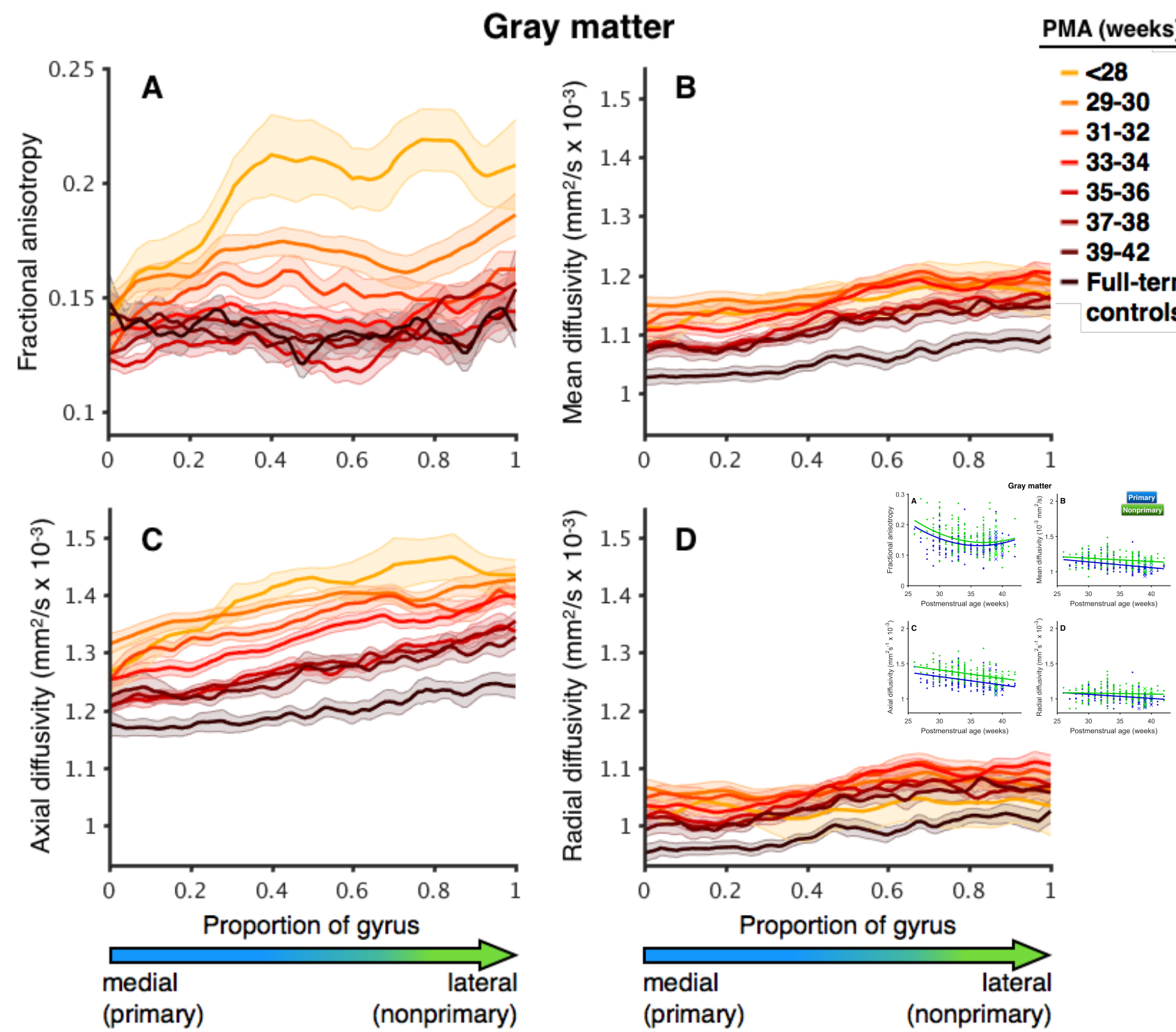


Putative transition from *primary* to *nonprimary* auditory cortex

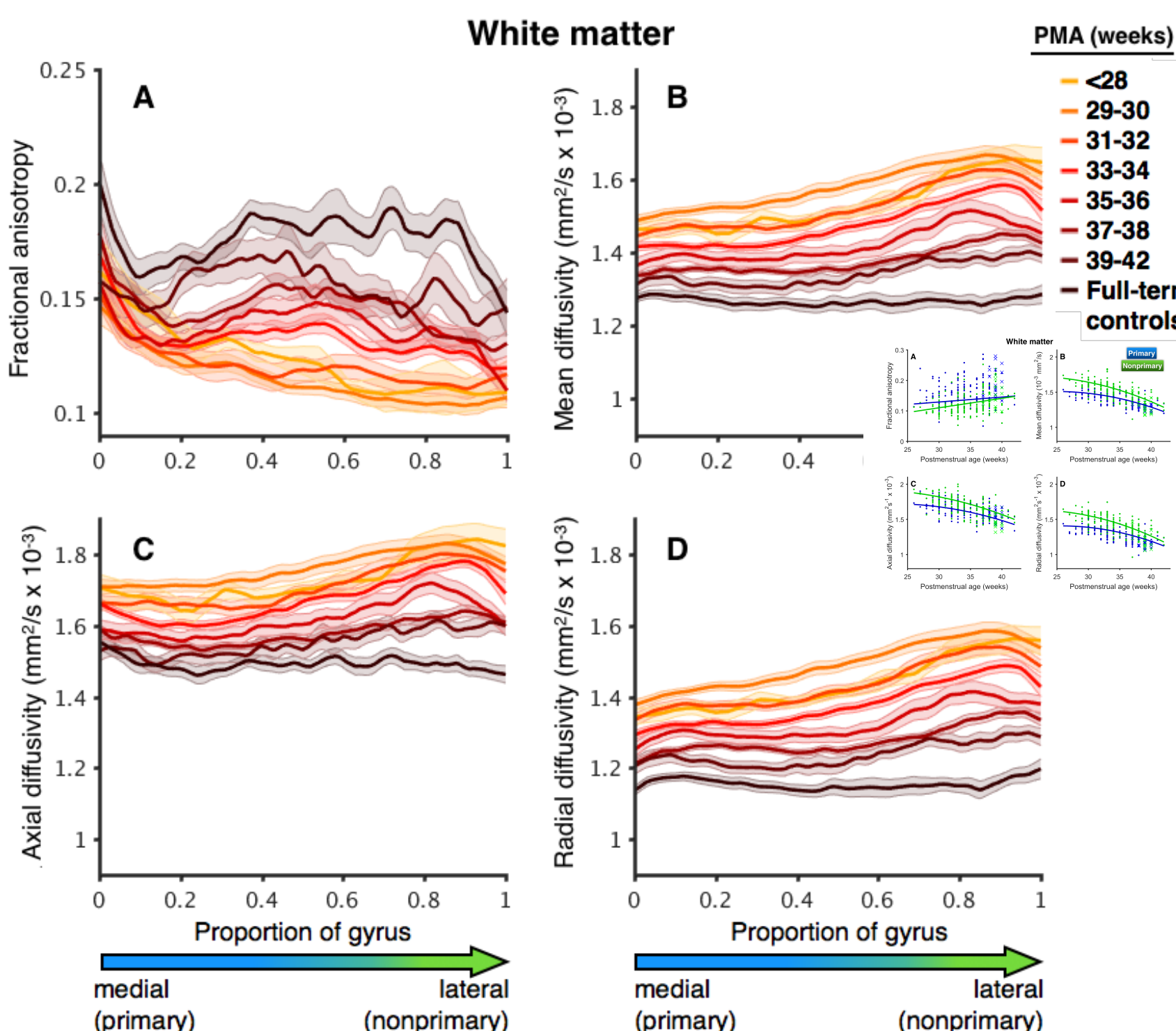
Results



Development of Heschl's gyrus (HG) macrostructure from 28 weeks to 40 weeks postmenstrual age (PMA)

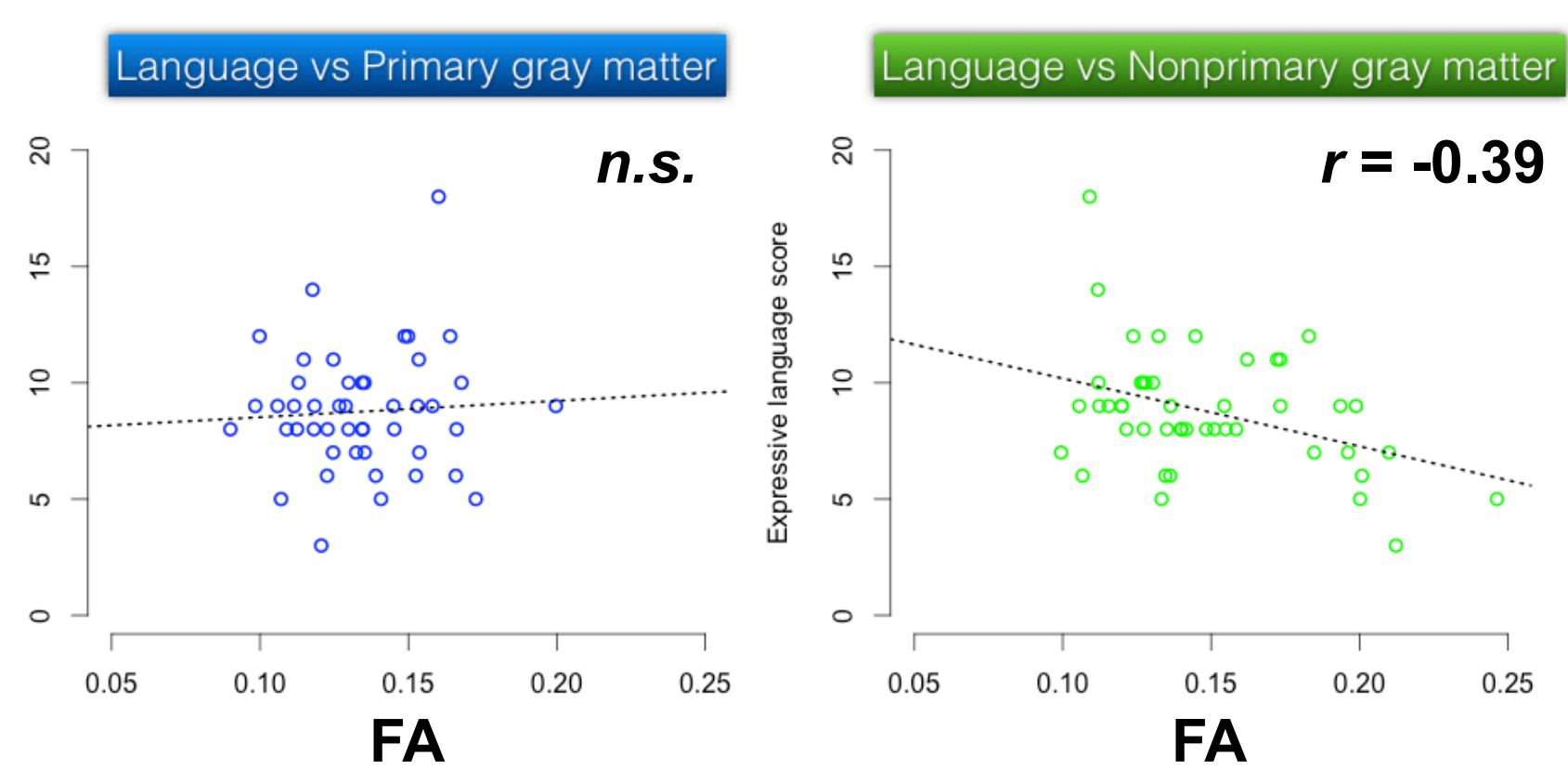


Gray matter fractional anisotropy and mean/axial/radial diffusivity changes along Heschl's gyrus by age group



White matter fractional anisotropy and mean/axial/radial diffusivity changes along Heschl's gyrus by age group

Results (continued)



Relationship between gray matter maturation in infancy (as measured by FA) and language development at age 2 years.

Conclusions

- pAC matures in advance of nAC.
- As a result, nAC exhibits more rapid maturational changes in diffusion measures from 26 to 42 weeks PMA than pAC.
- Premature birth is associated with higher diffusivity values in pAC and nAC at term age, consistent with a delay in maturation of cortical microstructure in preterm infants.
- A rapid maturation rate during the perinatal period may render nAC white matter more vulnerable to disruption or injury.
- nAC (but *not* pAC) maturation in infancy is associated with language development at age 2.

References

Ball *et al.* (2013) Development of cortical microstructure in the preterm human brain. *Proceedings of the National Academy of Sciences*, 110 (23):9541-9546.
McKinstry *et al.* (2002) Radial organization of developing preterm human cerebral cortex revealed by non-invasive water diffusion anisotropy MRI. *Cerebral Cortex*, 12:1237-1243.
Smyser *et al.* (2015) Cortical gray and adjacent white matter demonstrate synchronous maturation in very preterm infants. *Cerebral Cortex* (advance access online). doi: 10.1093/cercor/bhv164
Sun & Hevner (2014) Growth and folding of the mammalian cerebral cortex: from molecules to malformations. *Nature Reviews Neuroscience*, 15:217-232.

