

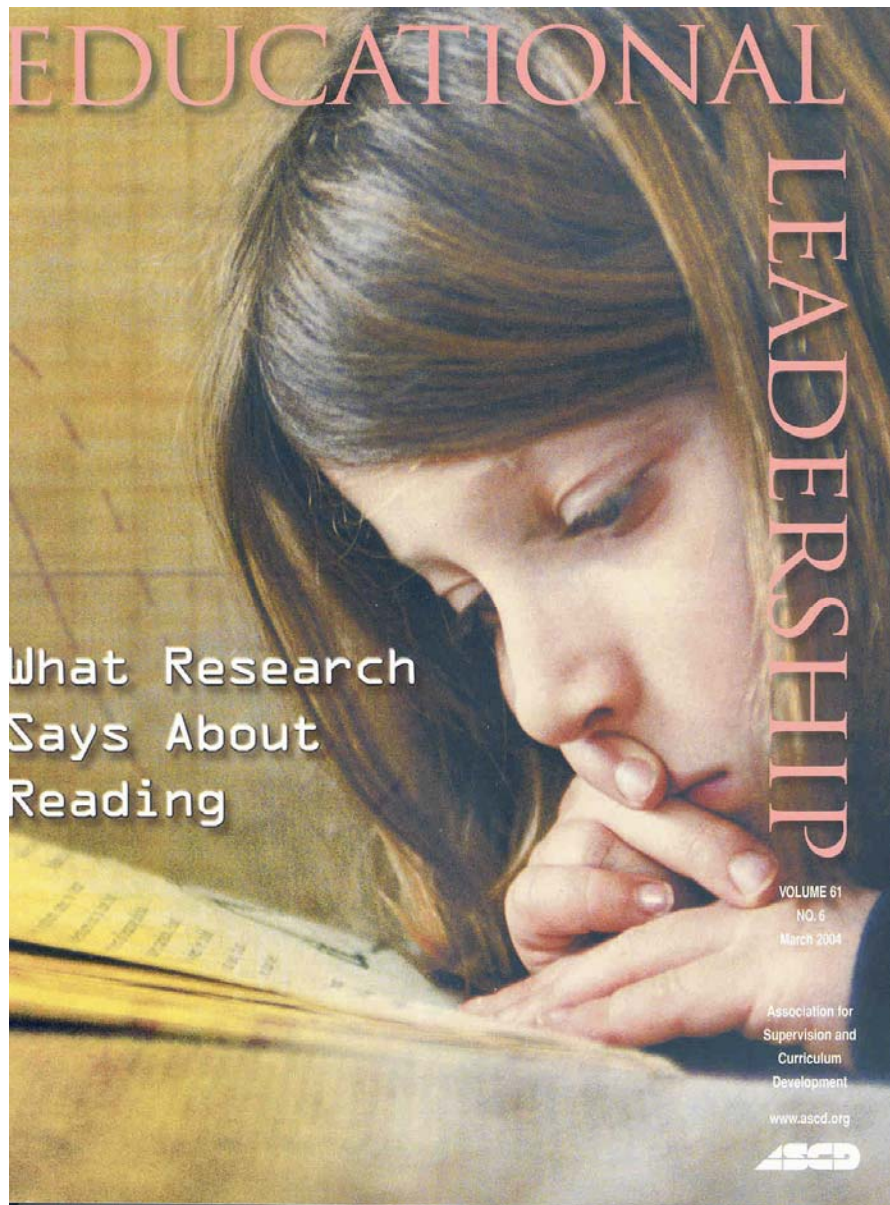
# Reading Circuitry in the Child's Developing Brain

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*Stanford University*

## Collaborators

*Robert Dougherty, Michal Ben-Shachar, Gayle Deutsch, Arvel Hernandez*



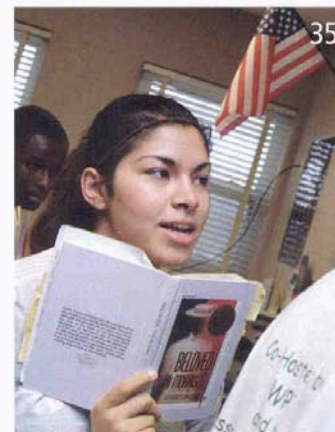


# EDUCATIONAL LEADERSHIP

March 2004 • Volume 61 • Number 6

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# Consensus: Phonology and reading

“... the specification of the role of phonological processing in the earliest stages of reading acquisition is one of the more notable scientific success stories of the last decade.” (Stanovich, 1991, p. 78)

“To my mind, the discovery and documentation of the importance of phonemic awareness ... is the single most powerful advance in the science and pedagogy of reading this century.” (Adams, 1990).

# Phonemic awareness and reading

- Hearing, distinguishing and manipulating the sounds in words (also phonological awareness)
- Claimed to be the chief causal factor in early reading achievement
- Hope: If we train phonemic awareness, children will become better readers (also skills-training, decoding)

**Bradley, L. & Bryant, P. (1983) Categorizing sounds and learning to read - a causal connection. *Nature*, 301, 419-421.**

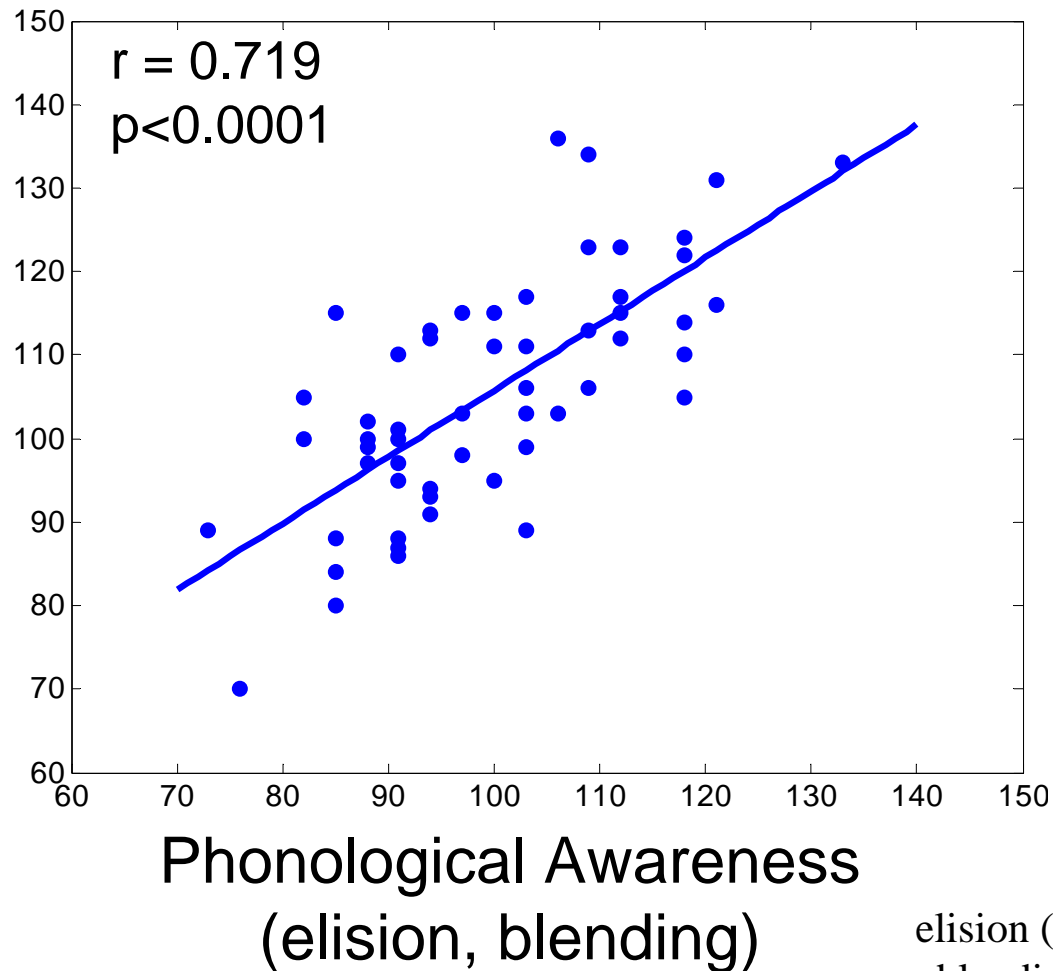
# Example Tasks

- Deletion: Remove the /r/ from rat. What do you have left?
- Add a /b/ to rat. What do you have?
- Rhyming
- Oddity: bud, bun, bus, rug
- Pronouncing pseudowords (dif, giz, dop, blif)

# Word reading correlates with phonological awareness

(Ben-Shachar, Deutsch, Dougherty, Wandell)

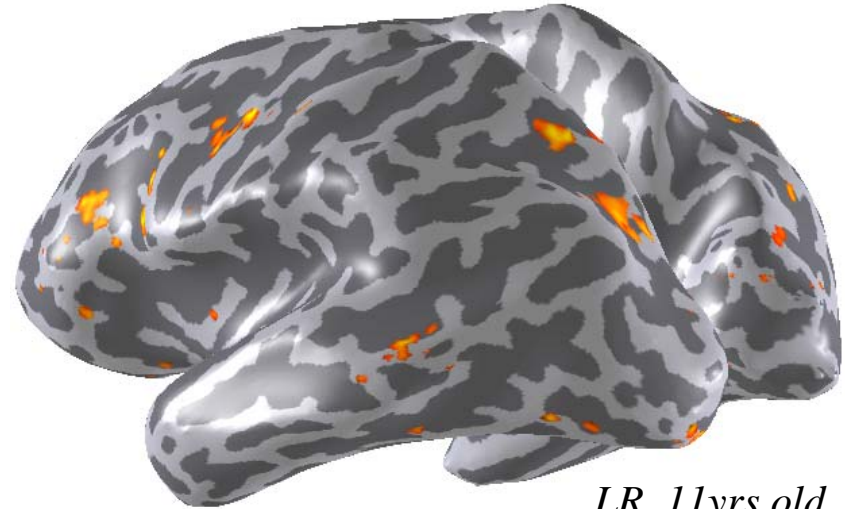
Reading  
(Word ID)  
read aloud:  
'..together,..  
enough, ..'



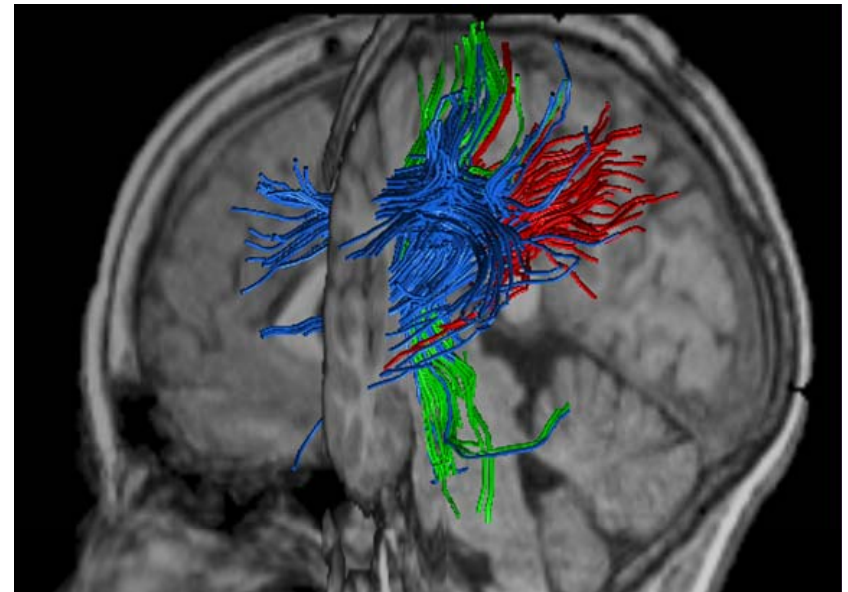
elision ( strain-/r/ = stain)  
blending ( /t+/oi/ = toy)

# Neural Basis of Reading

- Most variance explained by social factors and general cognitive ability
- Significant variance remains...
  - What are the specific neurological factors?
- Identify biological correlates
  - Explain individual differences
  - Predict & intervene before reading failure

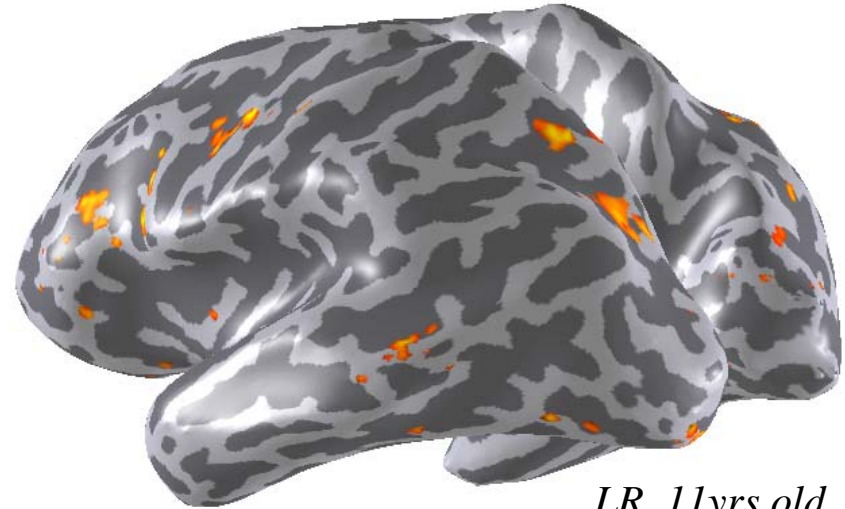


*LR, 11yrs old*

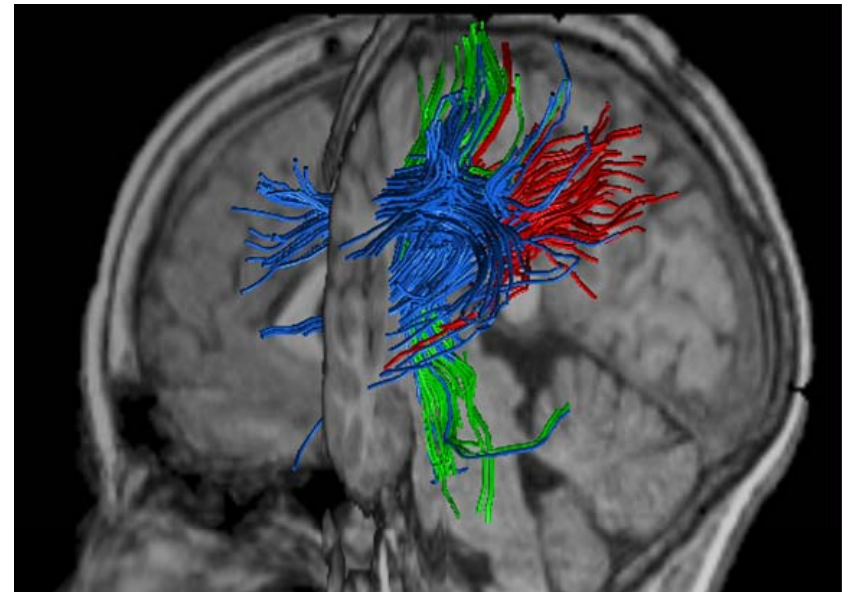


# Specific Reading Pathways

- Magnocellular hypothesis
- Temporo-parietal junction
- OTS - fMRI
- Diffusion Tensor Imaging



*LR, 11yrs old*



Lightness  
Perception  
(Lotto and Purves)



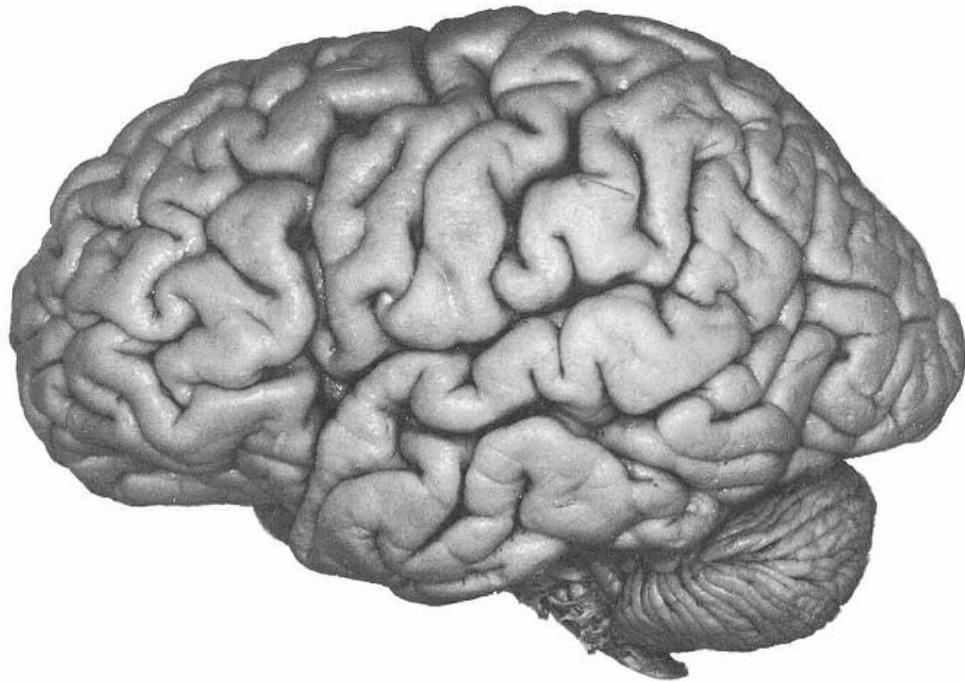
# The Astonishing Hypothesis

(Francis Crick; Braitenberg and Schutz, 1991)

*Neurons are the computational elements*

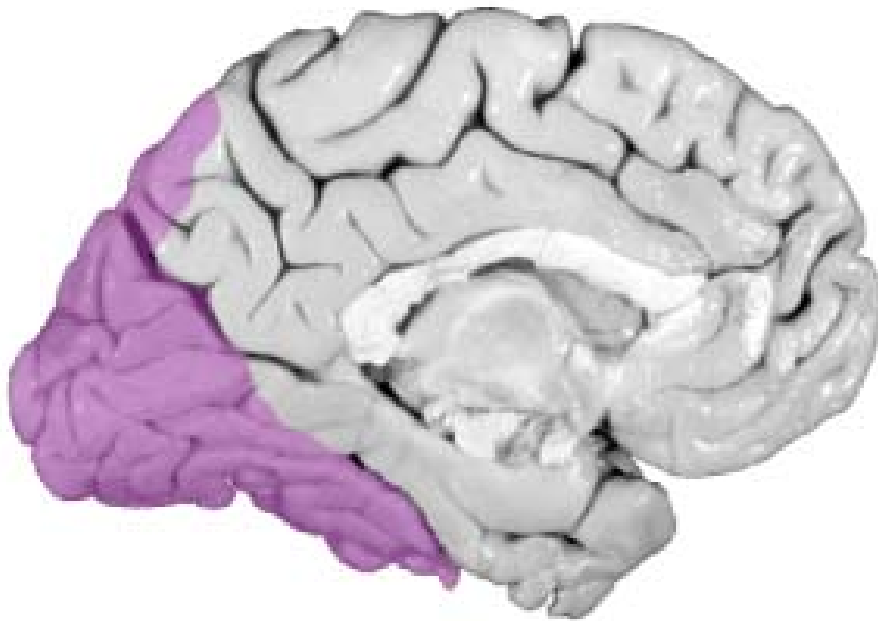
*White matter connects the neurons via axons*

*The connection is called the synapse*

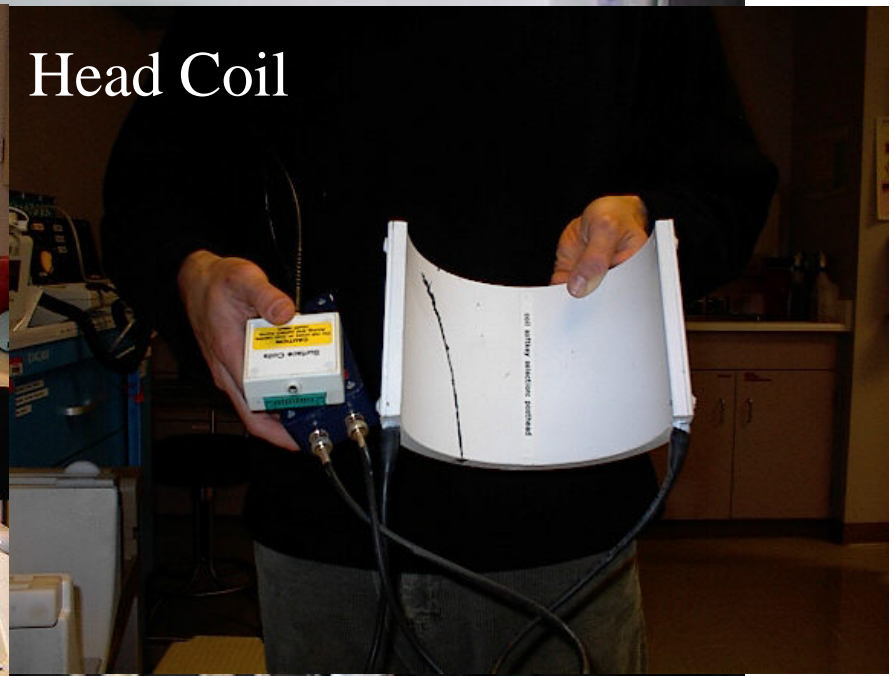
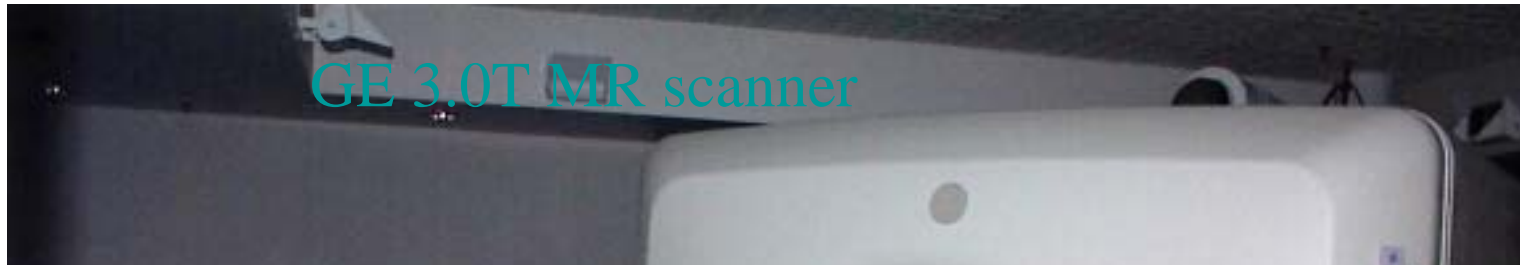


- Neurons:  $10^{11}$
- Synapses:  $10^{14}$
- Synapses/neuron  $10^3$
- Surface area of each hemisphere:  $25 \times 30 \text{ cm}^2$
- Most connections are local (10-100  $\mu\text{m}$ ); some span many cm
- Neurons/ $\text{mm}^3$ :  $10^4$ - $10^6$
- Axon length/ $\text{mm}^3$ : 3 km

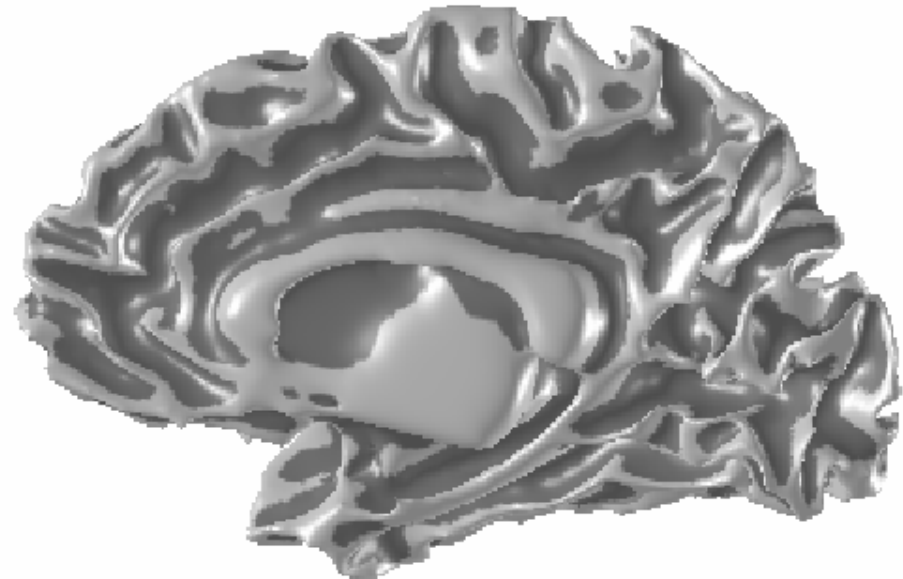
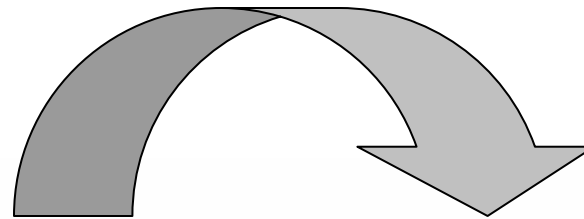
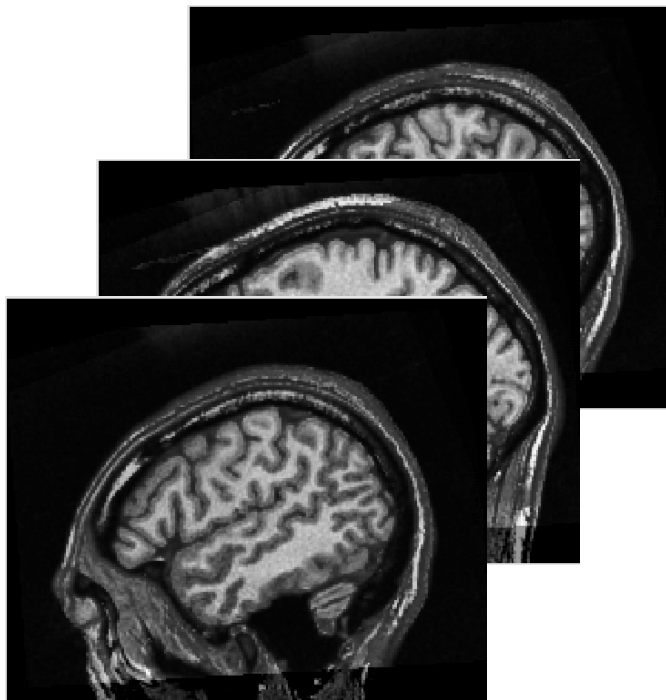
Visual cortex is about 20% of human neo-cortex



# FMRI Instrumentation



# Gray/white surface boundary



Brain inflation

# There Is An Increase In Oxygenated Blood Flow To Active Regions of Cortex

J.F. Fulton, M.D. (1928)

## Operation

On turning down a left occipital bone flap, a large angry-looking

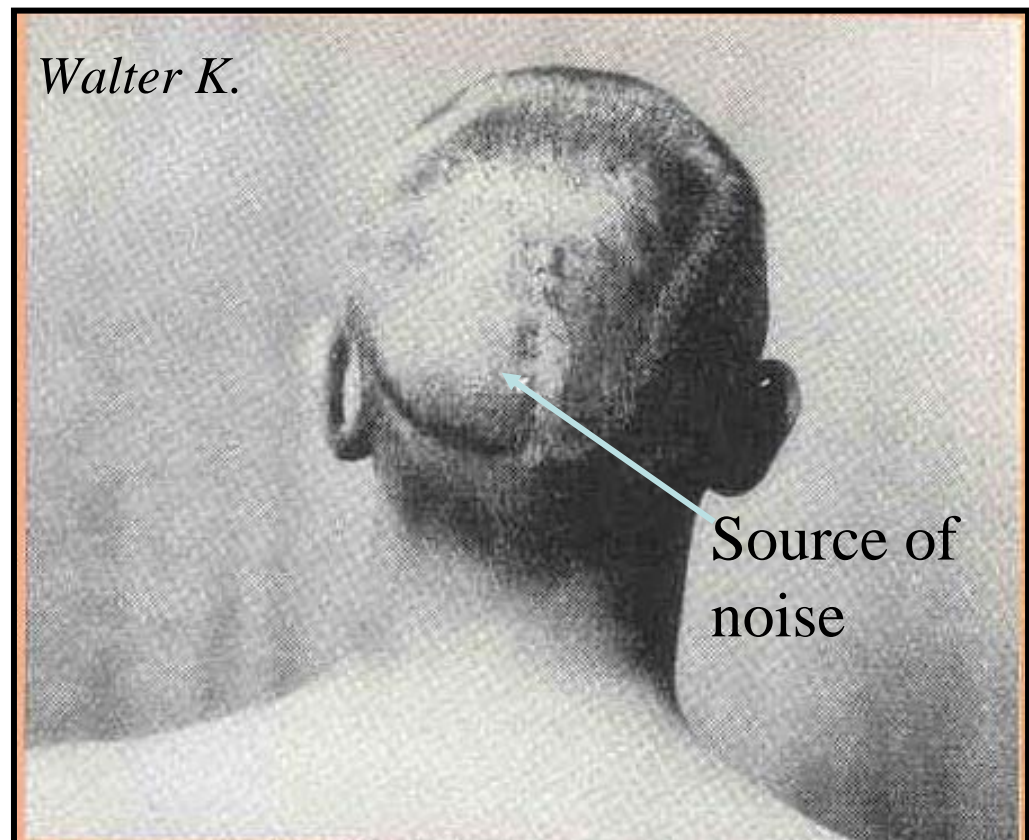
angioma arteriale racemosum of the left occ. Lobe was disclosed which extensively involved the visual cortex. The haemorrhage

decreased by the bone flap was so excessive that the operation had to be abandoned without touching the

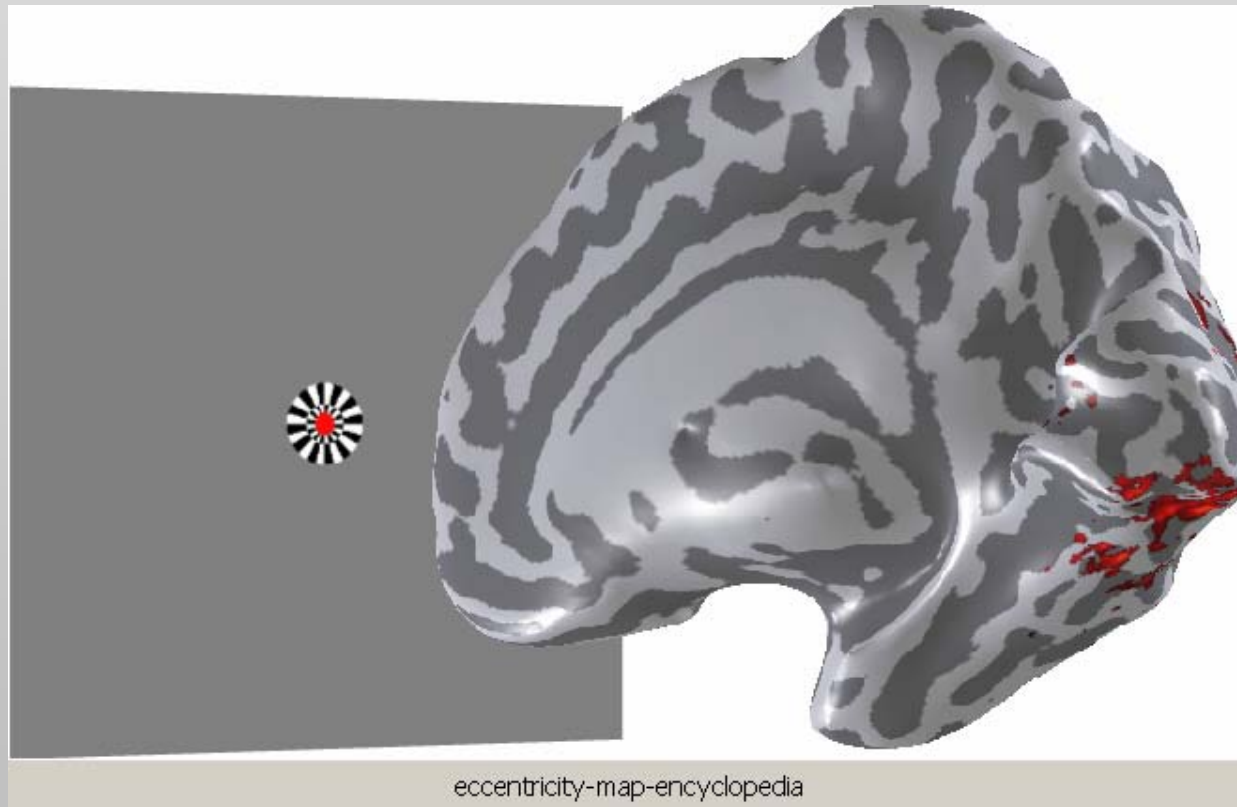
tumour. A decompression,

however, was made. The patient was discharged ... with greatly

improved vision.

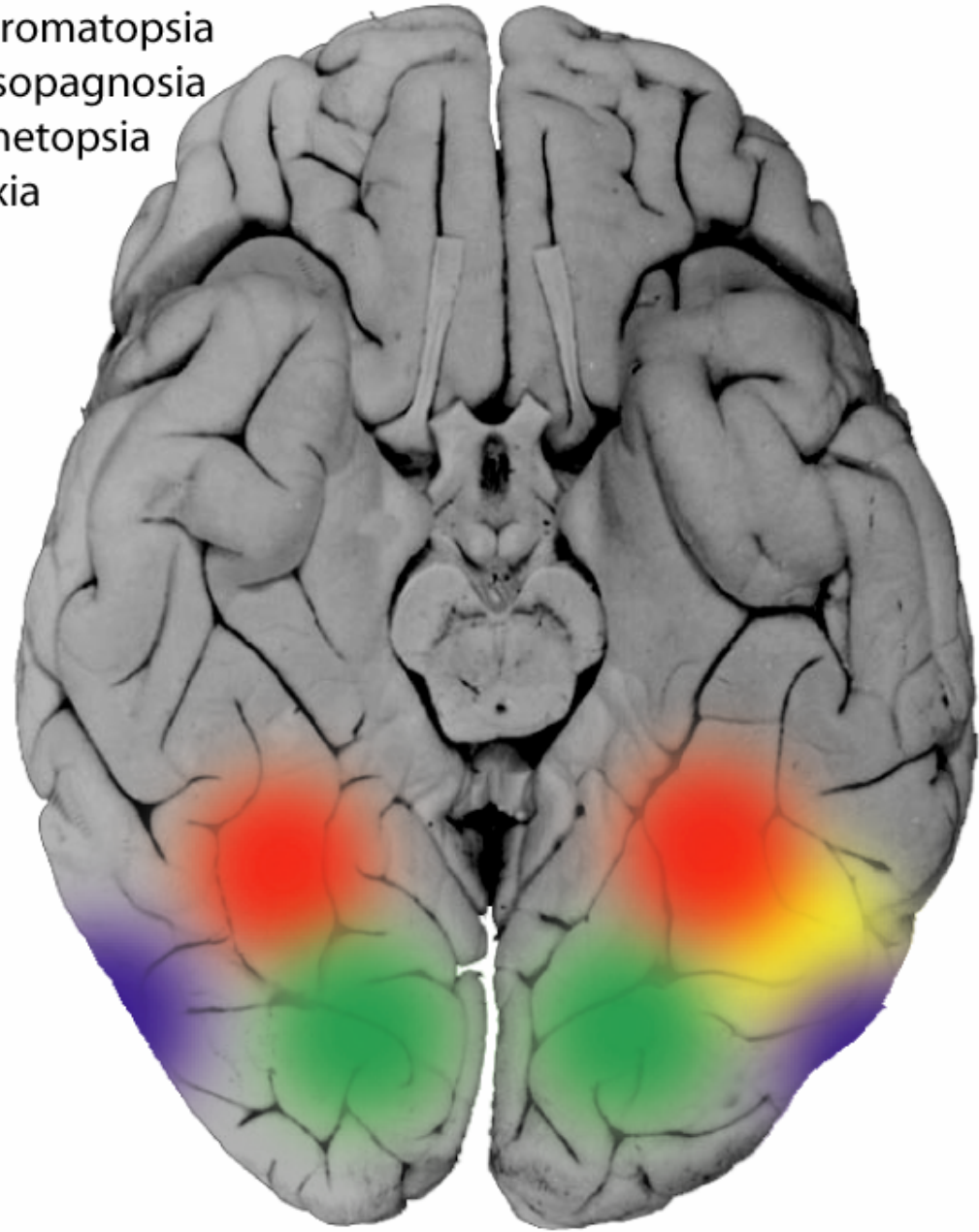


Using magnetic resonance imaging,  
we can measure responses in  
human visual cortex



Localized  
cortical  
damage  
produces  
very  
specific  
visual  
dysfunction

- Achromatopsia
- Prosopagnosia
- Akinetopsia
- Alexia

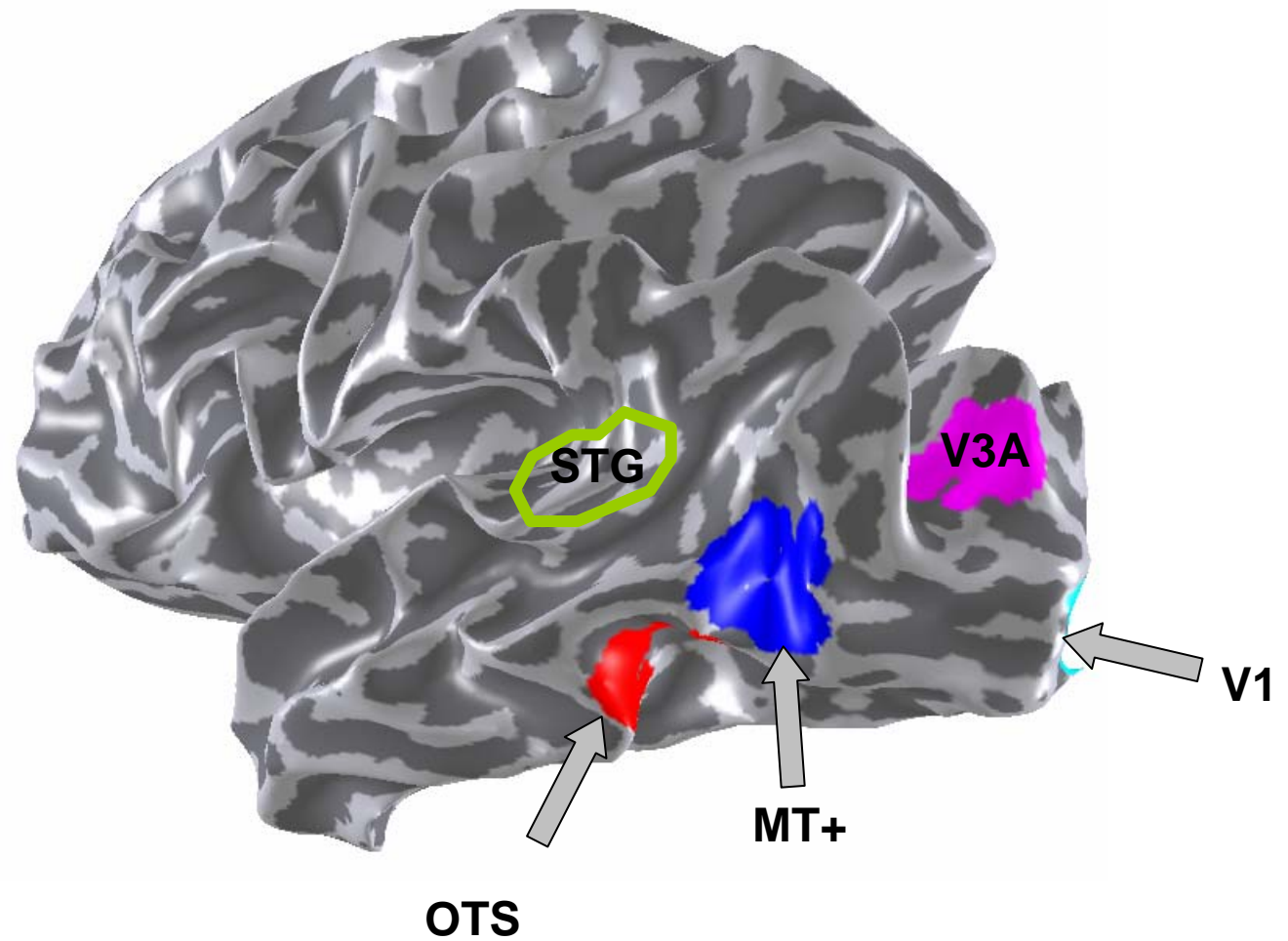
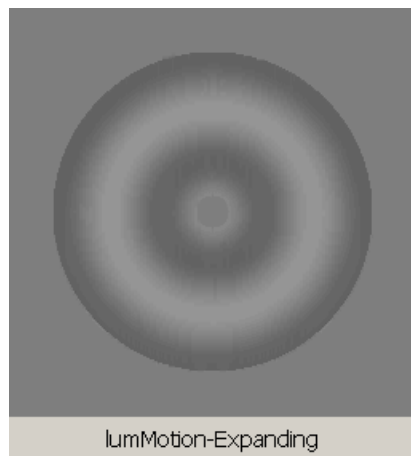


# Face-blindness



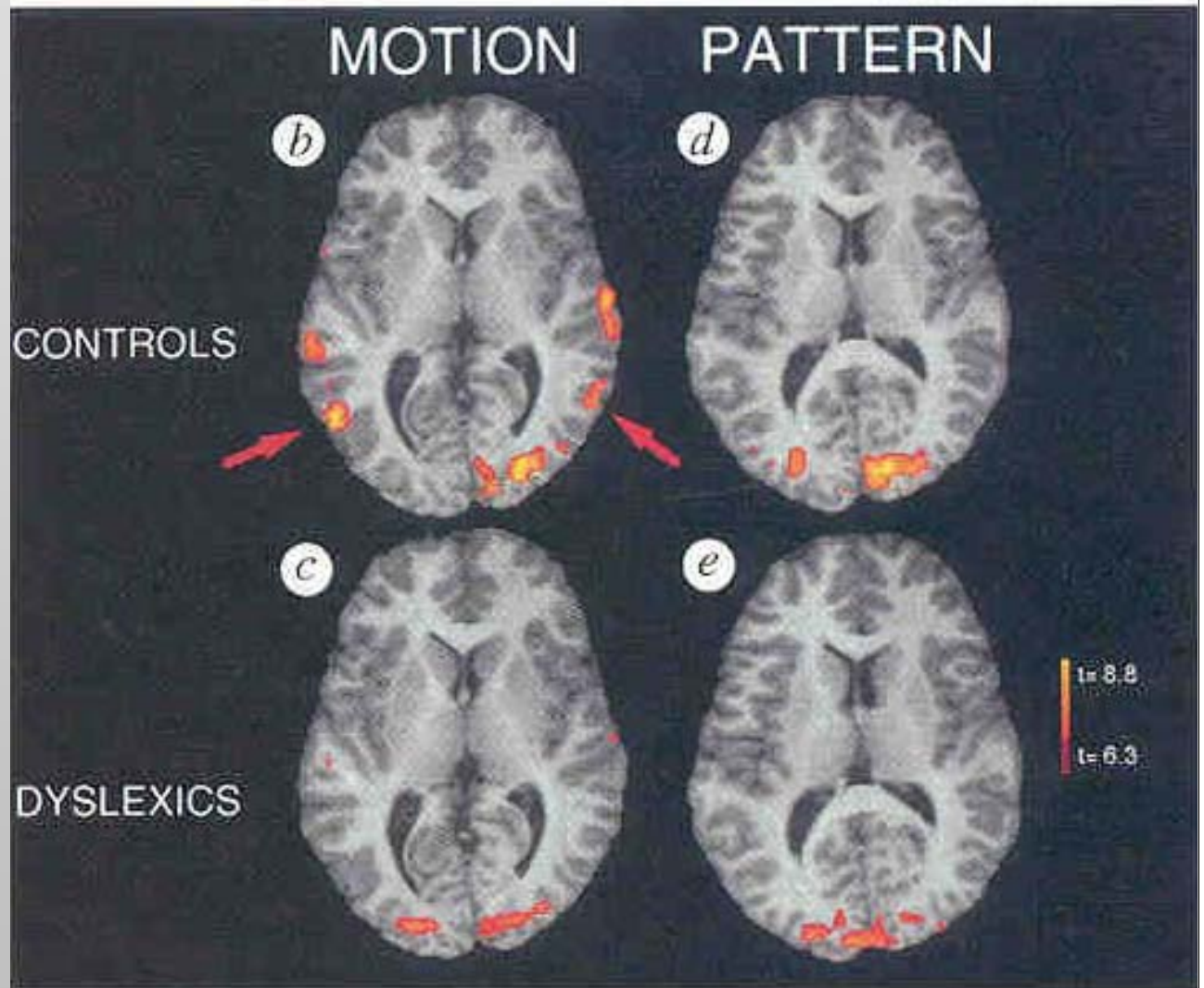
prosopagnosia

# Motion responses in childrens' MT+



MT+  
activation  
reduced in  
poor  
readers  
(Eden et  
al.)

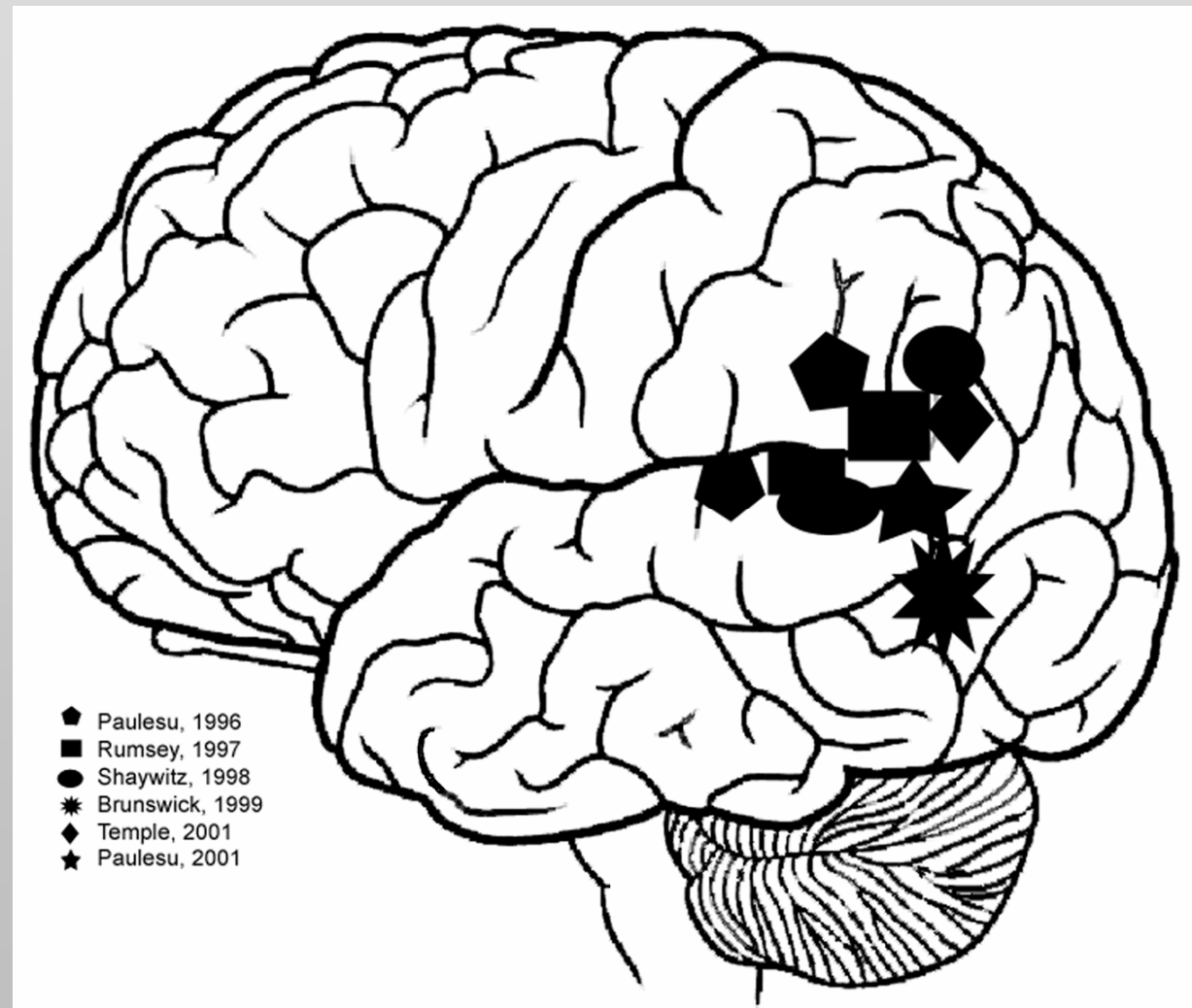
Replicated by  
Demb et al.,  
Ben-Shachar et  
al.



# Response differences during phonological tasks

Temple, 2001, CONB

*Locations where dyslexics had decreased activity relative to normal readers during phonological processing.*

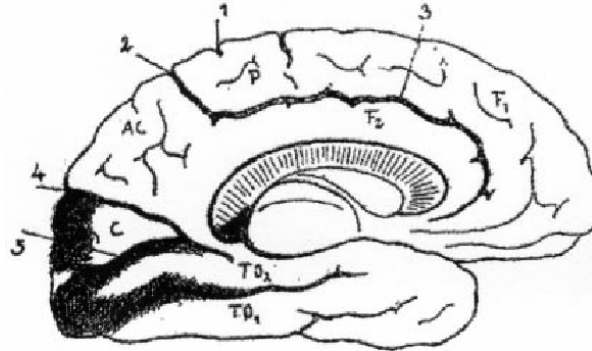


# Neurology of word reading

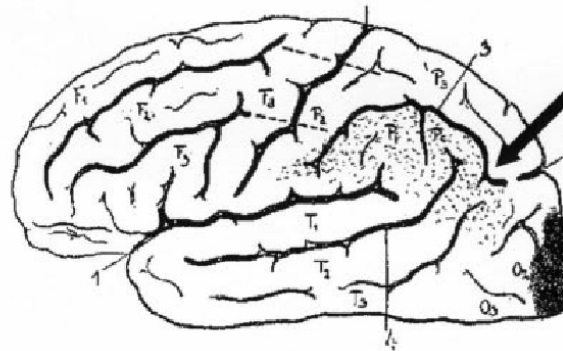
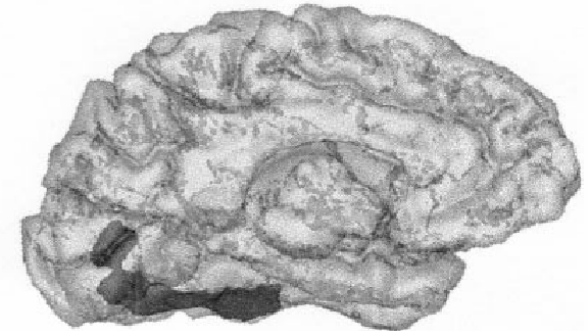
(Cohen et al, 2003; Cerebral Cortex)

• *Dejerine described a patient with a left occipito-temporal lesion who could see, but not process letters and words*

• *Callosal patients have a visual field loss of reading*



Letters vs. Checkerboards



# Remediation results are plagued with several weak results

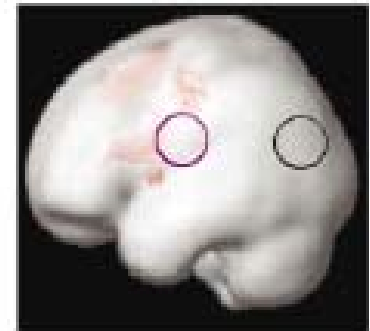
(Temple et al., PNAS, 2003; see  
also Shaywitz papers in PNAS and  
Biological Psychiatry)

Do 'T' and 'D' rhyme?  
vs.  
Are 'P' and 'P' the same?

Normal readers:  
rhyming vs. identity



Poor readers:  
rhyming vs. identity

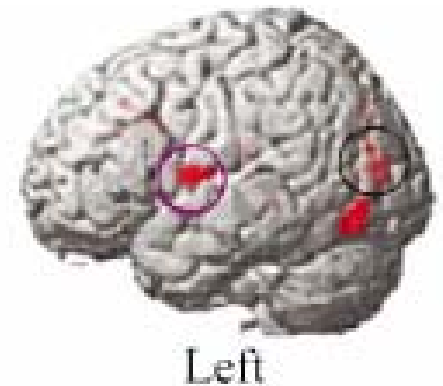
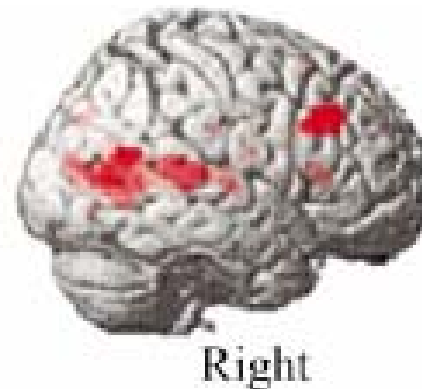


# Remediation results are plagued with several weak results

(Temple et al., PNAS, 2003; see  
also Shaywitz papers in PNAS and  
Biological Psychiatry)

Do 'T' and 'D' rhyme?  
vs.  
Are 'P' and 'P' the same?

Poor readers increases after  
remediation

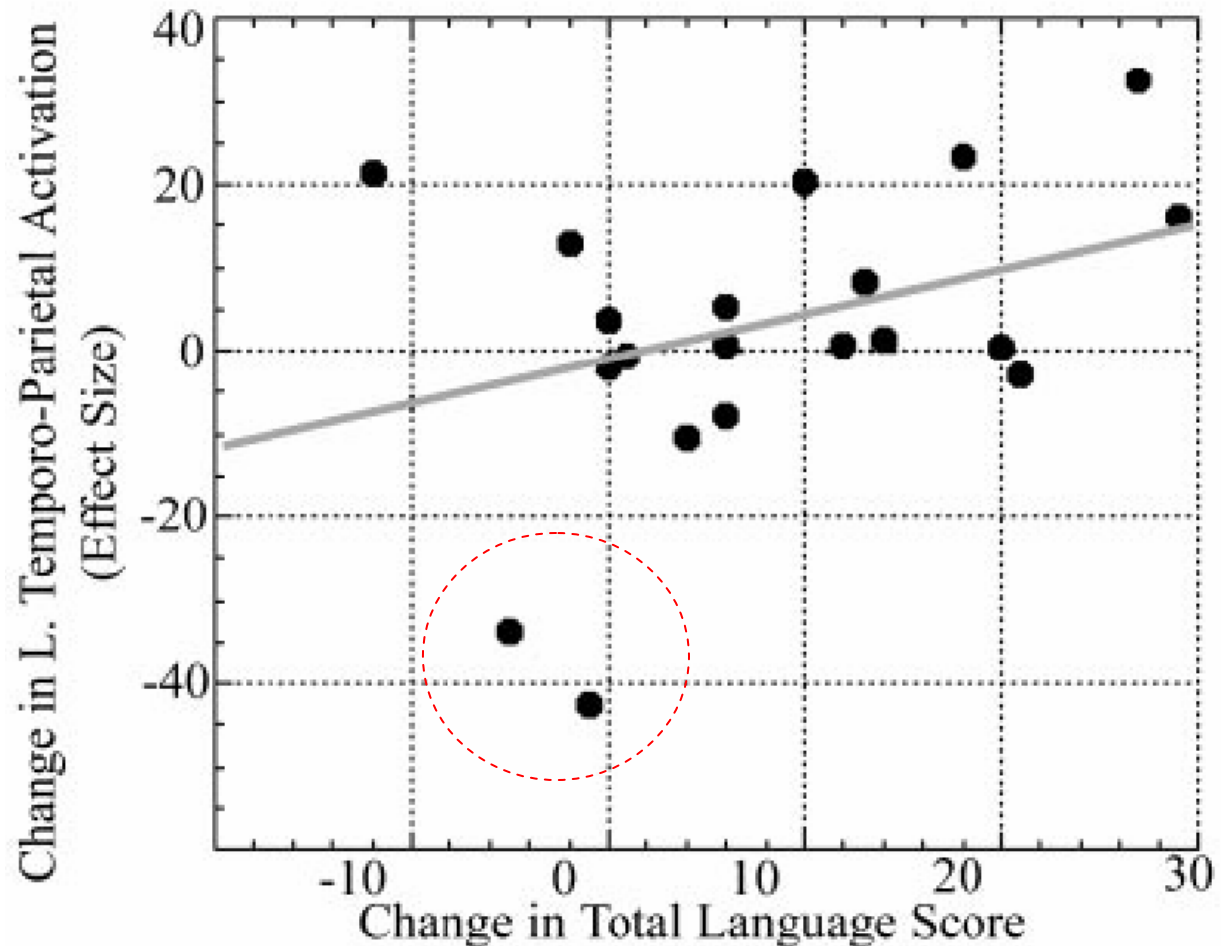


# Remediation results: fMRI signal size and language score

• *Removing the outliers, the correlation coefficient becomes non-significant.*

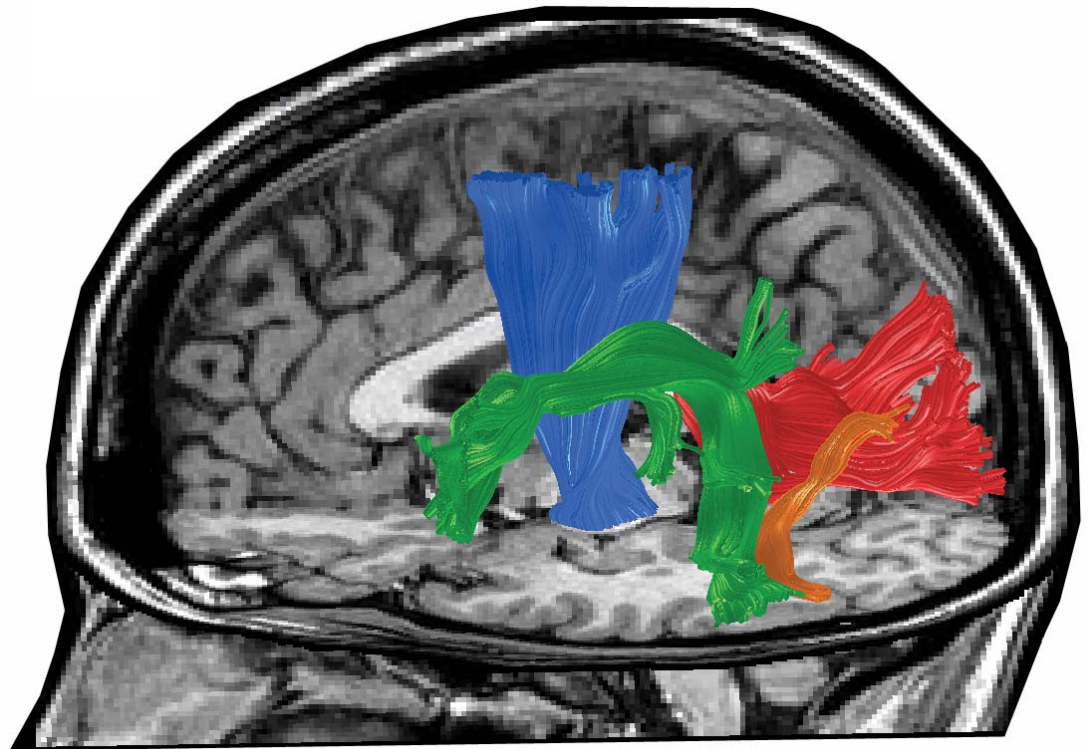
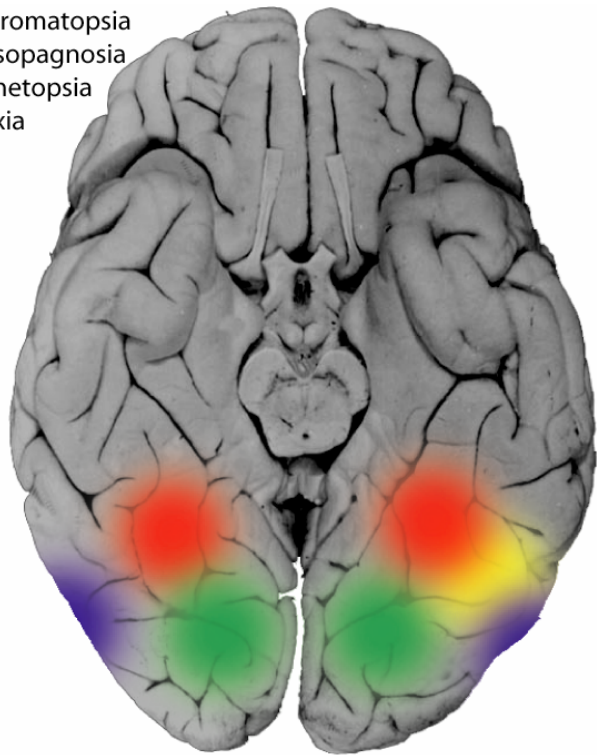
• *Removing two random points, the correlation remains significant*

*Figure 2. Temple et al. (2003) PNAS*

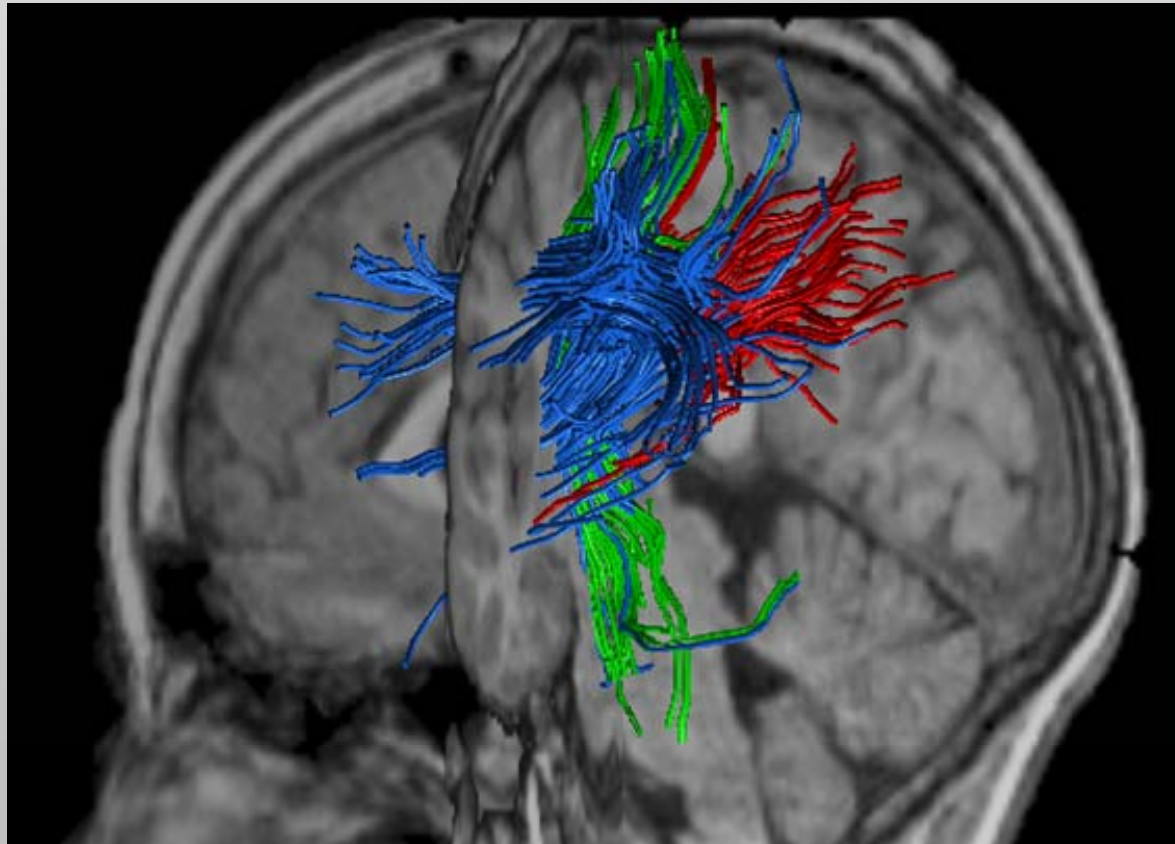


# Tracking the entire network of signals

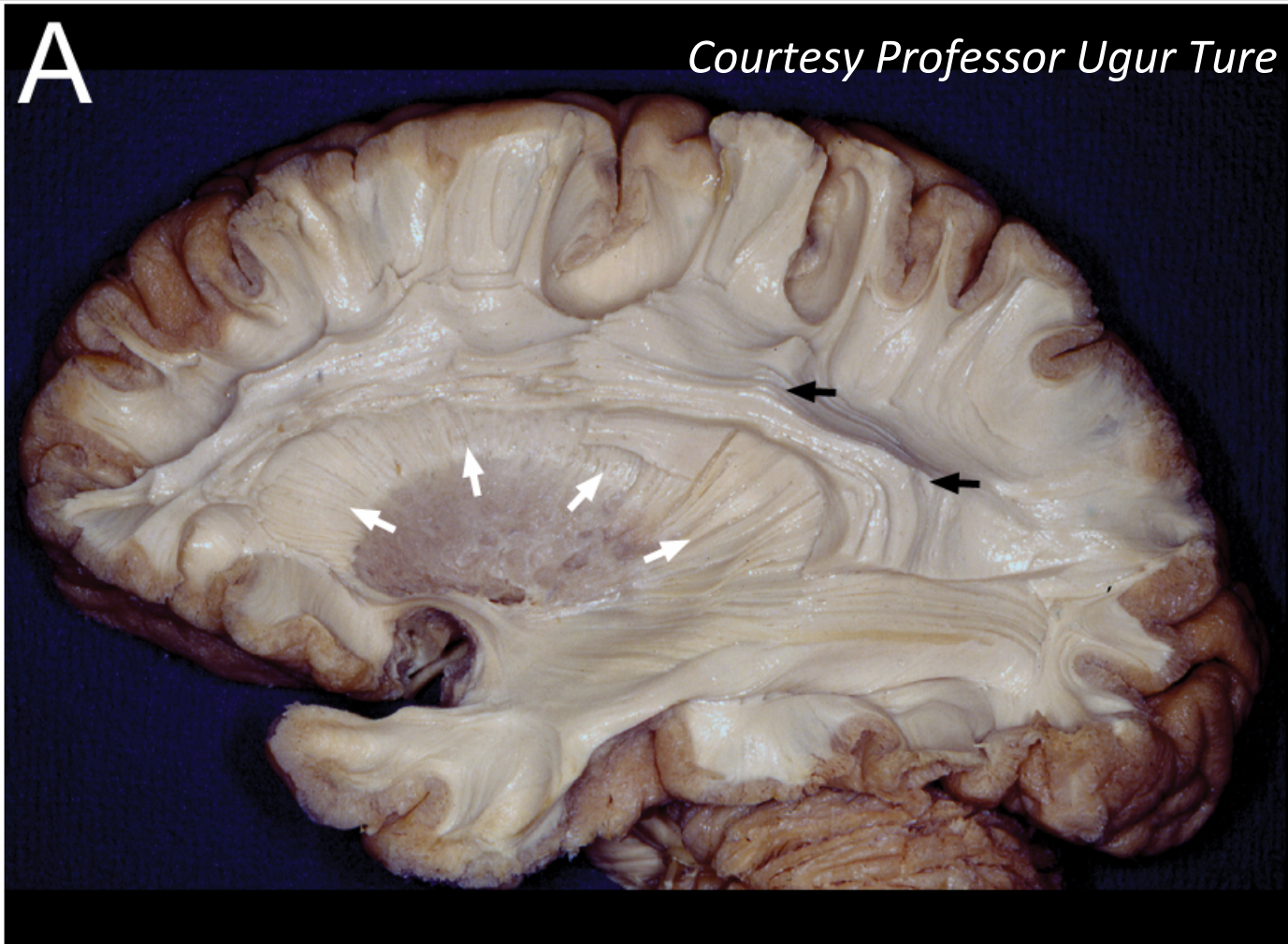
- Achromatopsia
- Prosopagnosia
- Akinetopsia
- Alexia



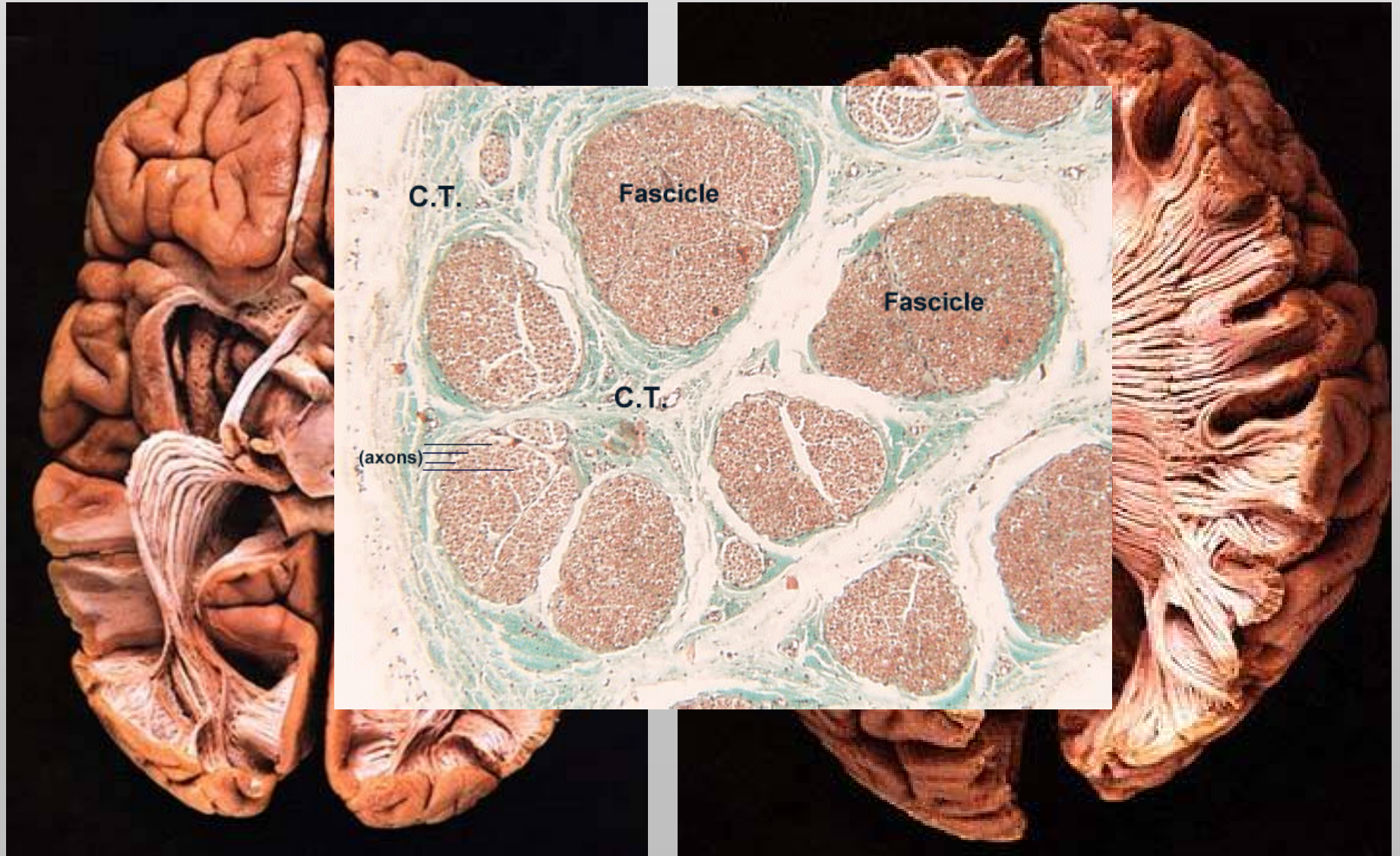
# A New Structural Imaging Method: Diffusion Tensor Imaging



# Human fiber tracts



# Human fiber tracts

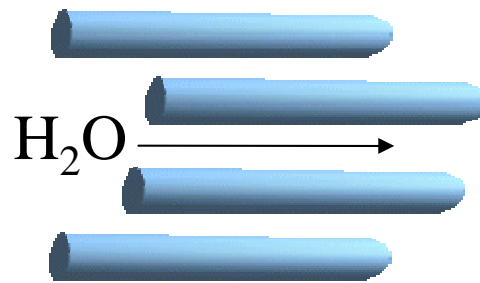


From: The Virtual Hospital ([www.vh.org](http://www.vh.org)); TH Williams, N Gluhbegovic, JY Jew

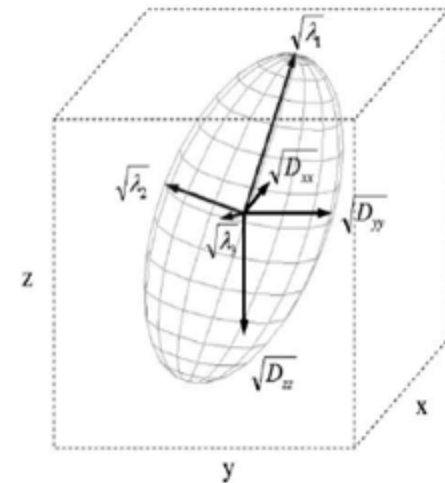
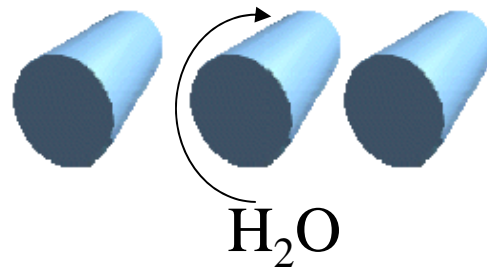
# H<sub>2</sub>O Diffusion Probes Microscopic Structures In the Brain

Unimpeded direction- large ADC

White matter fibres

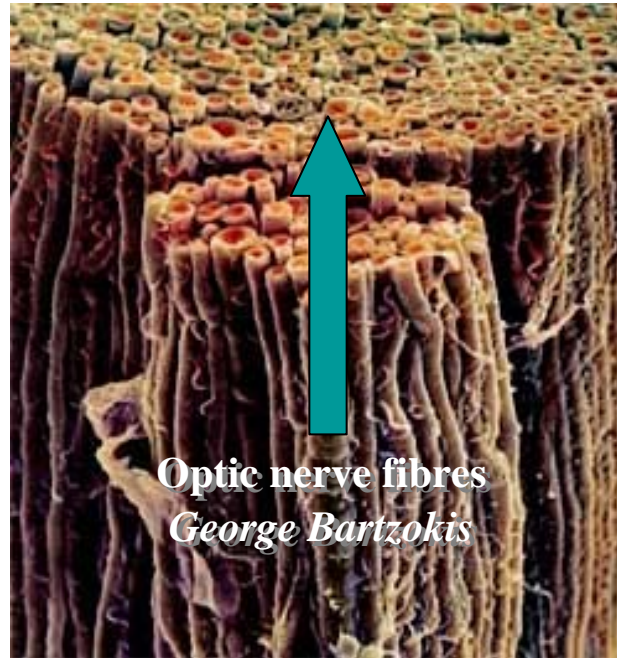


Impeded direction- smaller ADC

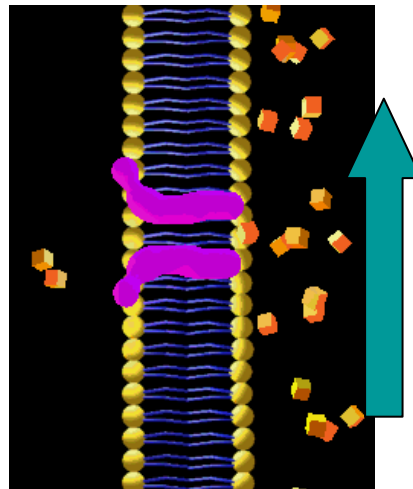


*Principal axis is usually aligned with a fibre bundle; Apparent Diffusion Coefficients (ADCs) are measured in at least 6 directions*

H<sub>2</sub>O  
diffusion  
probes  
membrane  
properties  
in the brain



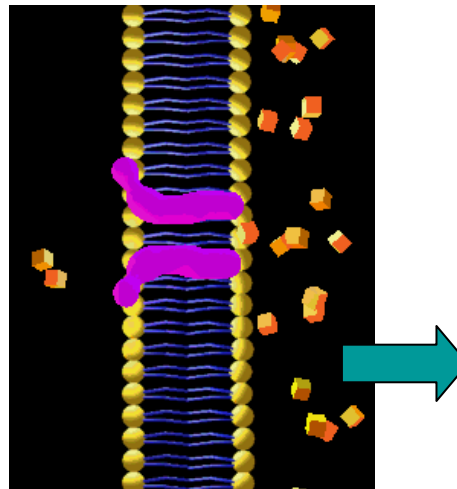
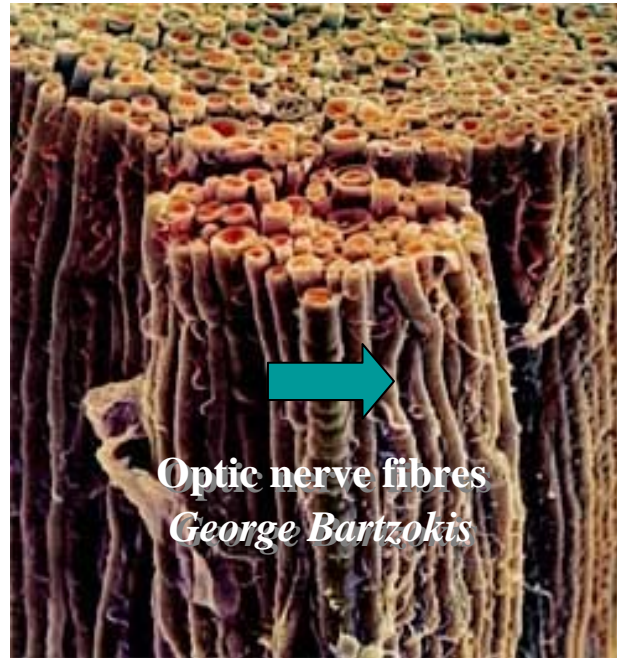
Optic nerve fibres  
*George Bartzokis*



Axial diffusivity

*In regions of high axial coherence, the cytoplasm within the axon limits diffusion and there is a large Apparent Diffusion Coefficient (ADC)*

H<sub>2</sub>O  
diffusion  
probes  
membrane  
properties  
in the brain



Radial diffusivity

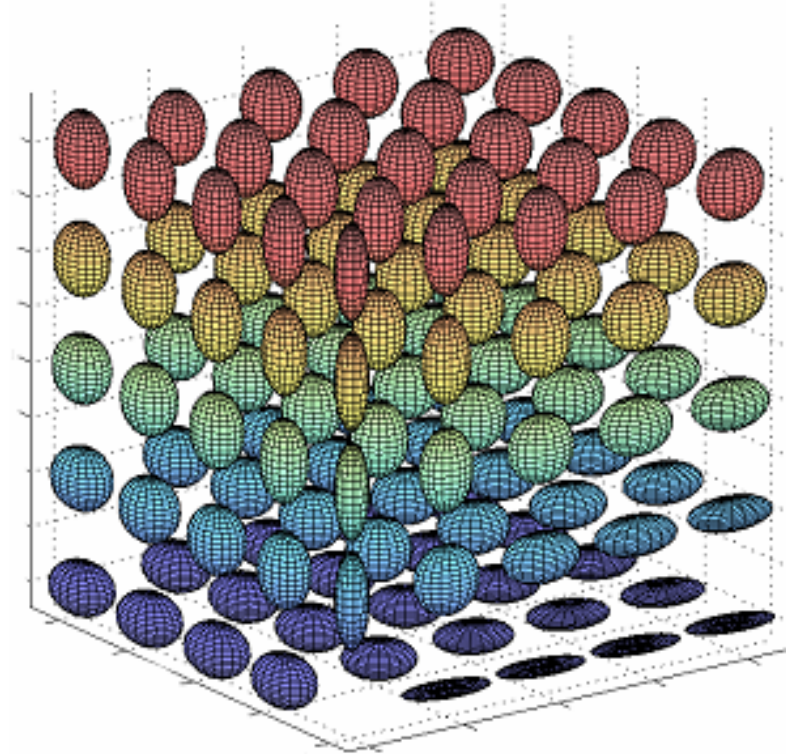
*In all other directions  
the bi-lipid cell  
membranes and myelin  
limit diffusion;  
perpendicular to the  
axon the ADC is  
smaller*

# DTI Data Sets Are Volumes of Diffusion Surfaces

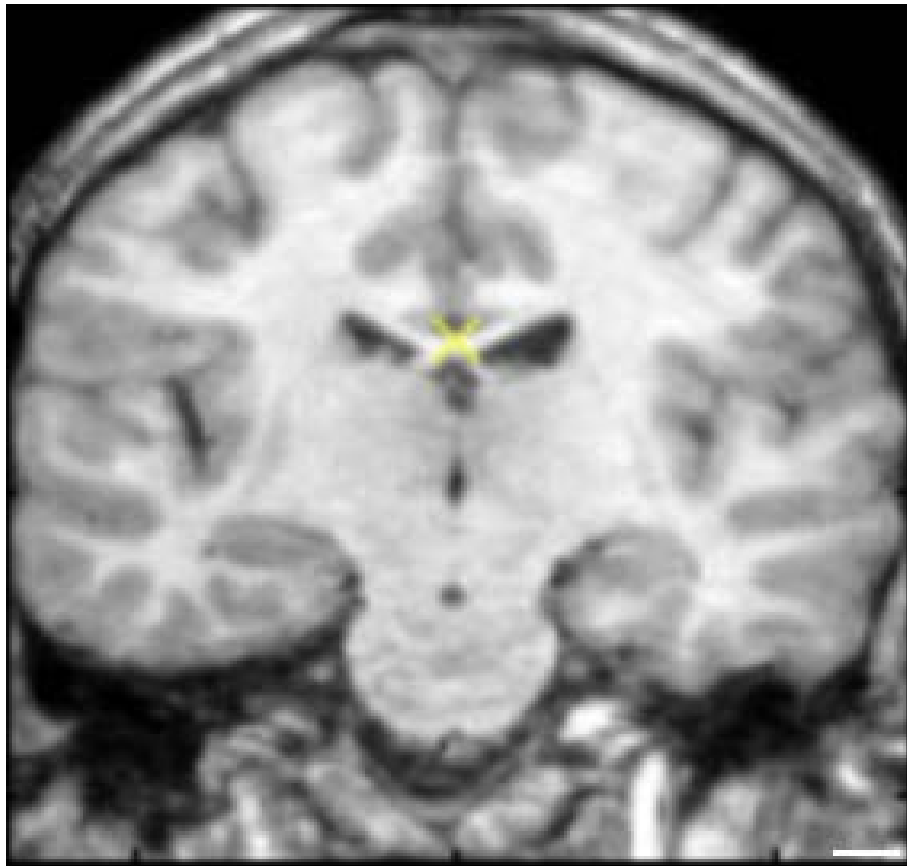
*Conventional MR volumes are real-valued*



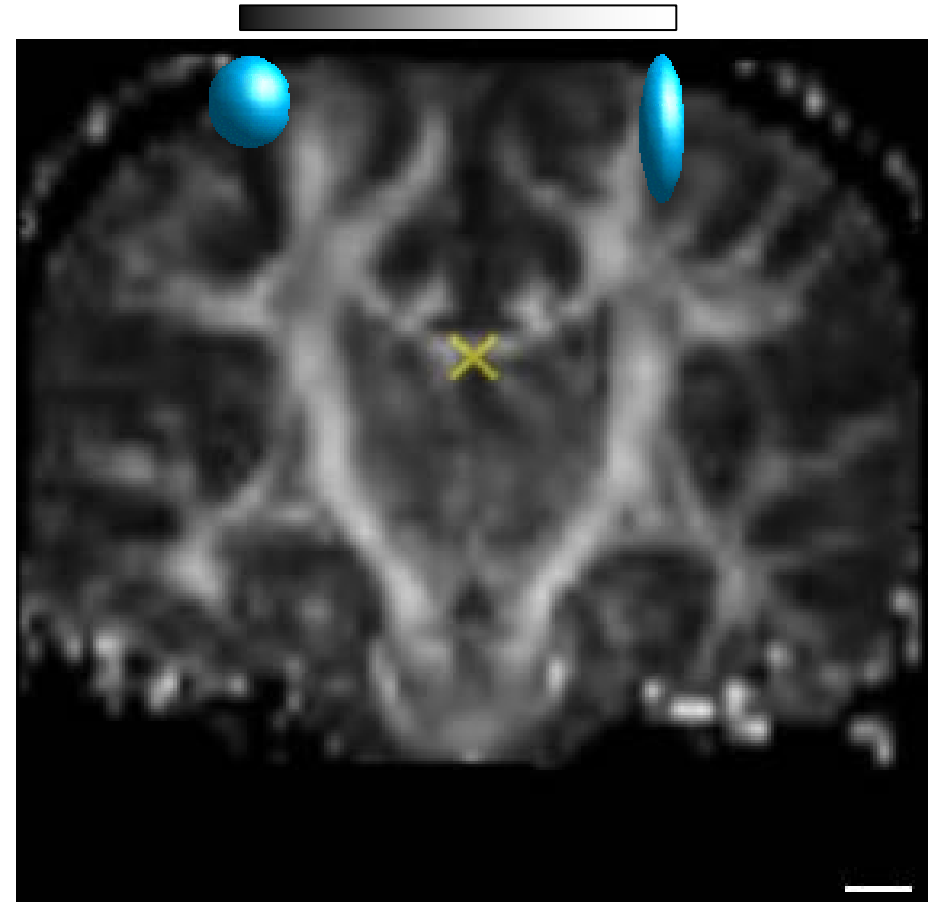
*DTI data are surfaces*



# DTI Reveals Differences In White Matter Structure



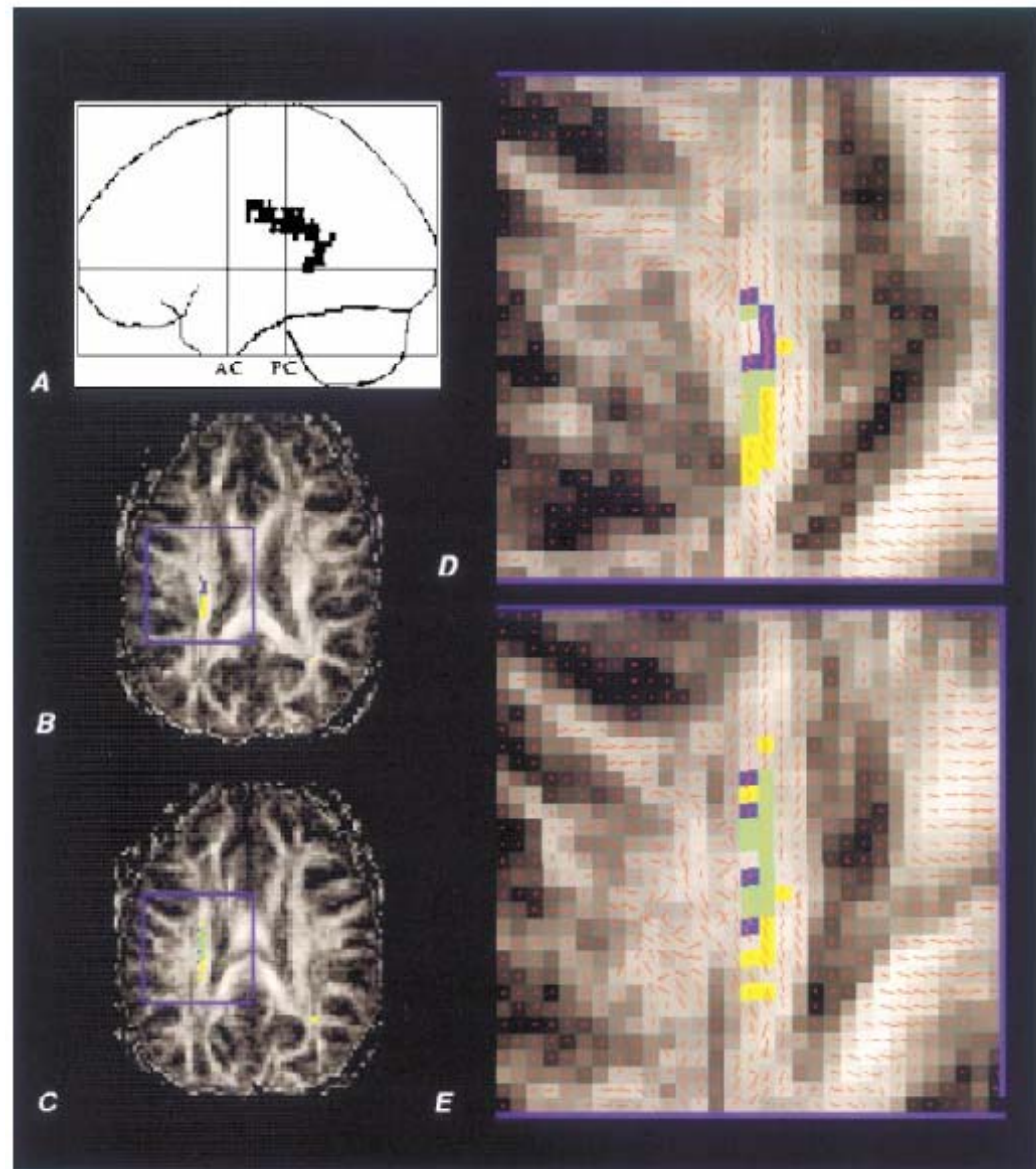
T1



DTI: Fractional Anisotropy

# In adults correlations exist between reading performance and FA

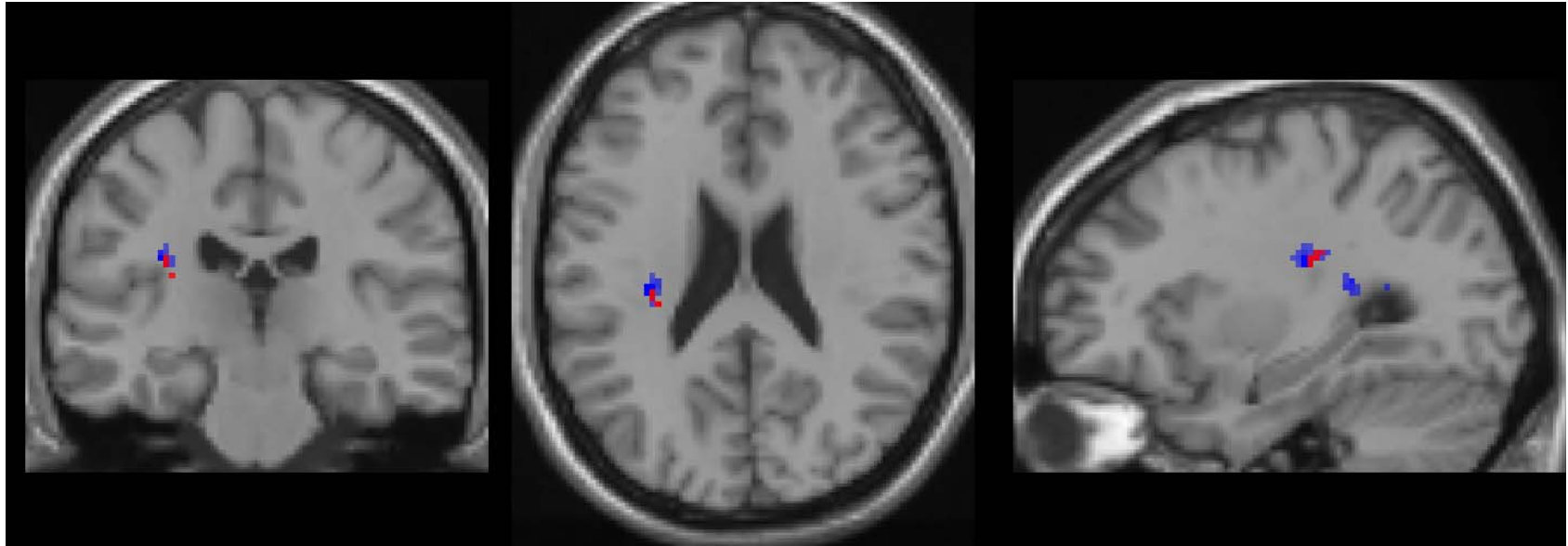
(Klingberg et al., 2000)



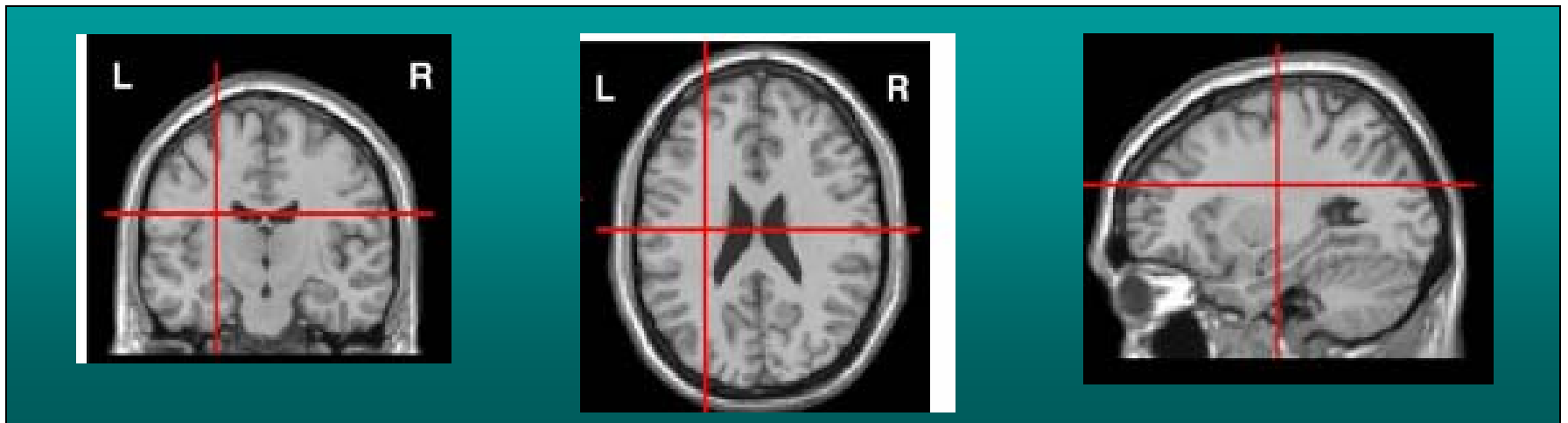
For the gray scale, lighter colors represent higher anisotropy. Green indicates voxels significant in both the between group analysis and the Word ID correlation analysis; yellow indicates voxels significant only in the between-group analysis; and blue indicates voxels significant only in the correlation analysis.

# FA correlates with reading skill in children, too

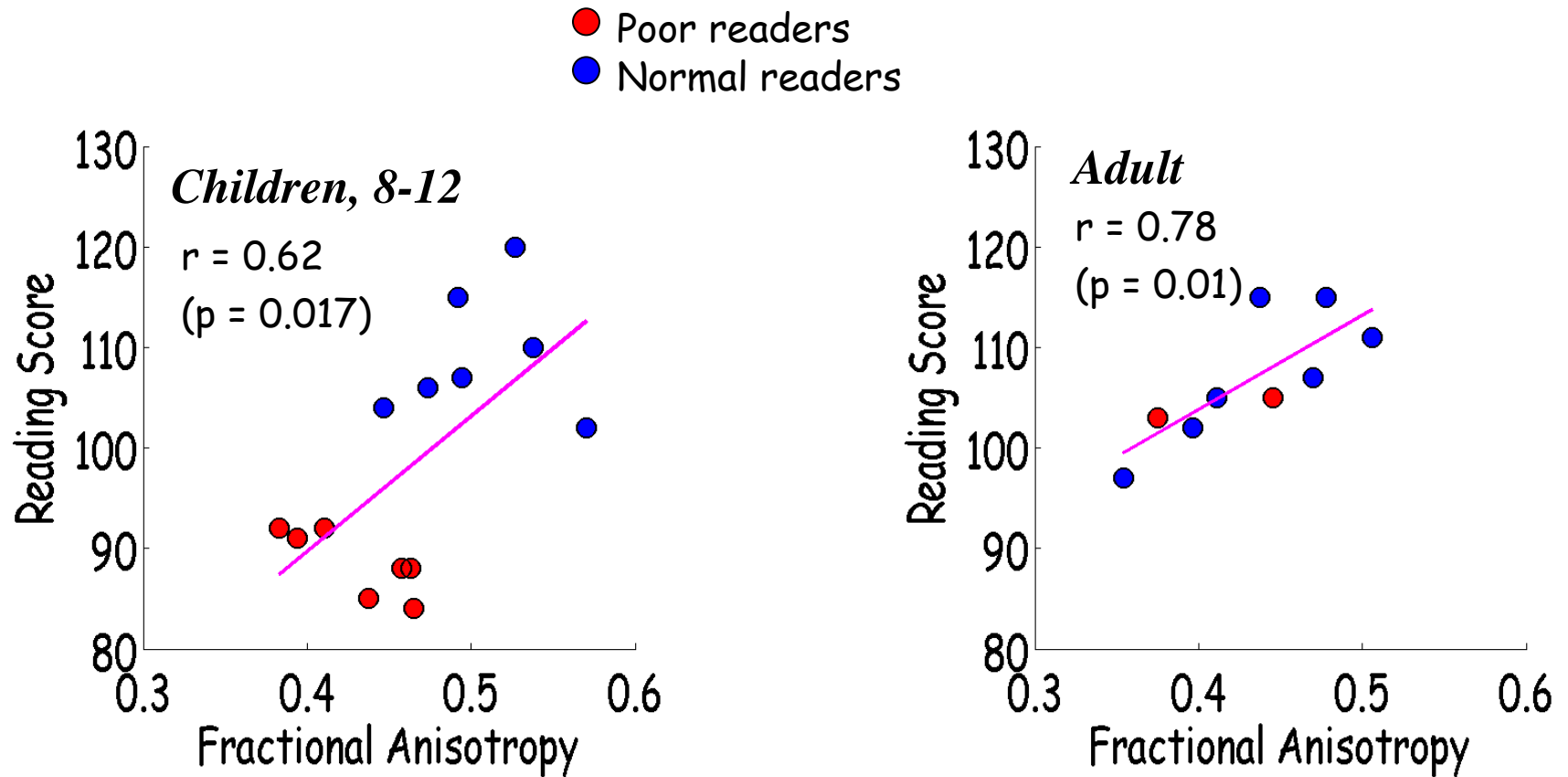
*Deutsch, Dougherty, Bammer, Siok, Gabrieli, Wandell (2005)*



*Beaulieu C, Plewes C, Paulson LA, Roy D, Snook L, Concha L, Phillips L. (2005).*

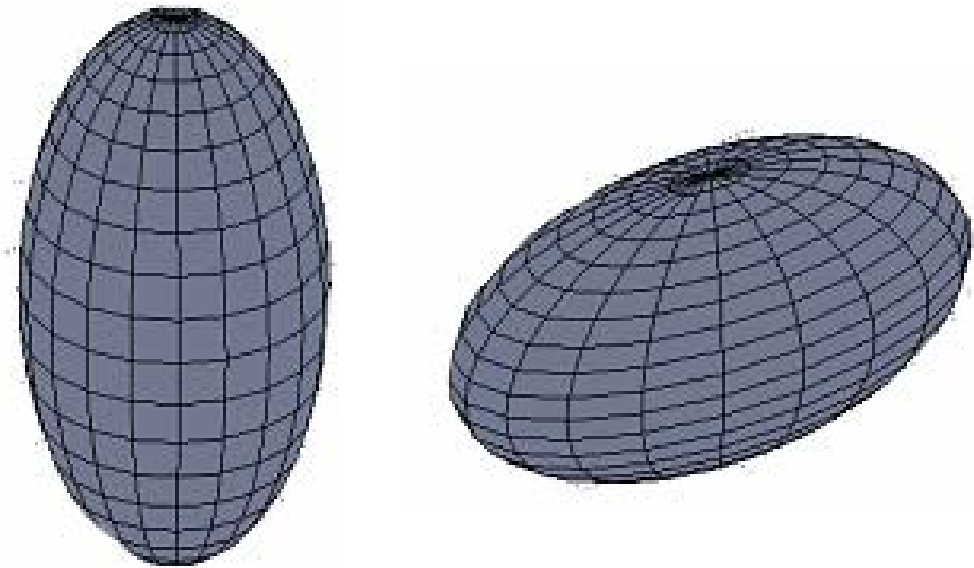


# This correlation is present in children and adults



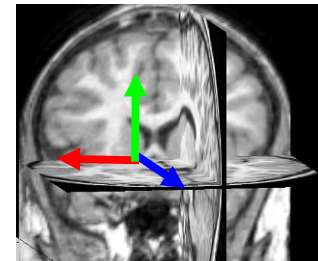
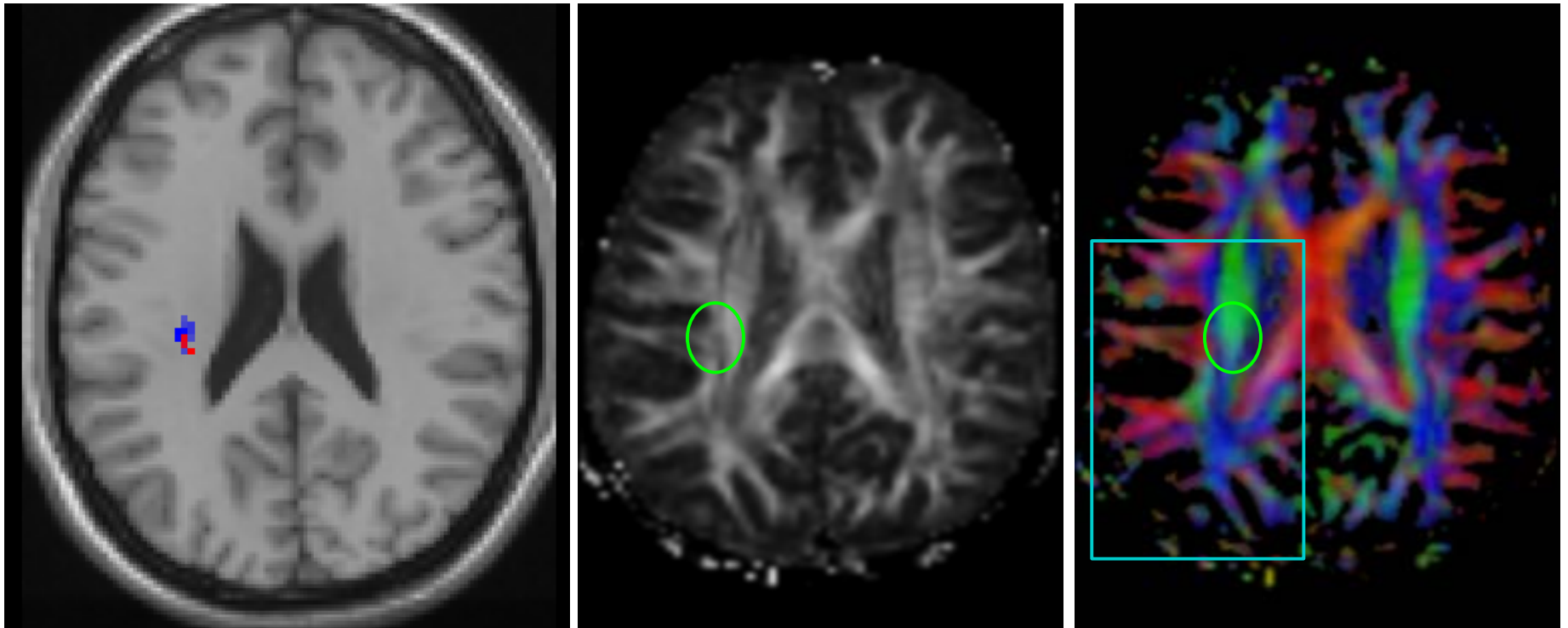
FA does not discriminate between ellipsoid orientations: What more can we learn from the principal diffusion direction?

$FA = 0.6$   
*Principal diffusion directions differ*



# Using direction we see much more

Account for Directional Data Requires New Statistical Methods

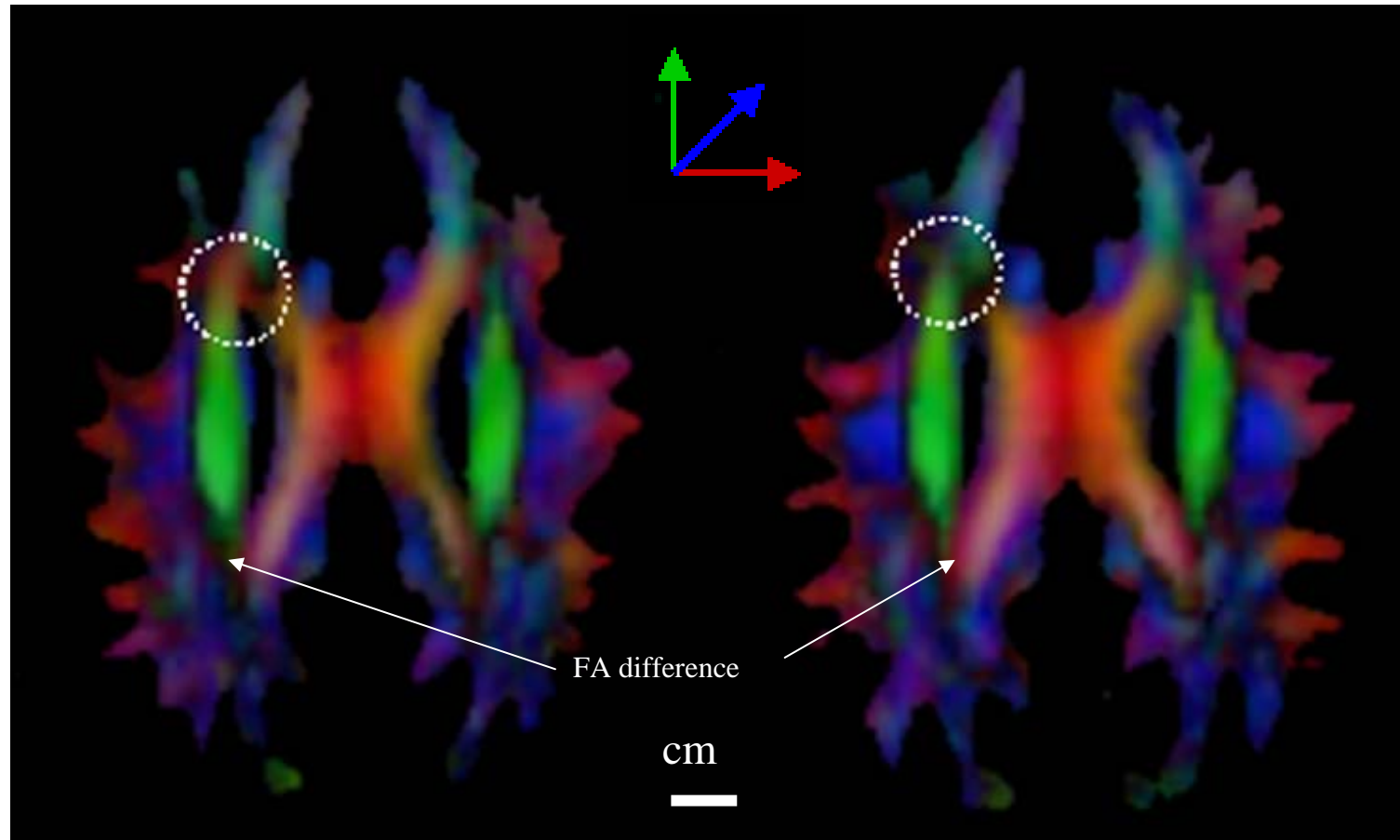


# There is a directional difference in anterior cortex (N=14)

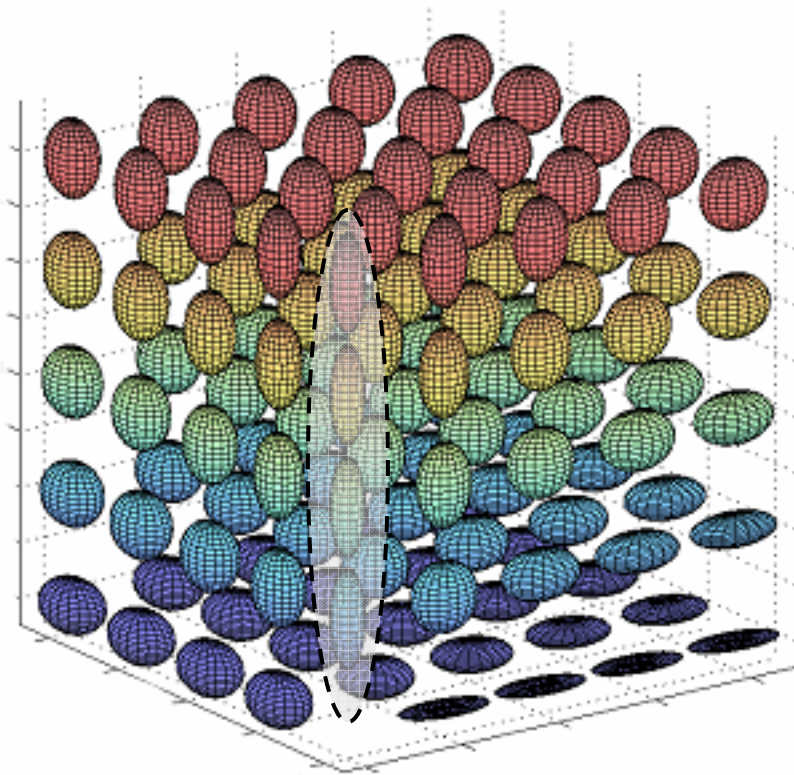
*(Schwartzmann, Dougherty, Taylor, 2005, MRM)*

*Good Readers*

*Poor Readers*



# Can we understand these data by analyzing fiber tracts (DTI-FT)?

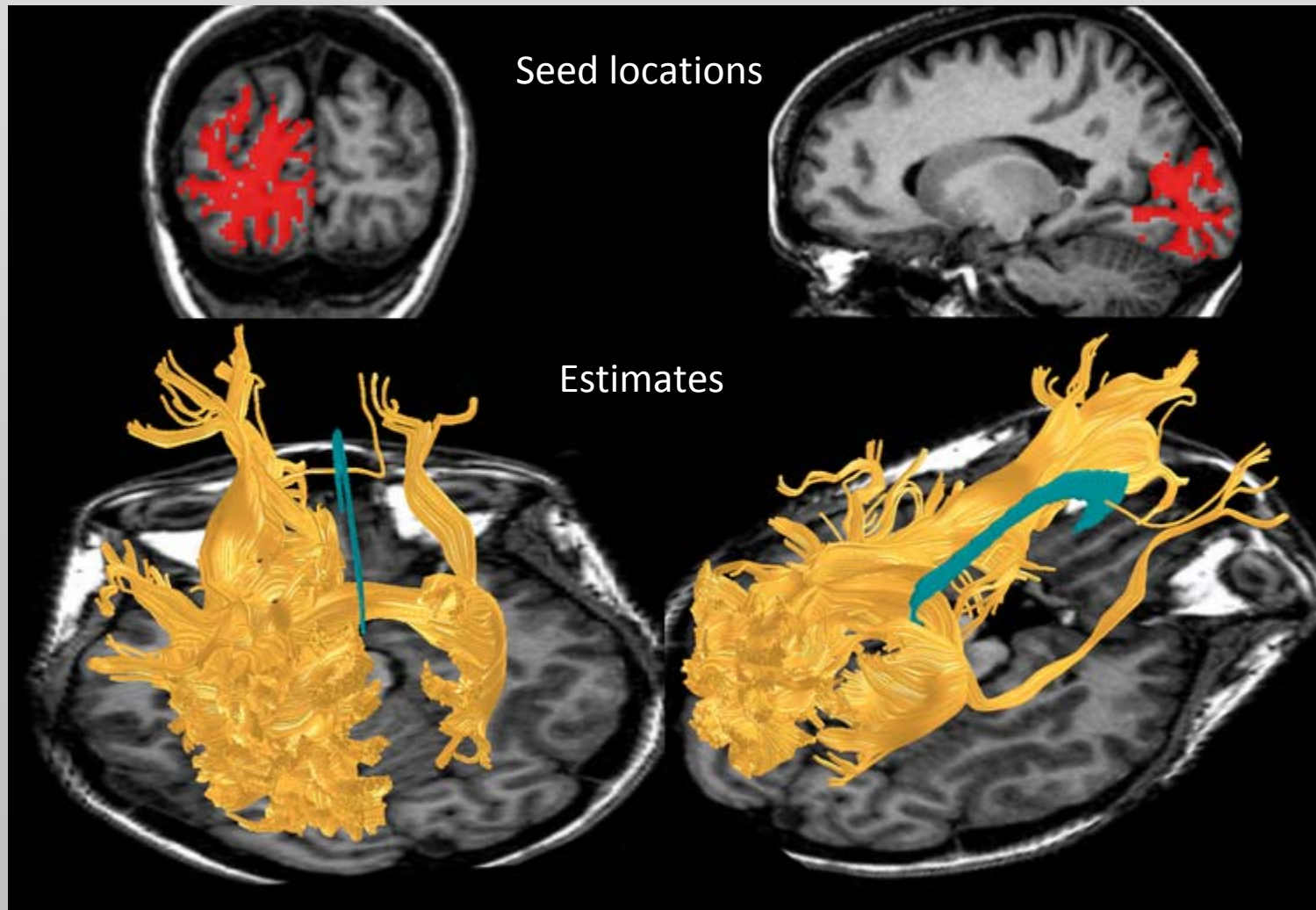


Many interesting algorithm issues

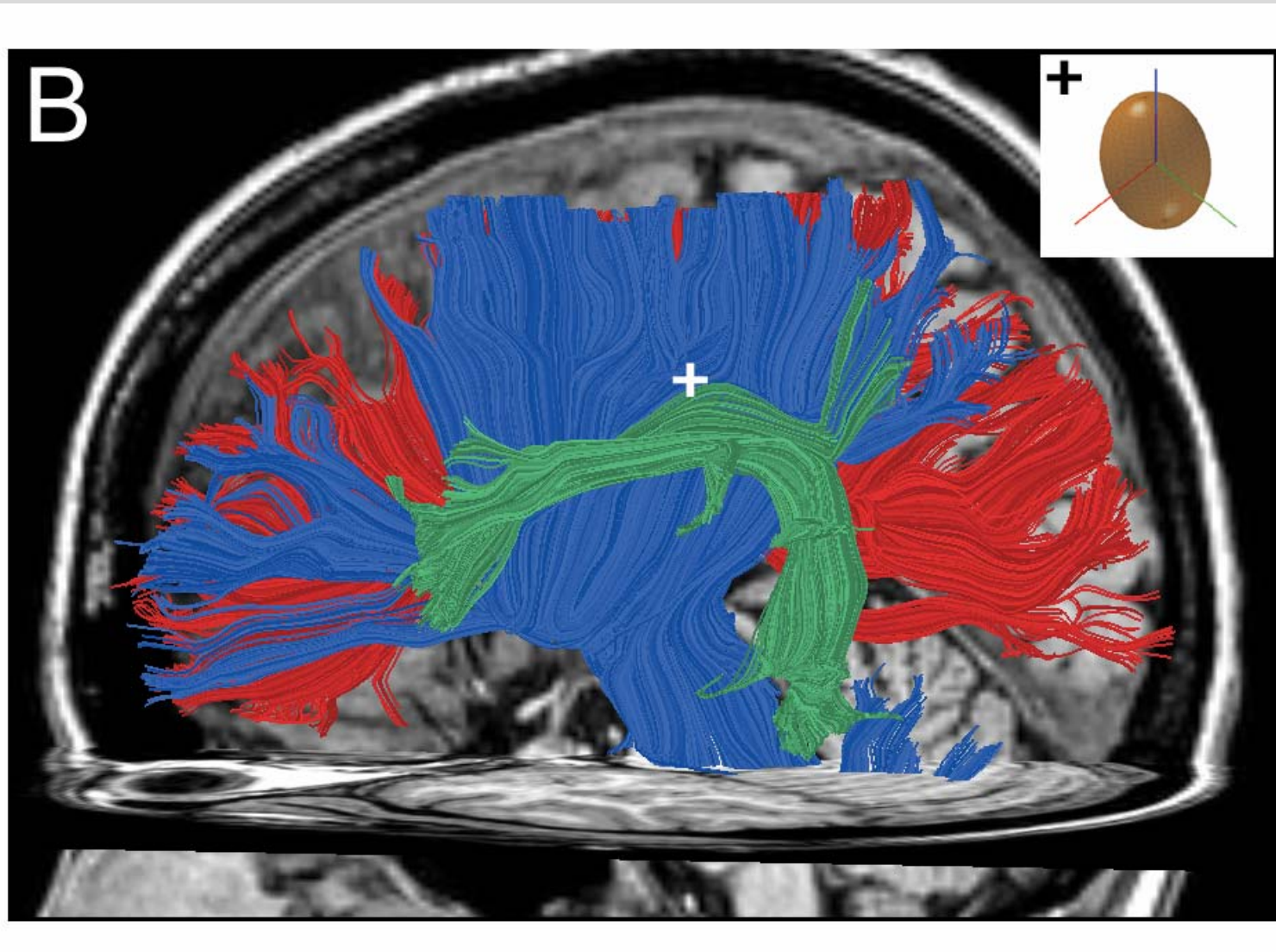
- Algorithm thresholds (direction, FA)
- Confidence intervals, probabilistic reasoning
- Spatial sampling
  - Samples are sparse, but directions are fine
  - Interpolation to intermediate positions
- Spatial co-registration between modalities (T1, fMRI) and subjects
- Validation needed

# Fiber tract estimates

(Dougherty et al., PNAS, 2005)

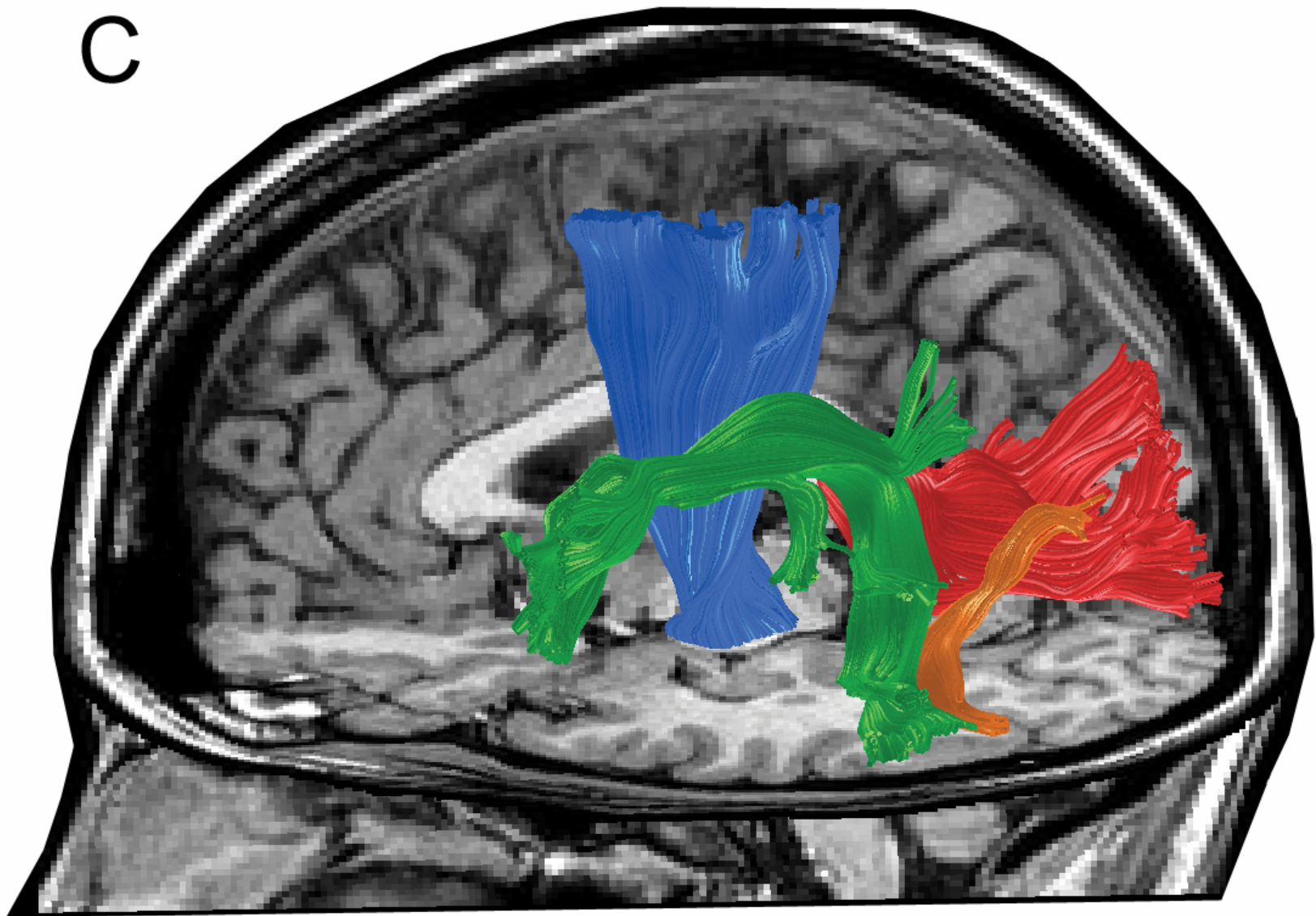


# Principal reading fiber tracts

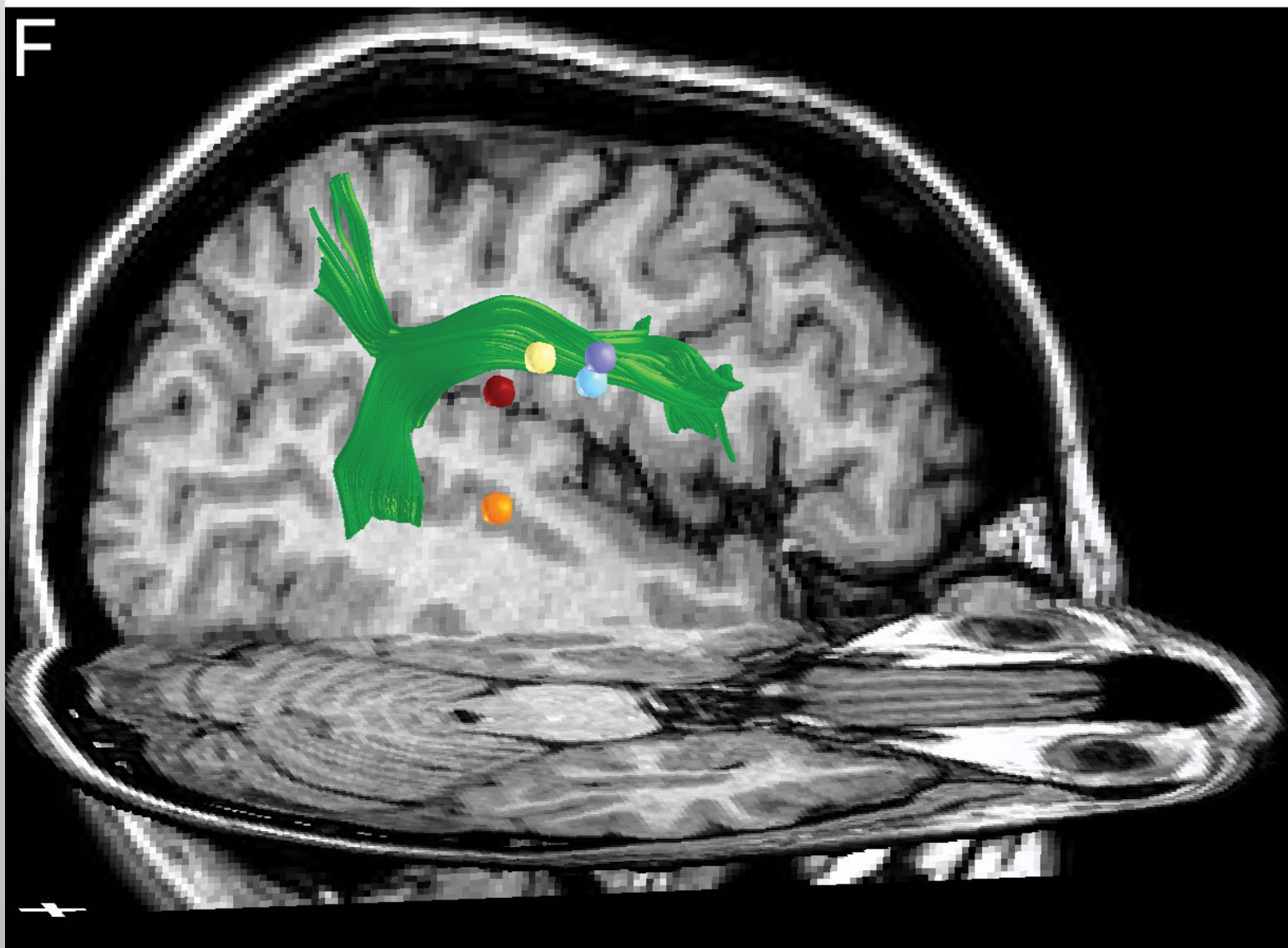


# Principal reading fiber tracts

C



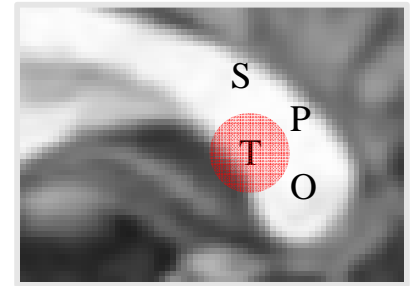
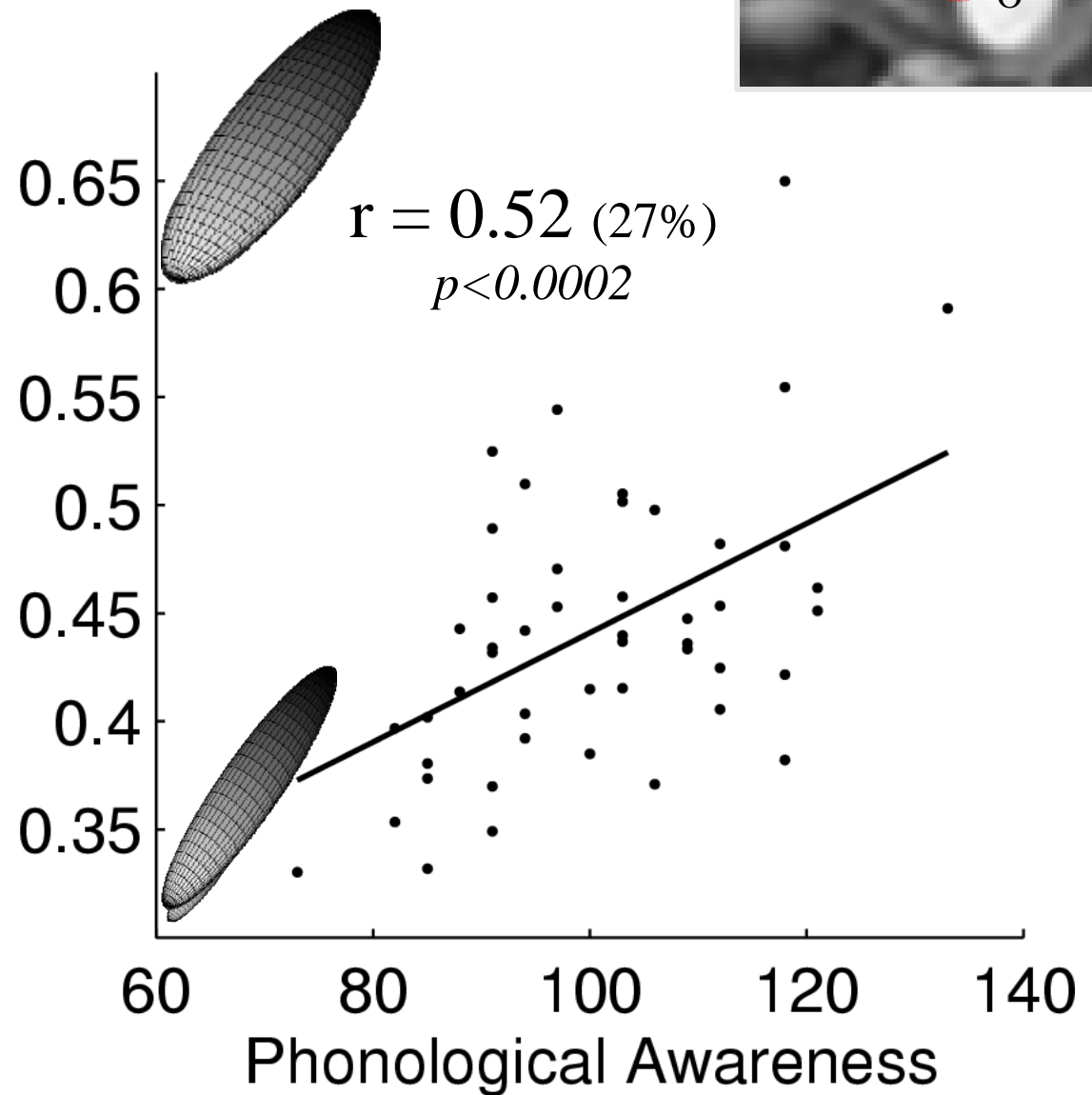
# Principal reading fiber tracts and FA differences



Radial diffusivity  
in temporal  
callosal  
projection zones  
correlates with  
phonological  
awareness

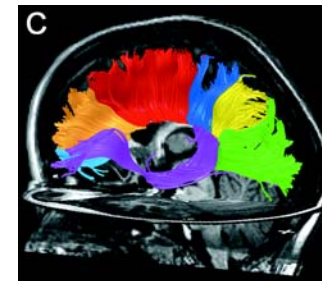
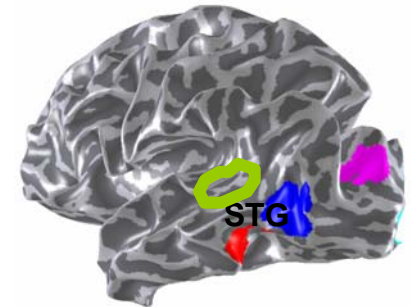
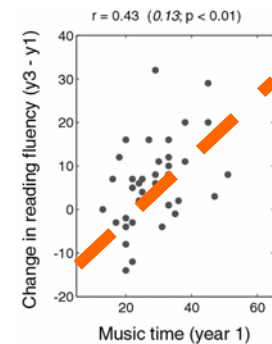
(Dougherty et al., in review)

$(\lambda_2 + \lambda_3)/2$  ( $\mu\text{m}^2/\text{msec}$ )



# What I told you

- We are making progress on identifying key locations in the reading pathway using both functional and structural measures
- We do not have a decisive set of findings about how training influences the development and responses in these locations



# What we would like to know

- Which parts of the reading pathway are enhanced by reading instruction?
  - Does specific training (e.g., phonemic awareness) influence a specific circuit? Is one training technique more or less efficient than the other? Do they combine well?
  - What physiological change result from this training? Synaptic coupling? Sprouting? Myelination and membrane changes?
- How can we improve fMRI and DTI techniques to link neuroimaging data precisely to neural changes
  - Synaptic release and sub-threshold signals, spiking activity, feed forward and backward signals mechanisms (fMRI)
  - Diffusion through myelin, cell membranes, cytoplasm (DTI)

# Collaborators

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