

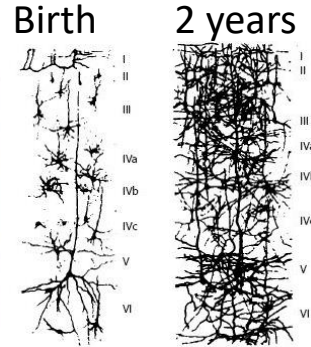
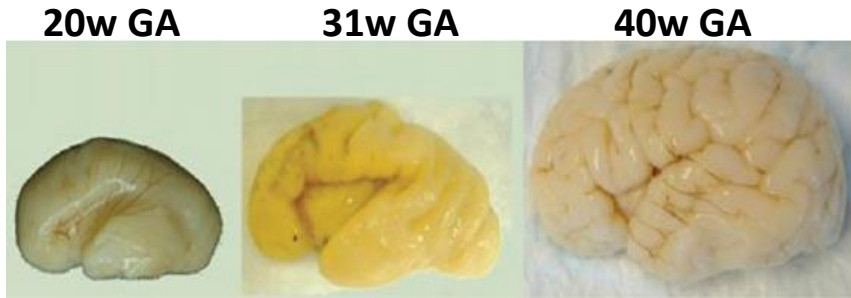
# Bridging the structural and functional correlates of infant brain development using MRI and EEG

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**Jessica Dubois, Ghislaine Dehaene-Lambertz**

**NeuroDiderot – INSERM U1141/Neurospin**

# From brain development to functional acquisitions

## Gray matter maturation



## White matter maturation

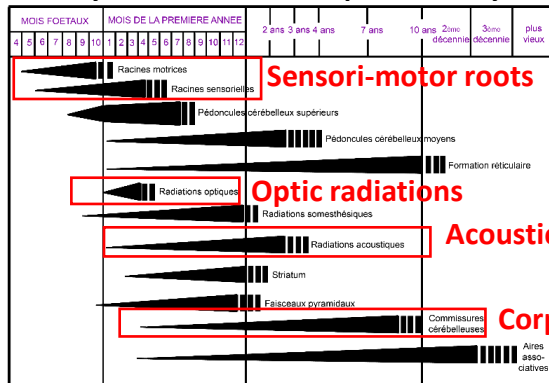


adult

Flechsig, 1920



## Myelination of pathways



Asynchronies in maturation

Yakovlev and Lecours, 1967

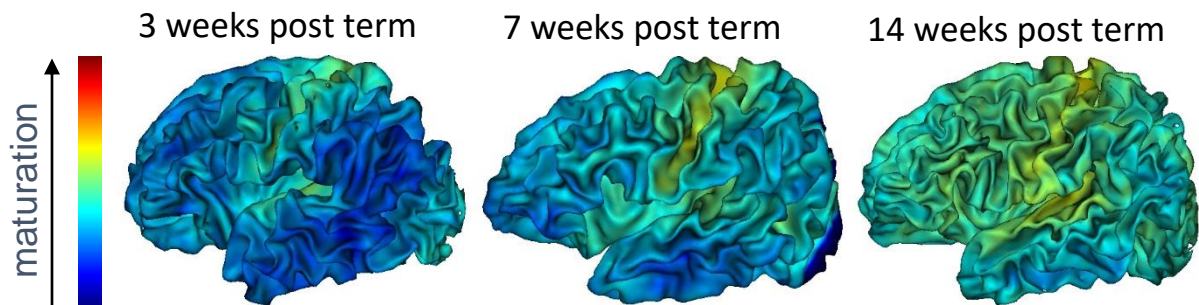
How neuroanatomical changes are translated into behavioral acquisitions ?

➤ Non-invasive neuroimaging of the developing brain *in vivo*

# Brain development during infancy

## Structural maturation – MRI

### Maturation of gray matter

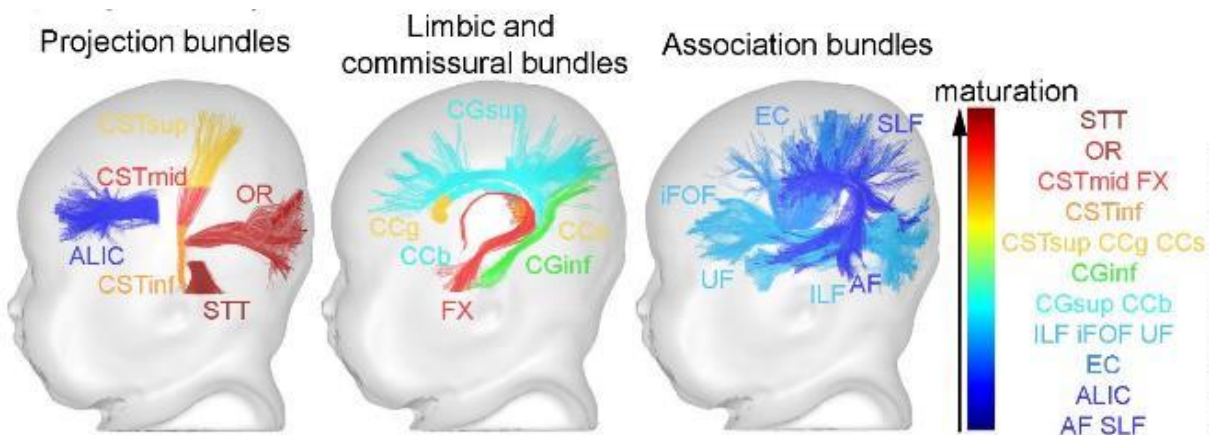


Leroy *et al.*, J. Neuroscience 2011

## Functional maturation – EEG/MEG

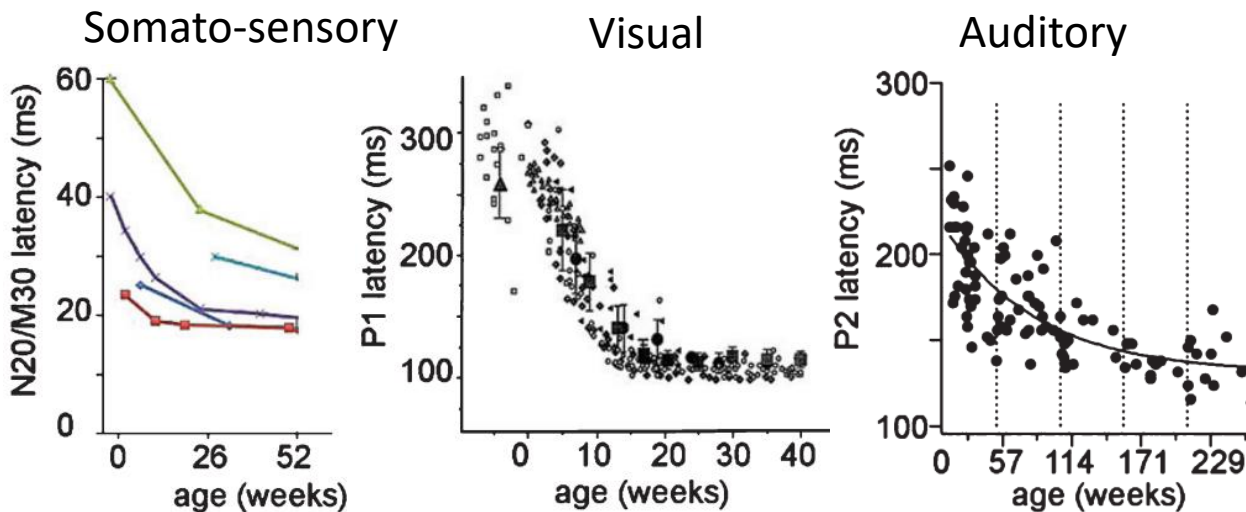


## Maturation of white matter pathways – diffusion MRI



Dubois *et al.*, Brain mapping, 2015

## Response latencies



Nevalainen *et al.*, Front Hum Neuroscience 2014

McCulloch *et al.*, Vision Research 1999

Shafer *et al.*, Int J Psychophysiol. 2015

➡ **Asynchronies in maturation**

# Aim

**Structural development**  
**diffusion MRI**



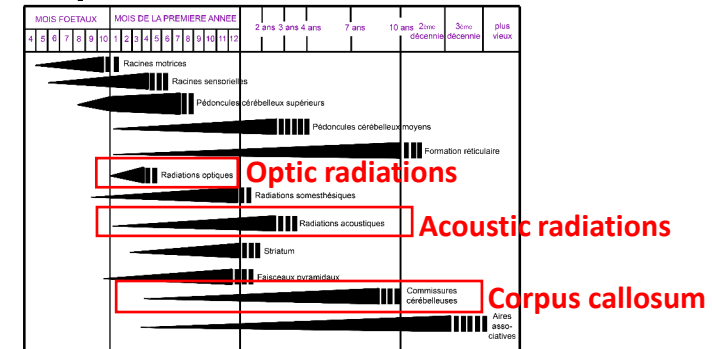
**Functional development**  
**EEG**

- Evaluating the maturation of brain responses with respect to structural maturation of white matter pathways and cortical gray matter :
  - Regressing out the effect of age
  - **In white matter:** Myelination of neuronal fibers increases the speed of neural information that travel through the neurons
  - **In gray matter:** Increase in synaptic density, myelination of intra-cortical fibers or the changes in intracellular organelle density could affect cortical responses.

• Visual and auditory networks

**1. Early responses**

**2. Inter-hemispheric transfer of the responses**



# Functional relevance of structural changes

MRI



Maturation of white matter

diffusion MRI

Myelination ↑



Maturation of brain responses

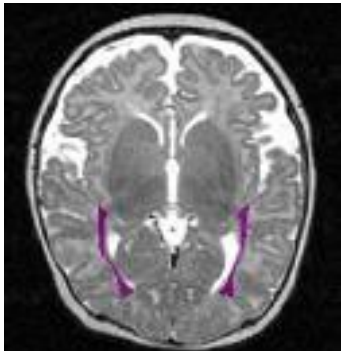
EEG

Neural conduction speed ↑

EEG

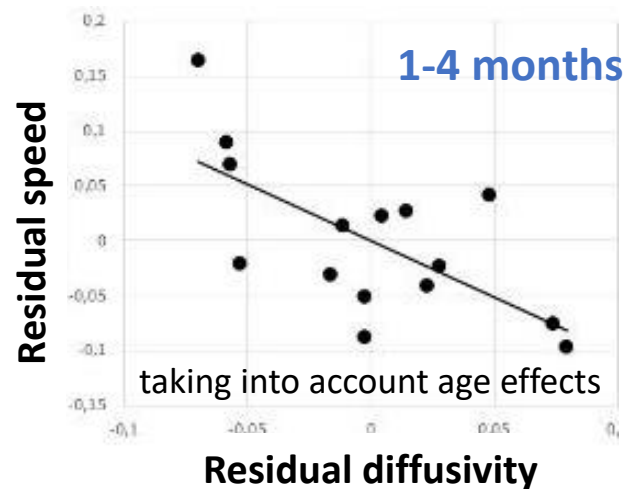


DTI in the optic radiations

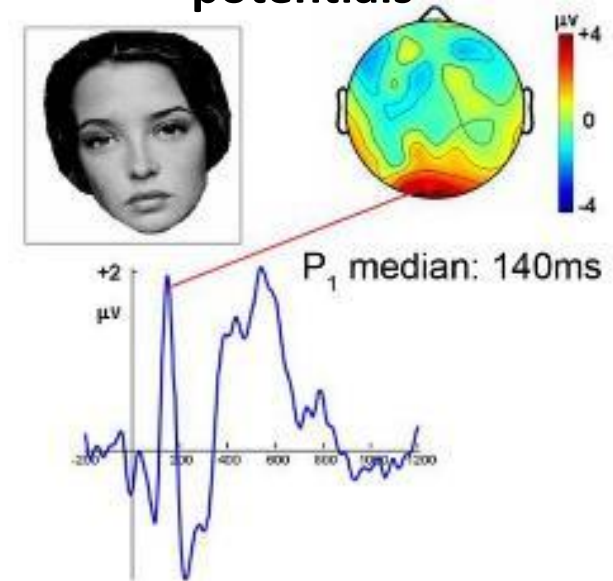


Correlations between functional and structural development

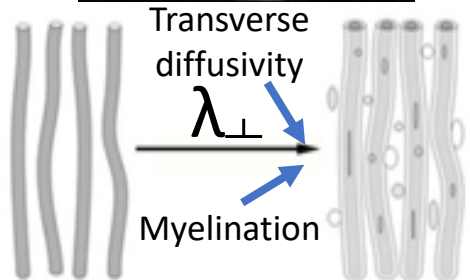
Latency  $\xrightarrow{\text{Anatomical Distances}}$  Conduction Speed



EEG – Visual evoked potentials

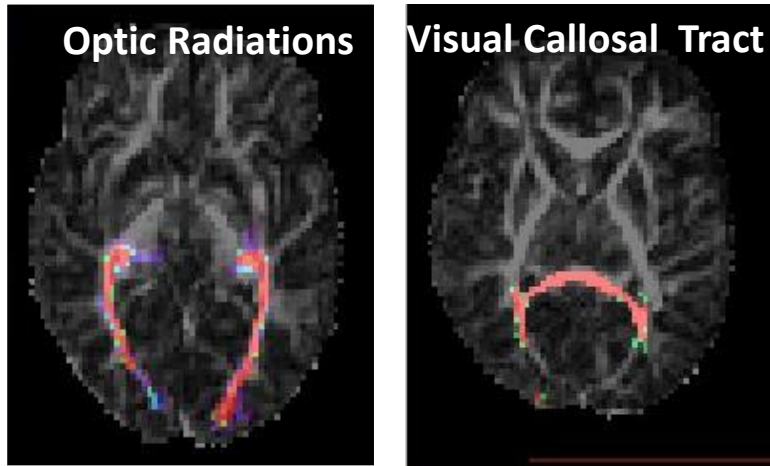


Dubois *et al.*, J Neuroscience, 2008

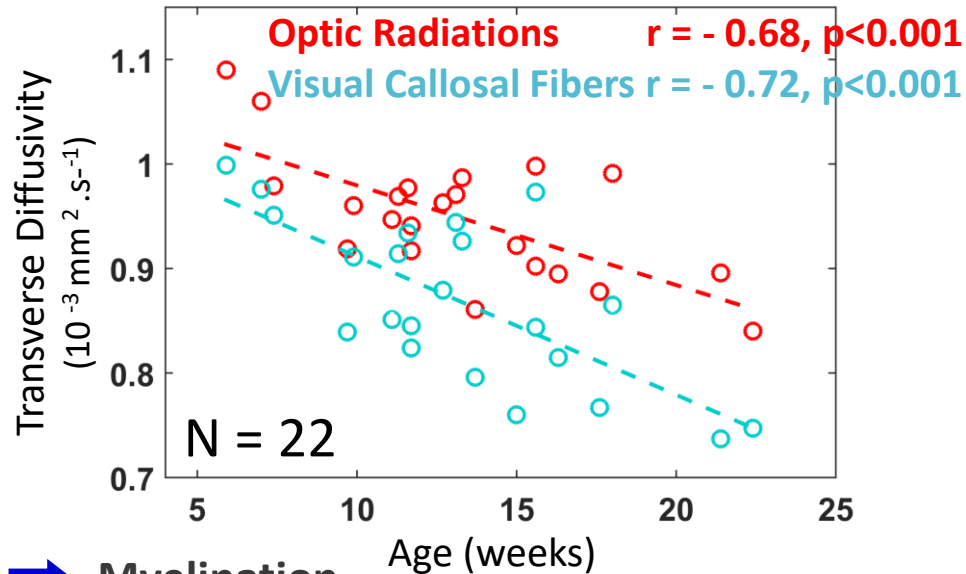


# MRI and EEG correlates of visual development

Pathways reconstruction: Tractography

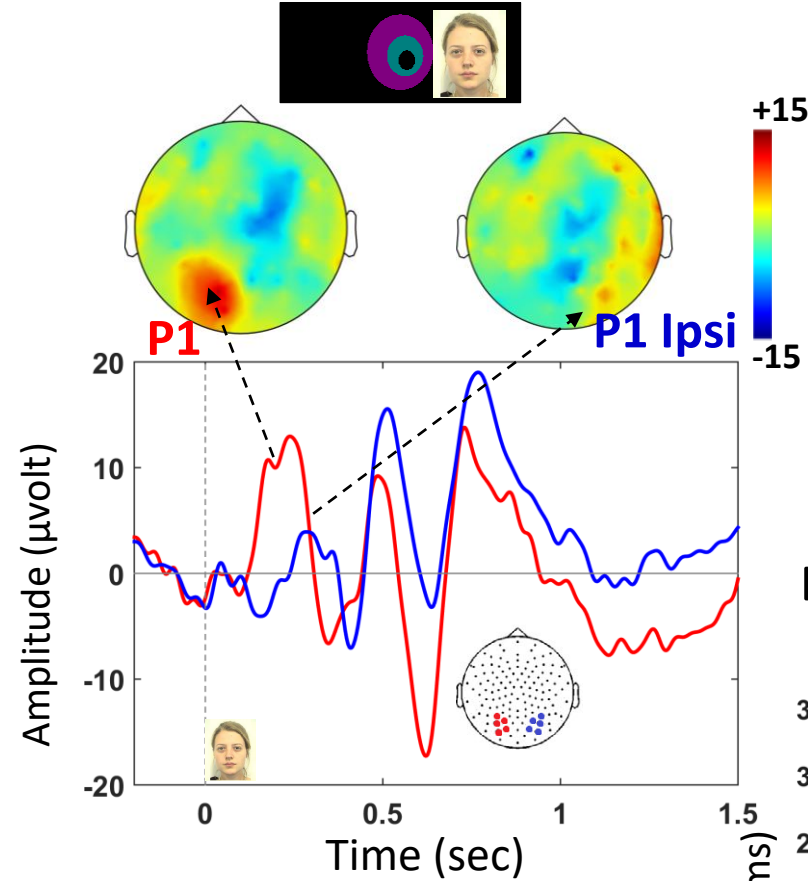


Decrease in transverse diffusivity across development



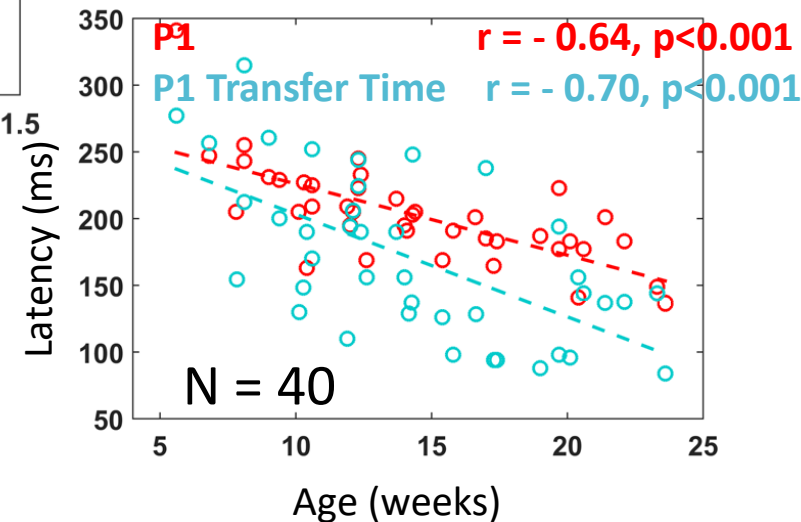
➔ Myelination

Visual evoked responses • 40 infants : 1-6 months old



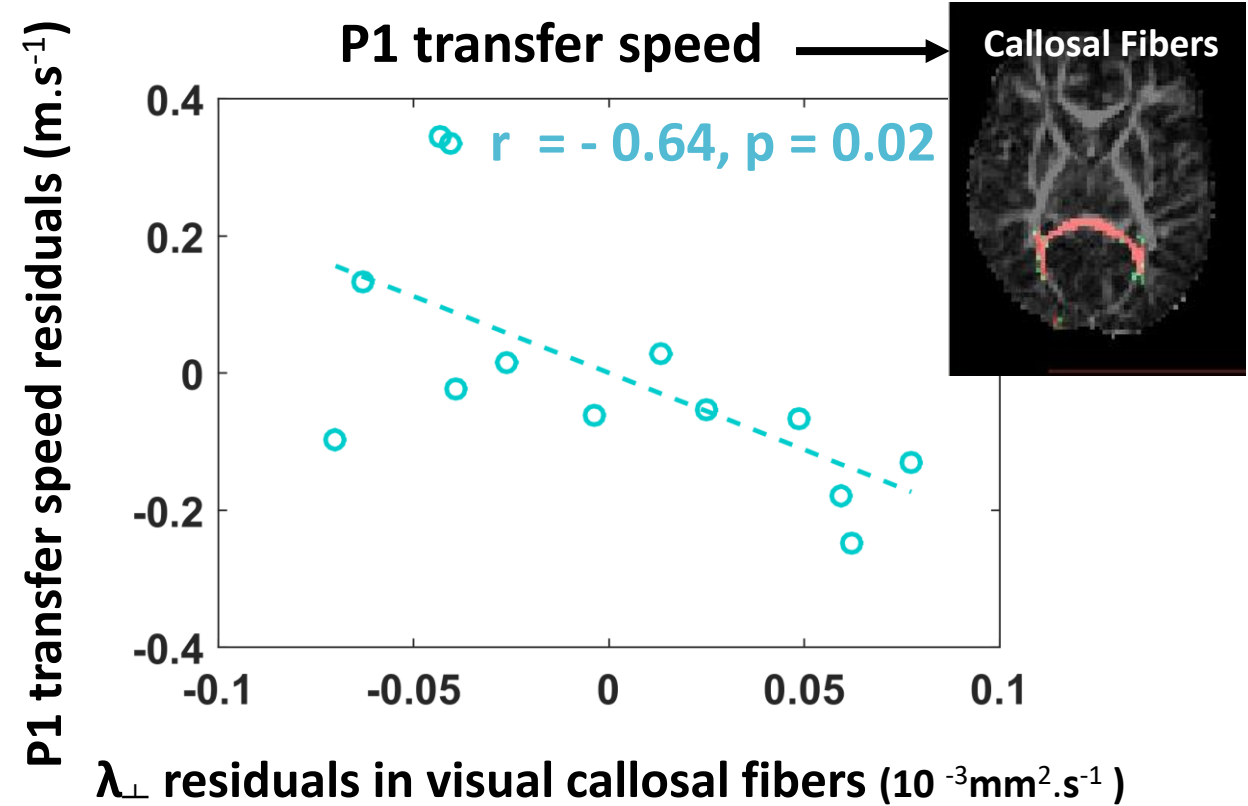
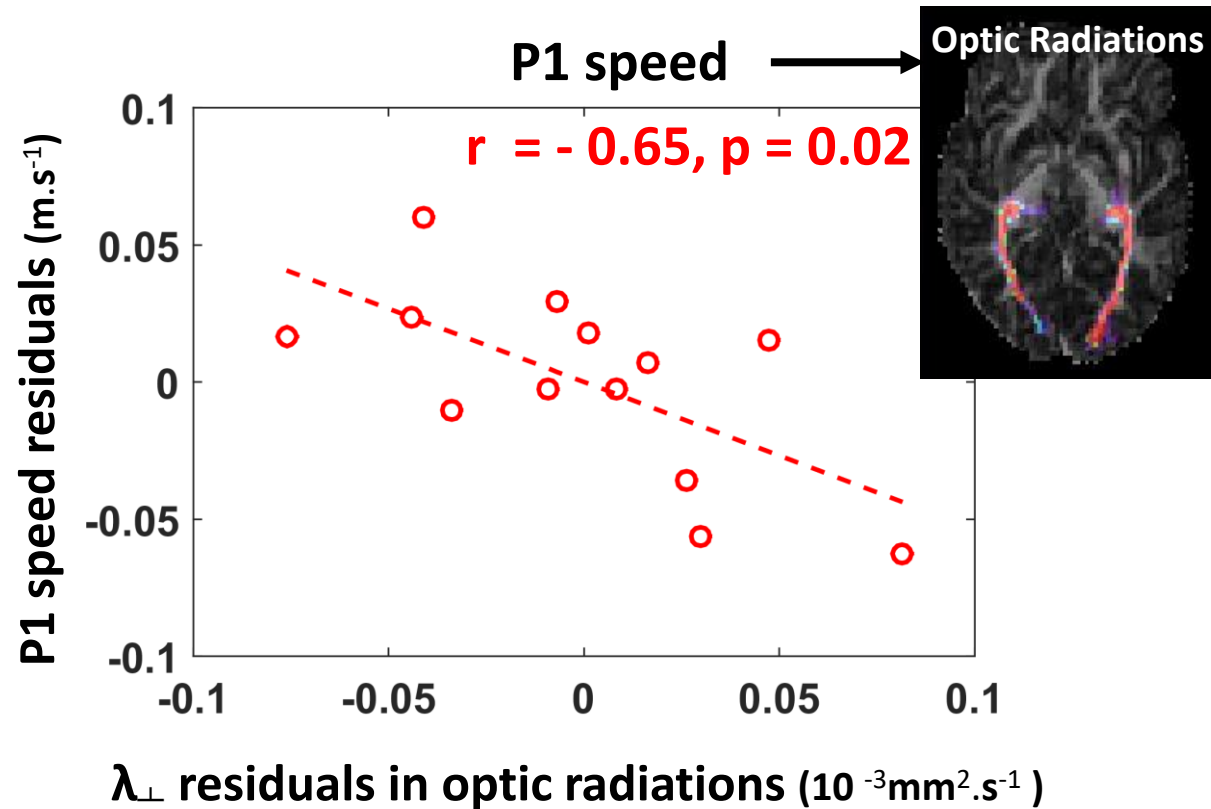
P1 transfer time:  
P1 Ipsi – P1 contra

Decrease in response latencies across development



# Correlations between structural and functional maturation

- 13 infants completed both EEG & MRI experiments



Controlling for age effects

- Variability in the response speeds relates to the maturation of corresponding pathways

# Structure-function relationship across multiple brain networks

Maturation of white matter

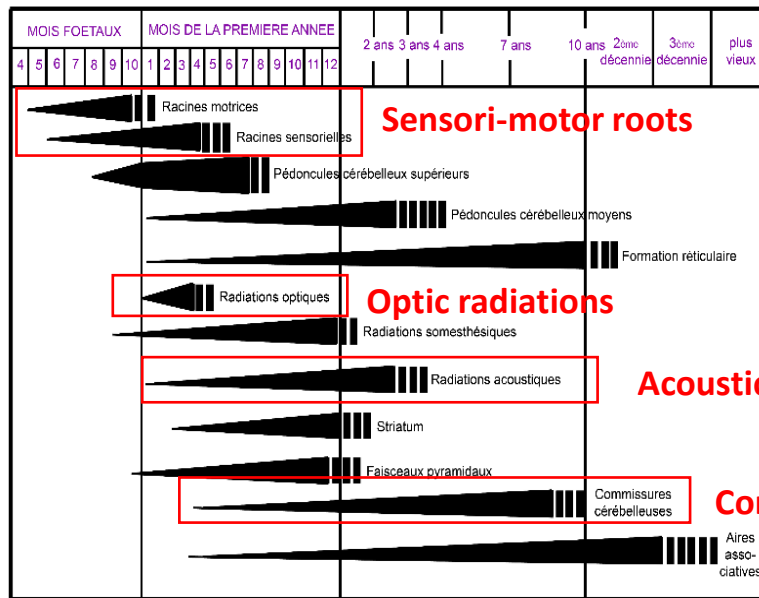
diffusion MRI

Maturation of brain responses

EEG



## Myelination of pathways



Visual network

Auditory network

Different Maturational calendars

Different *in utero/ex utero* inputs

Yakovlev and Lecours, 1967



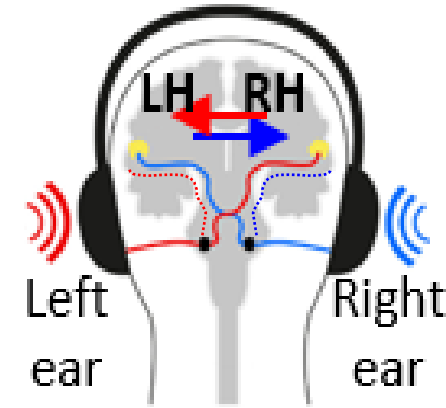
# MRI and EEG correlates of auditory development

- 19 infants (1-6 months)

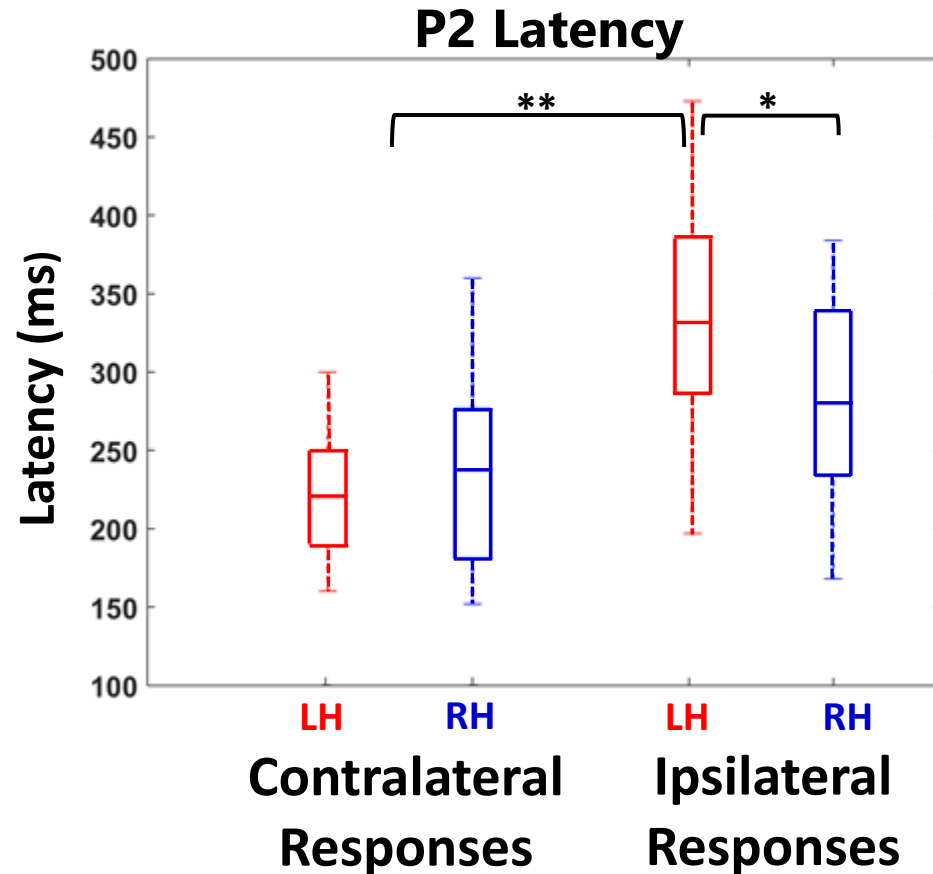
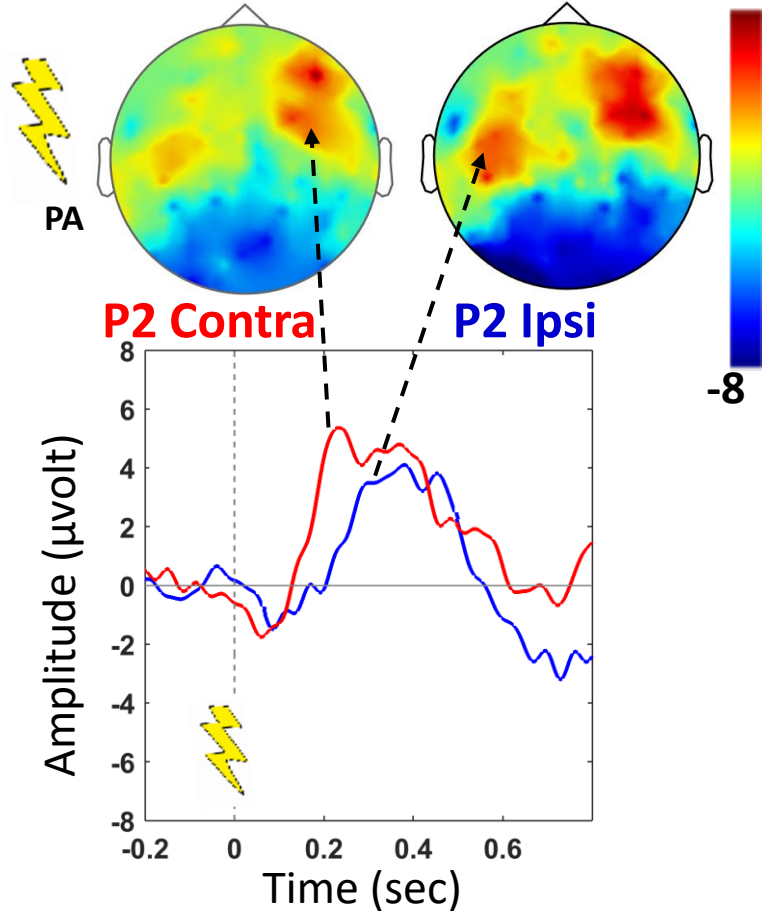


## Auditory pathways:

- Contralateral and ipsilateral pathways
- Responses from the ipsilateral pathways superpose to the transfer of responses



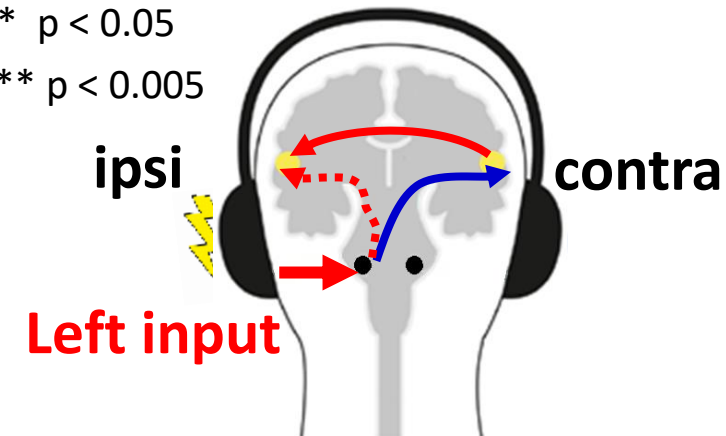
## Auditory evoked responses to monaural syllables



Left hemisphere (LH)  
Right hemisphere (RH)

\*  $p < 0.05$

\*\*  $p < 0.005$



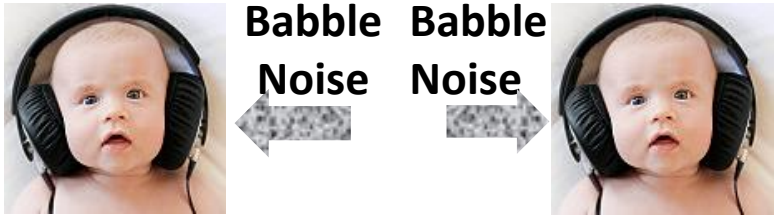
# Response latency differences between typical and AgCC infants

13 infants with **Agnesis of corpus callosum AgCC**

18 **Typical** infants

3-4 months

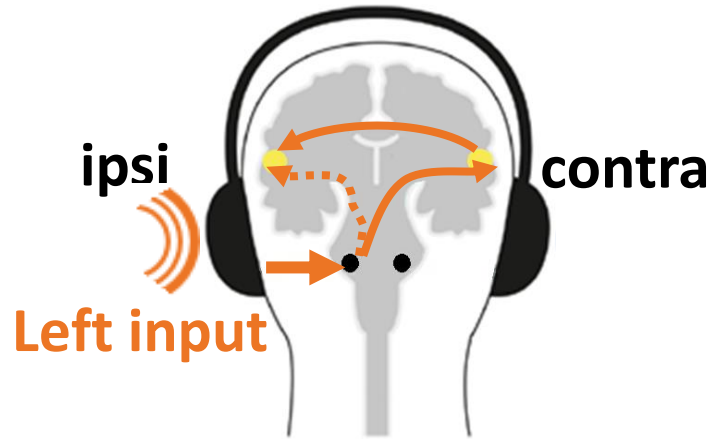
Auditory evoked responses to monaural noise



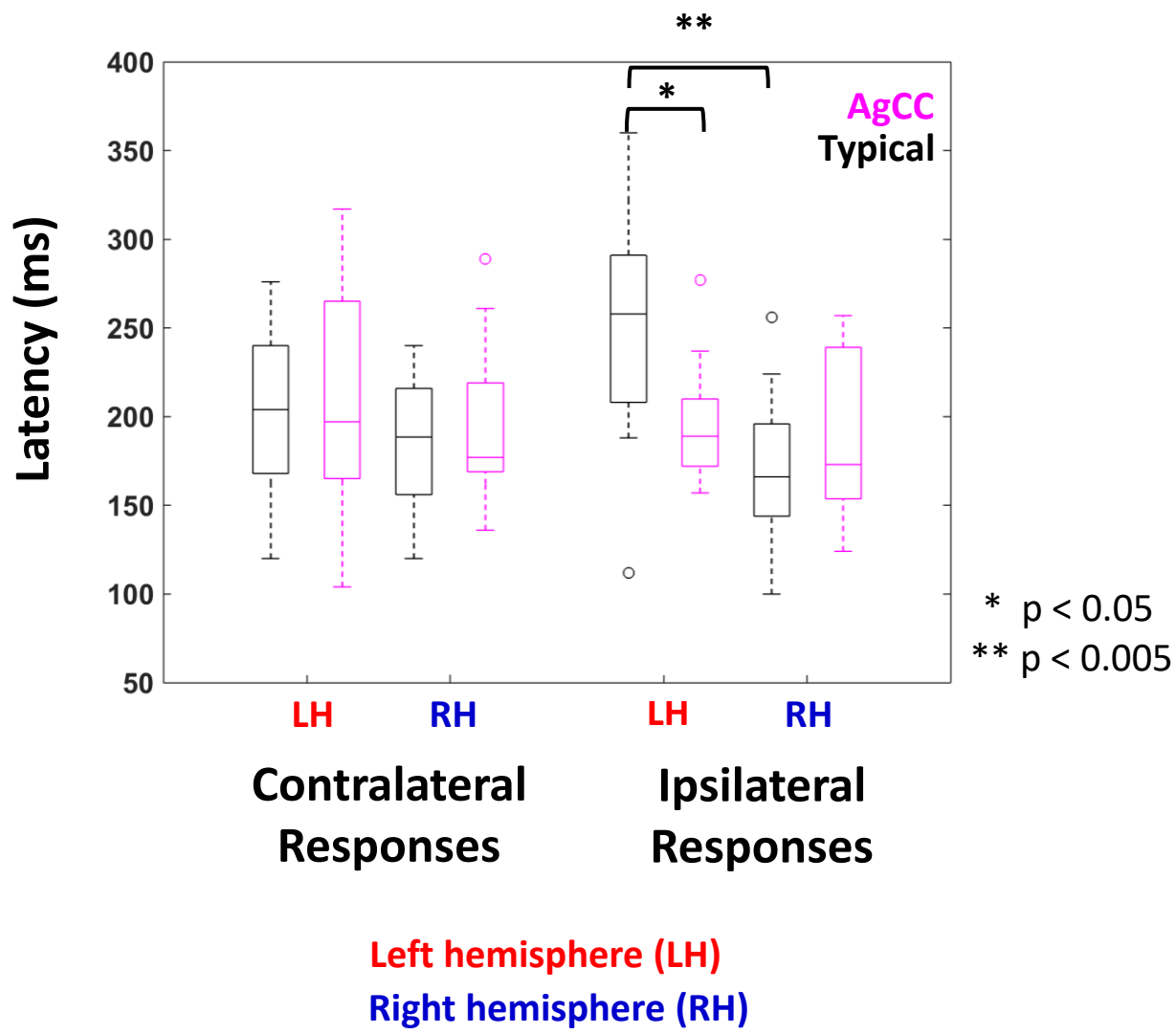
➤ **Delayed ipsilateral responses in the left (but not right) hemisphere in typical but not AgCC infants.**



Involvement of callosal pathways

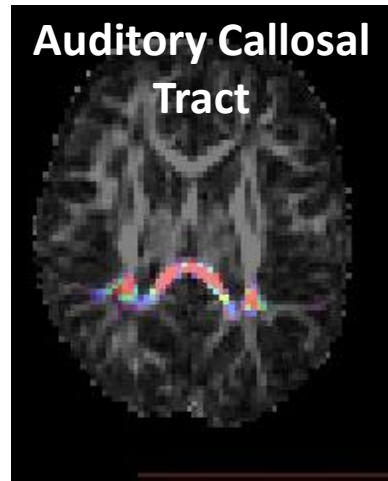
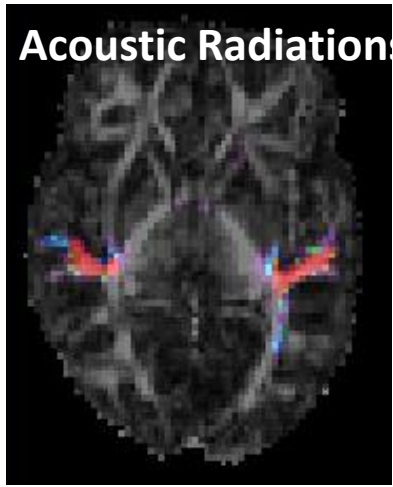


**P2 Latency in AgCC vs. typical infants**

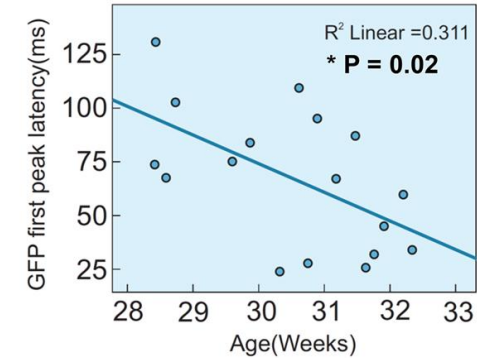
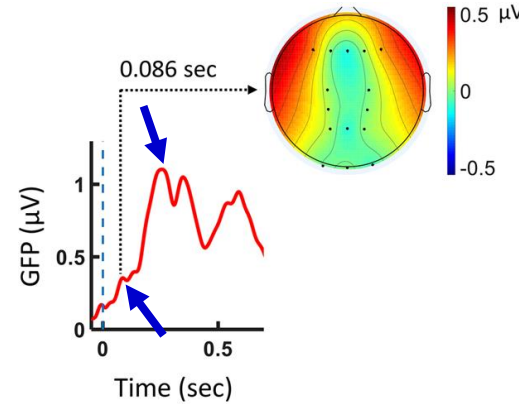


# Correlations between structural and functional maturation

## Maturation of auditory white-matter pathways

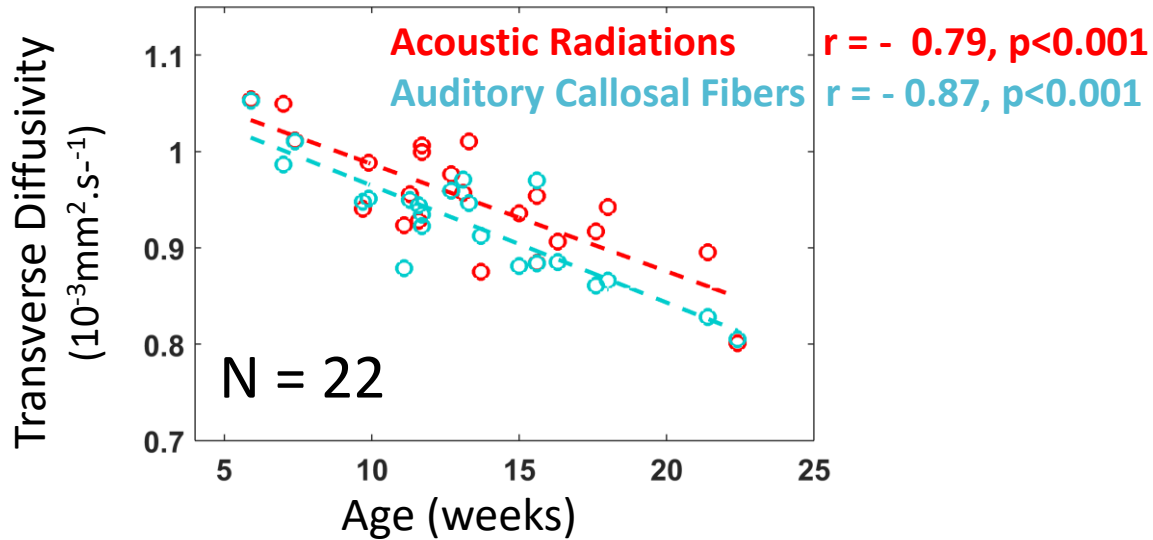


➤ No relation between the speed of P2 responses and the maturation of acoustic radiations → P2 is a late response



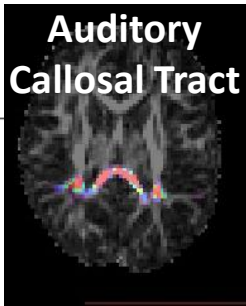
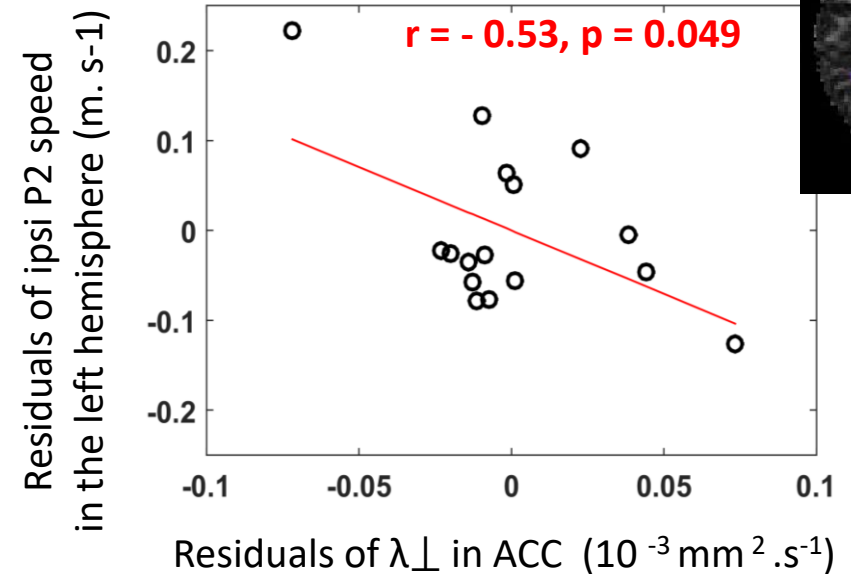
Daneshvarfard *et al.*, Brain Struc and Func, 2020

## Age-related changes



Adibpour *et al.*, Developmental Cognitive Neuroscience, 2020

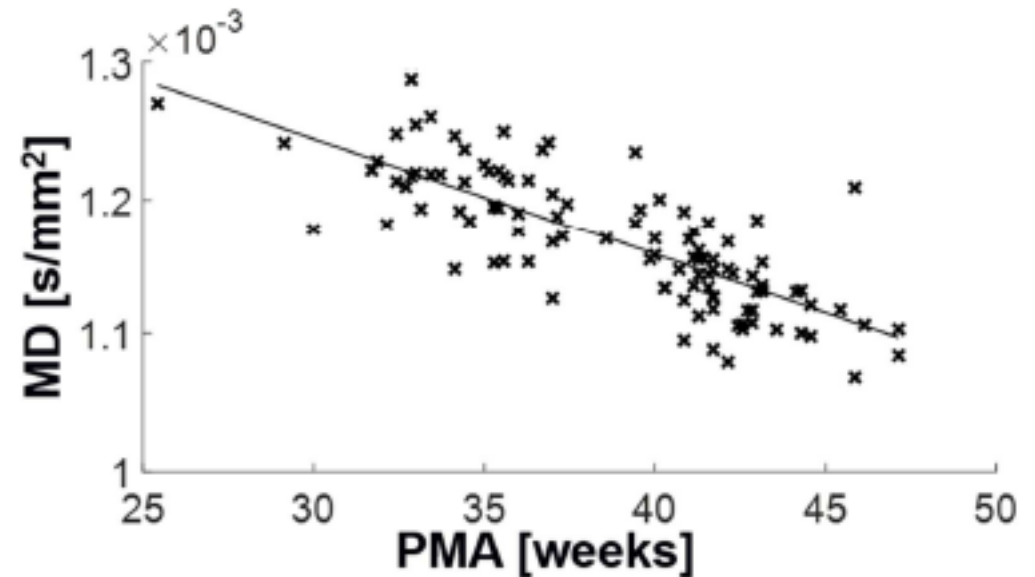
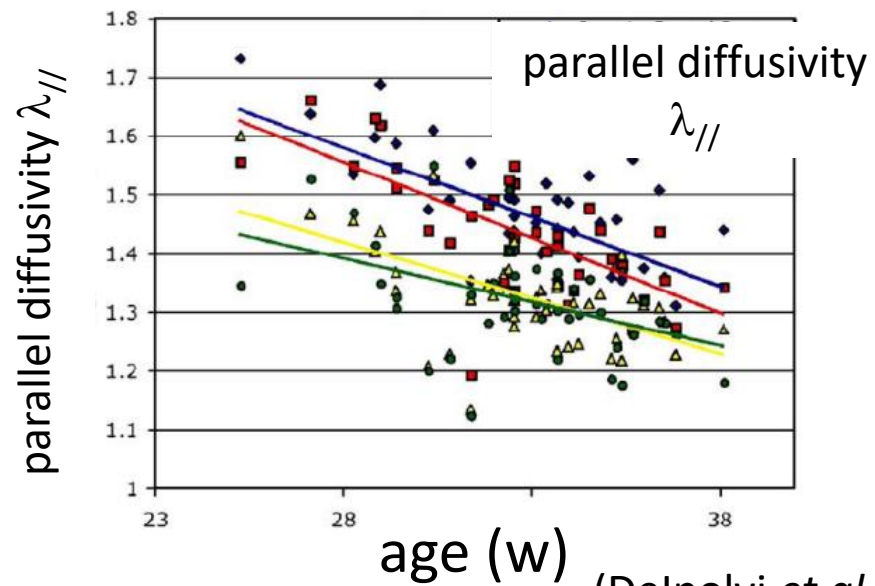
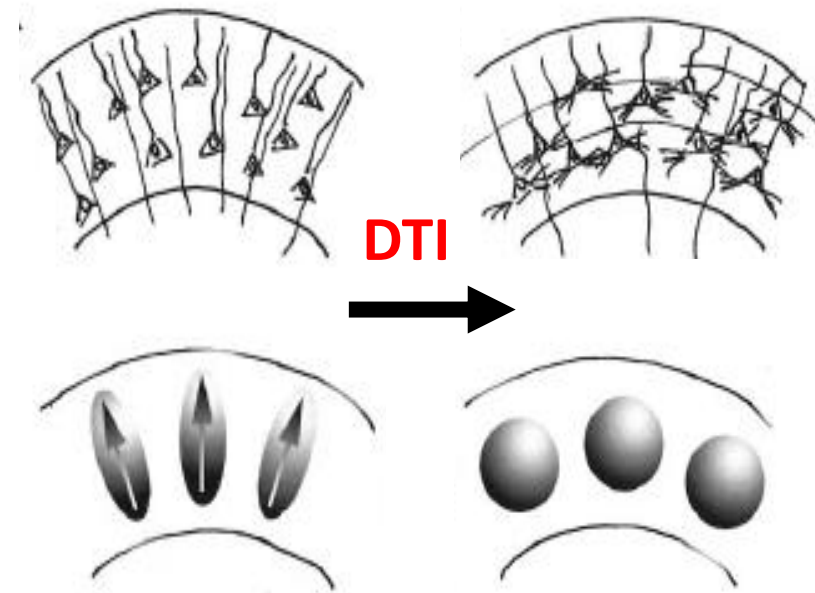
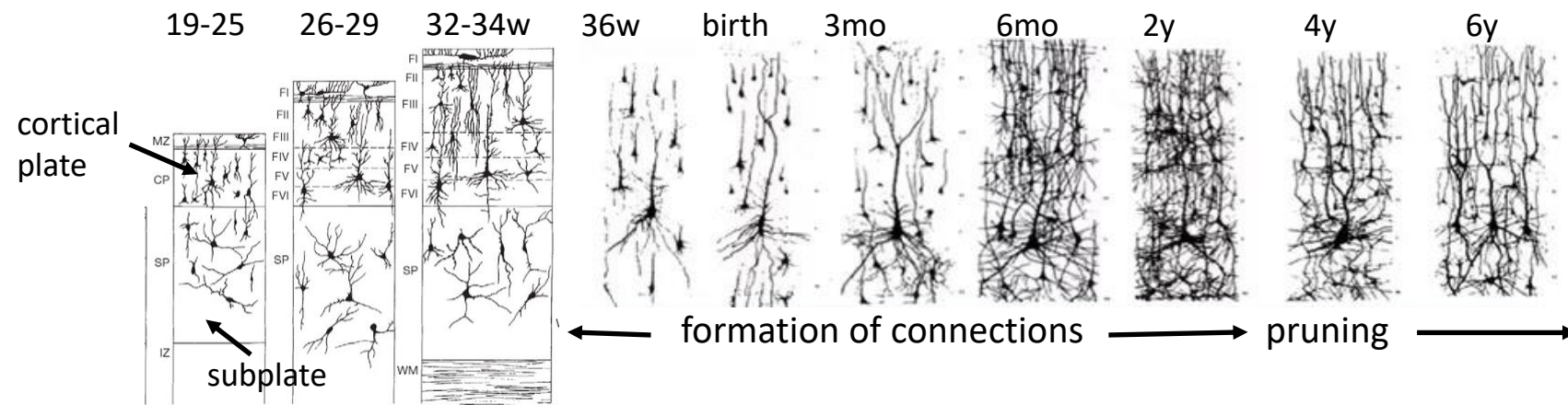
## Ipsilateral P2 latency in left hemisphere



# Cortical maturation

## Growth of dendritic arborization and synaptogenesis

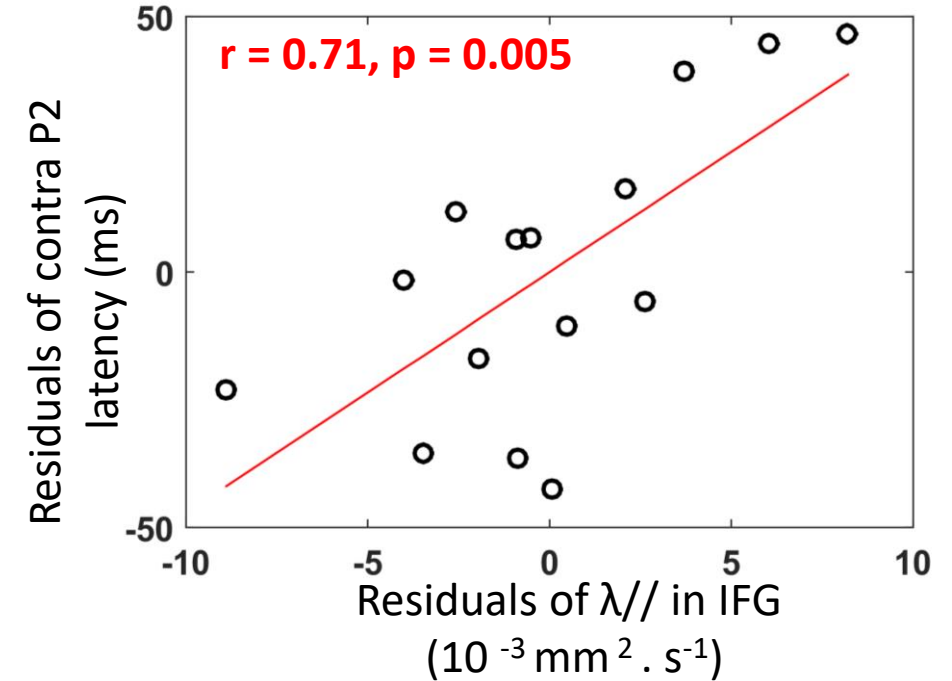
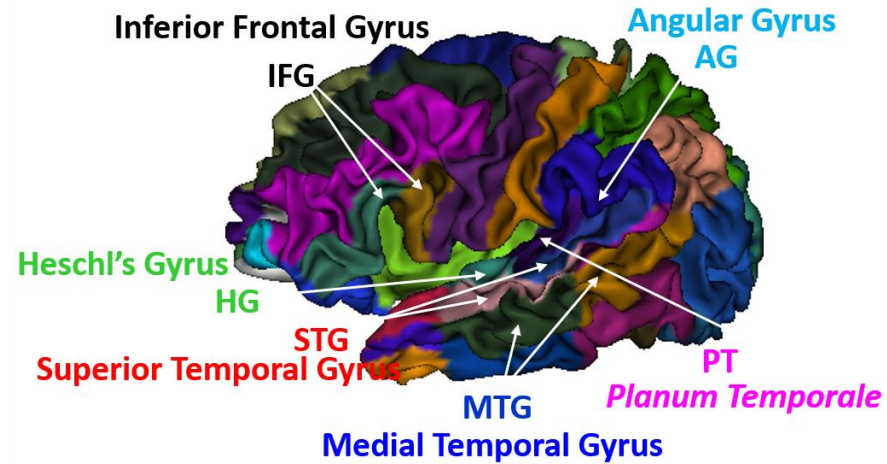
Cortex : during gestation / after birth



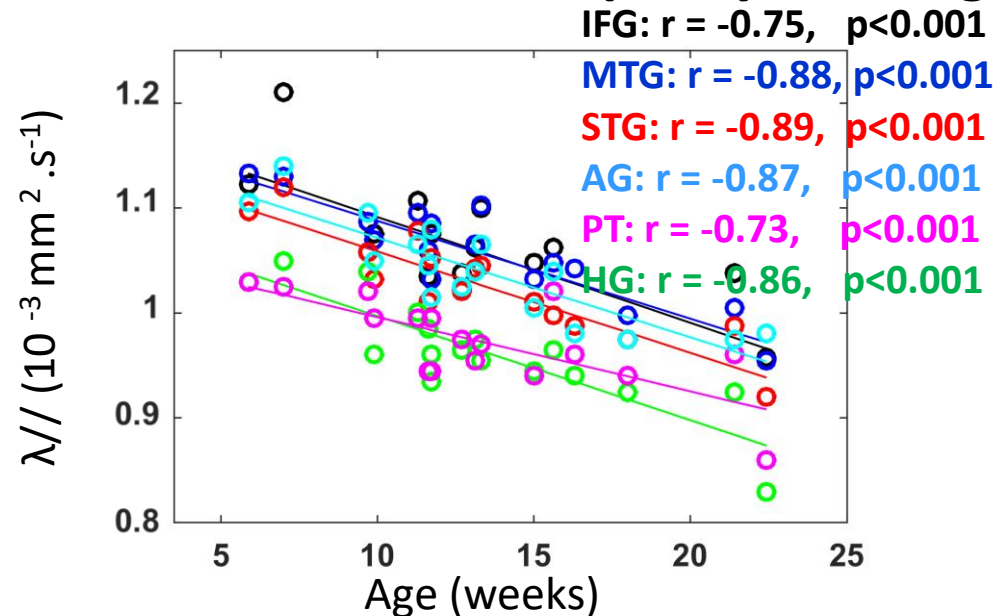
(Delpolyi *et al*, Neuroimage 2005; Bataille *et al*, Neuroimage, 2019)

# Correlations between structural and functional maturation

## Cortical parcellations in infants

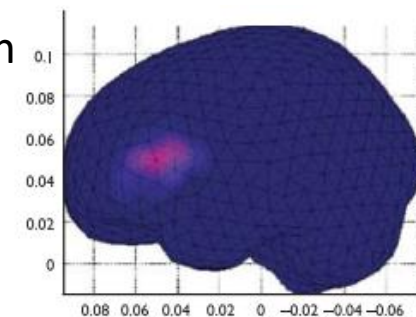


## Maturation of cortical perisylvian regions

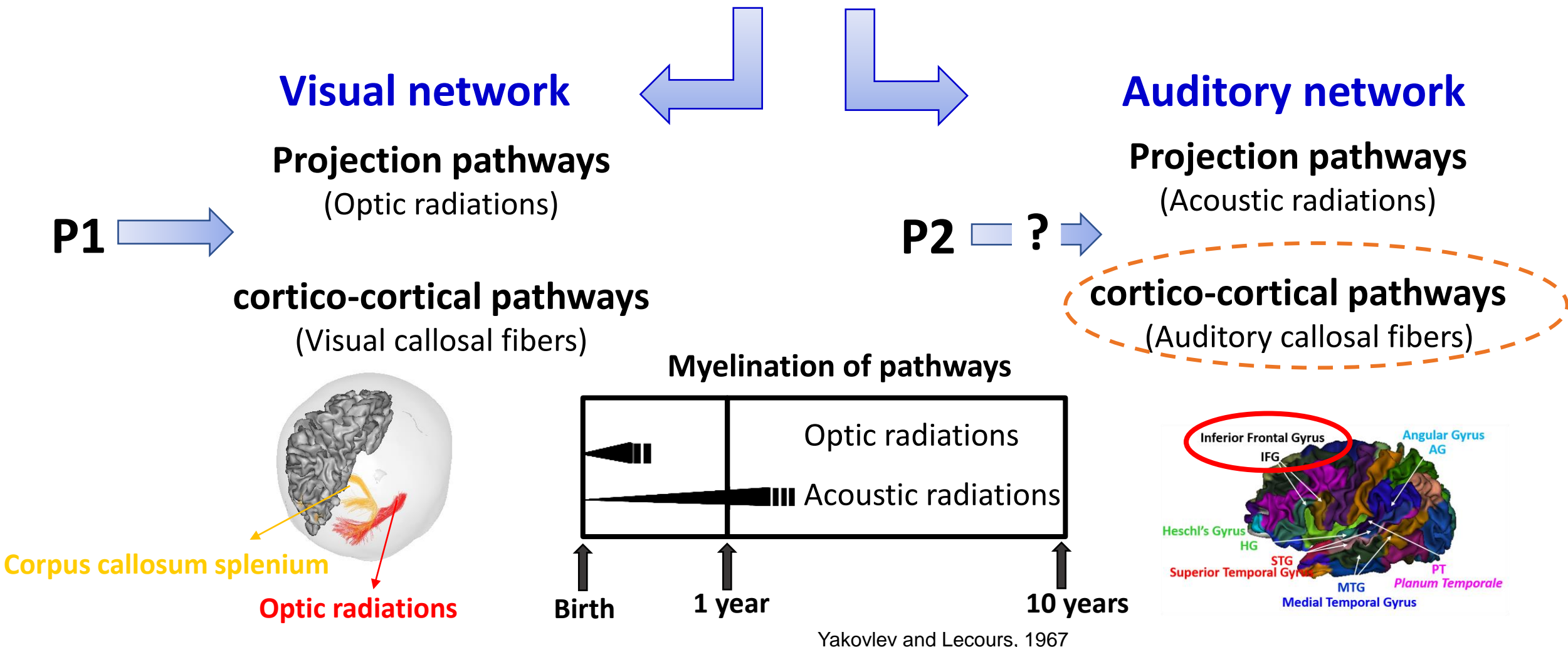


➤ Variability in the response latencies relates to the maturation of Inferior Frontal Gyrus

Source reconstruction of activity between 210-300 ms  
6 month old



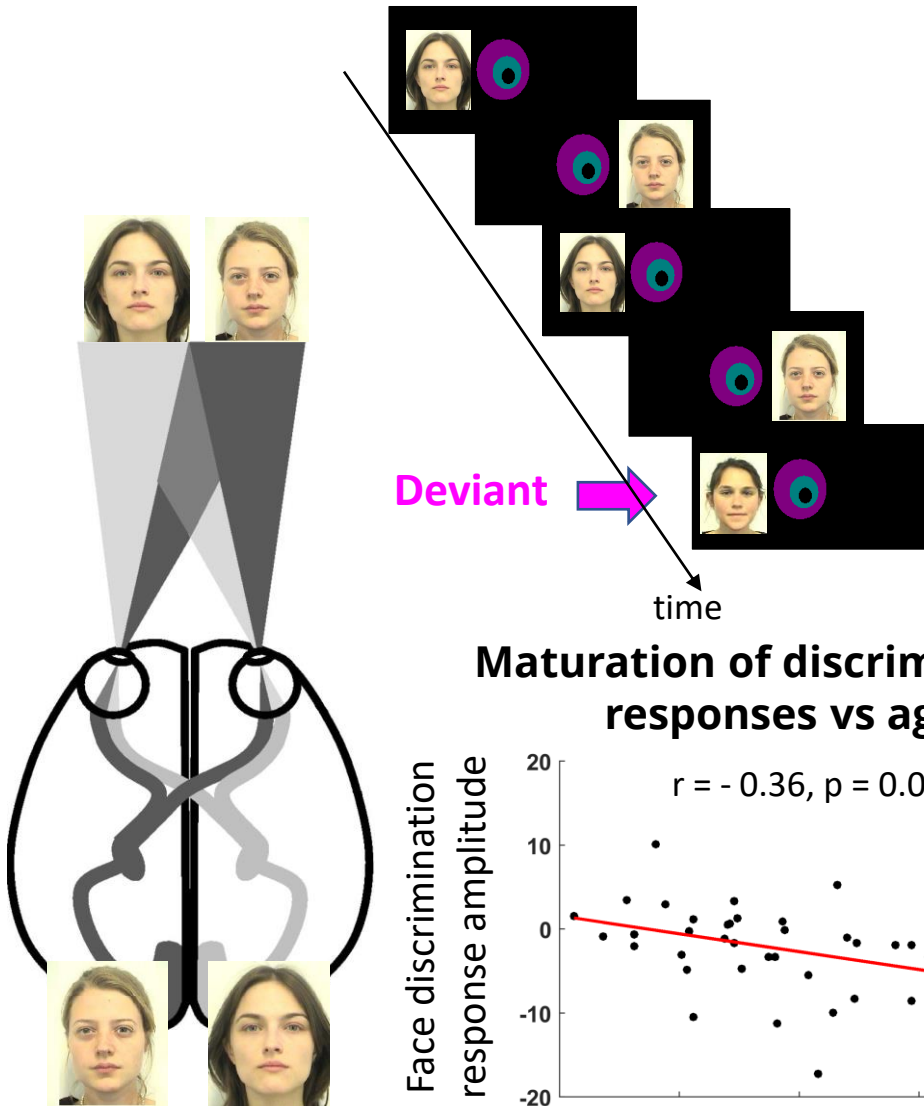
# Linking structural and functional maturation



➤ Early interhemispheric transfer of neural activity, supported by microstructural maturation

# Functional maturation beyond sensory processing

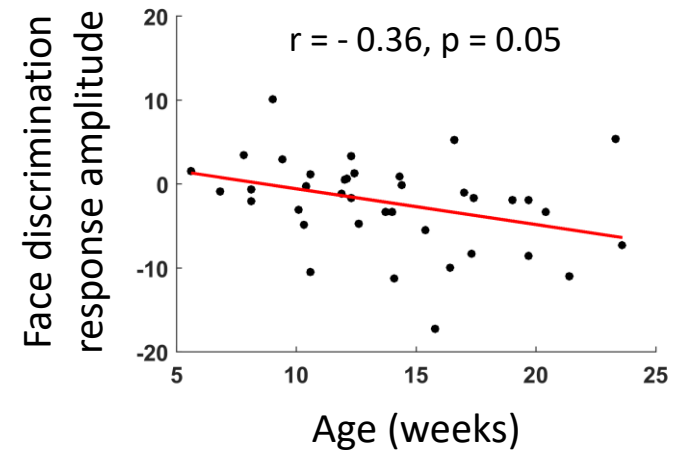
Face discrimination task



Deviant

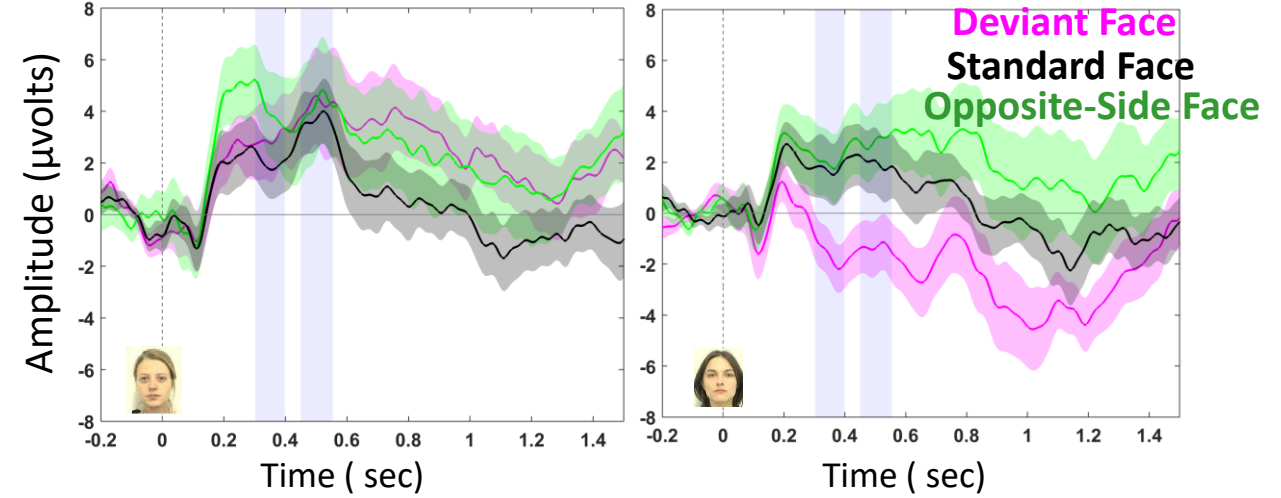
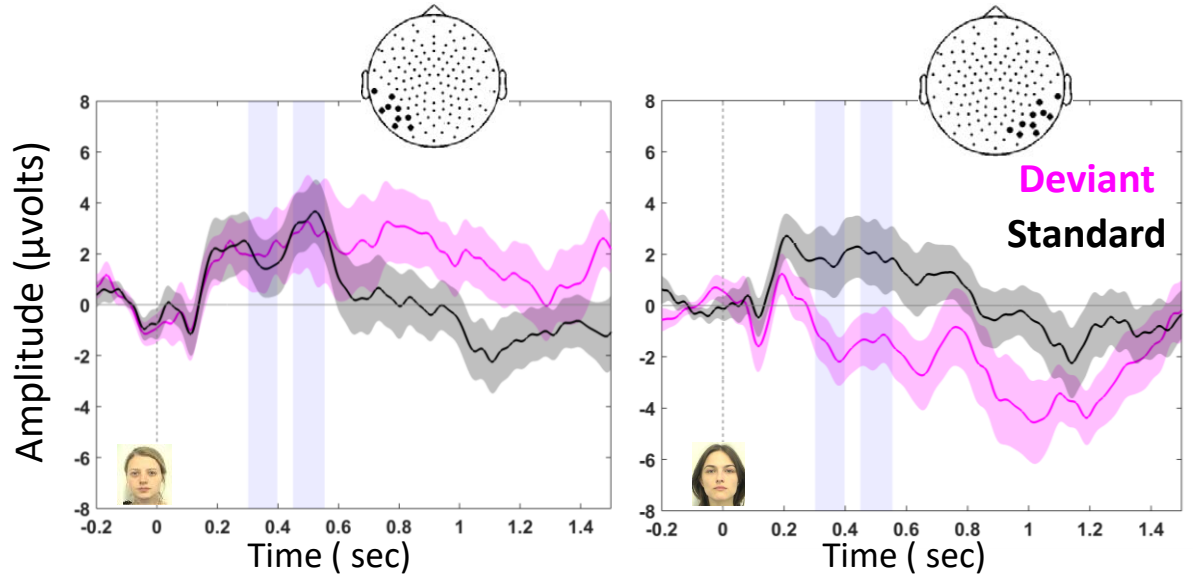
time

Maturation of discrimination responses vs age



Right Hemifield / Left Hemisphere

Left Hemifield / Right Hemisphere

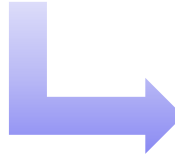


Missing link to structural maturation?

# Perspectives :

Structural maturation

MRI



Functional maturation

EEG/MEG (fMRI)



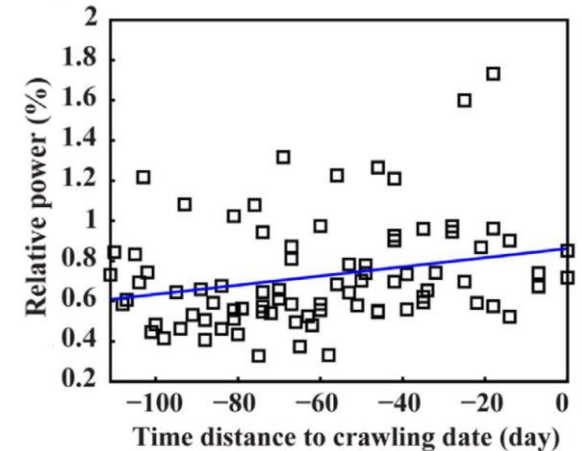
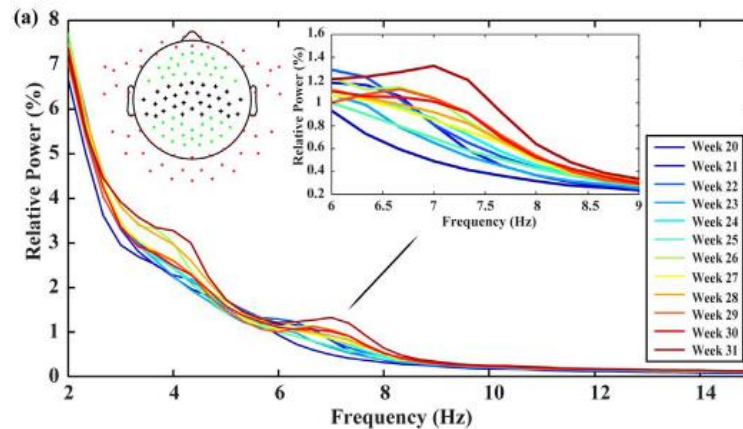
Behavioral acquisitions

## Sensori-motor acquisitions in relation with functional response maturation

### Mu rhythm and motor acquisitions

#### Crawling

- 5-7 months
- Rest-state recording
- The mu rhythm power specifically correlated to the crawling onset.



Xiao et al., Neuroimage, 2017

➤ In early developmental period, study of behavioral acquisitions are limited.

Structural maturation



Functional maturation

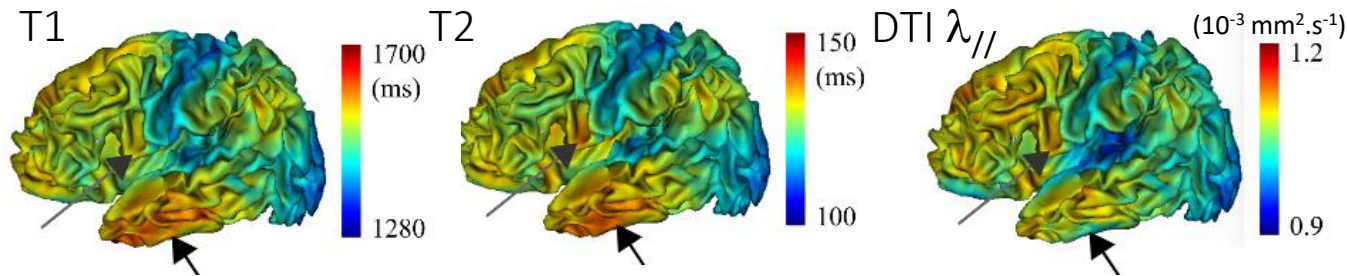


# Perspectives :

➤ Moving towards a finer description of structure-function relationships:

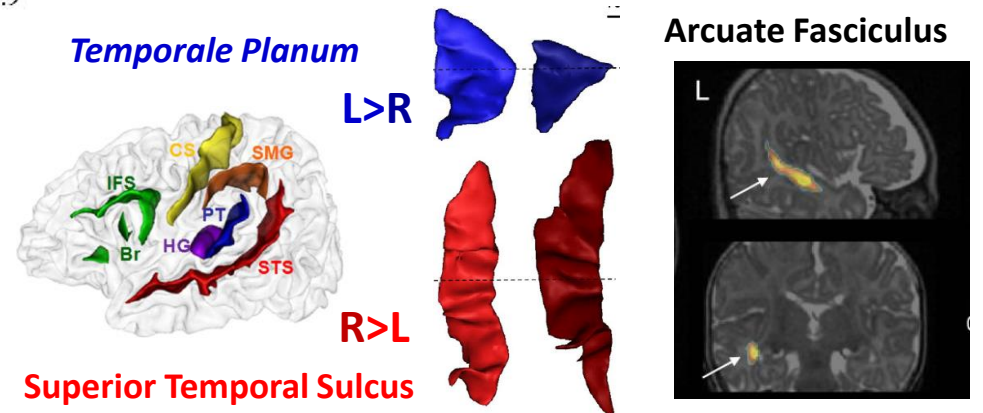
➤ Quantifying other functional parameters: e.g. morphology and amplitude of evoked responses

➤ Quantifying other microstructural parameters (Lebenberg et al., Brain Plasticity, 2016)



Lebenberg et al., IEEE ISBI 2015, Neuroimage 2019

➤ Assessing the structure-function relationships within lateralized networks, given the structural and functional asymmetries described in the infant brain.

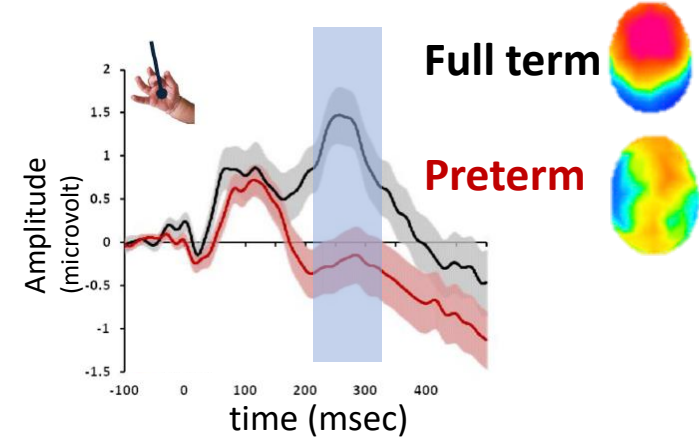


Glasel et al., Neuroimage, 2011 Dubois et al., Cerebral Cortex 2009

➤ Investigating the predictive values of the measures for later outcomes (e.g. language)

(O'Muircheartaigh et al., J Neuroscience, 2013)

Sensory(tactile) response maturation



Maitre et al., Current Biology, 2017

# Thank you for your attention.

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## Acknowledgements:

All infants and their parents

## Collaborators:

Jessica Dubois, Ghislaine Dehaene-Lambertz, Marie-Laure Moutard

Jessica Lebenberg, Claire Kabdebon, Francois Leroy

UNIACT-Neurospin team, Giovanna Santoro

## Funding resources:



European  
Research  
Council