



Brain & Cognitive Development: A Focus on Myelination and Executive Functions

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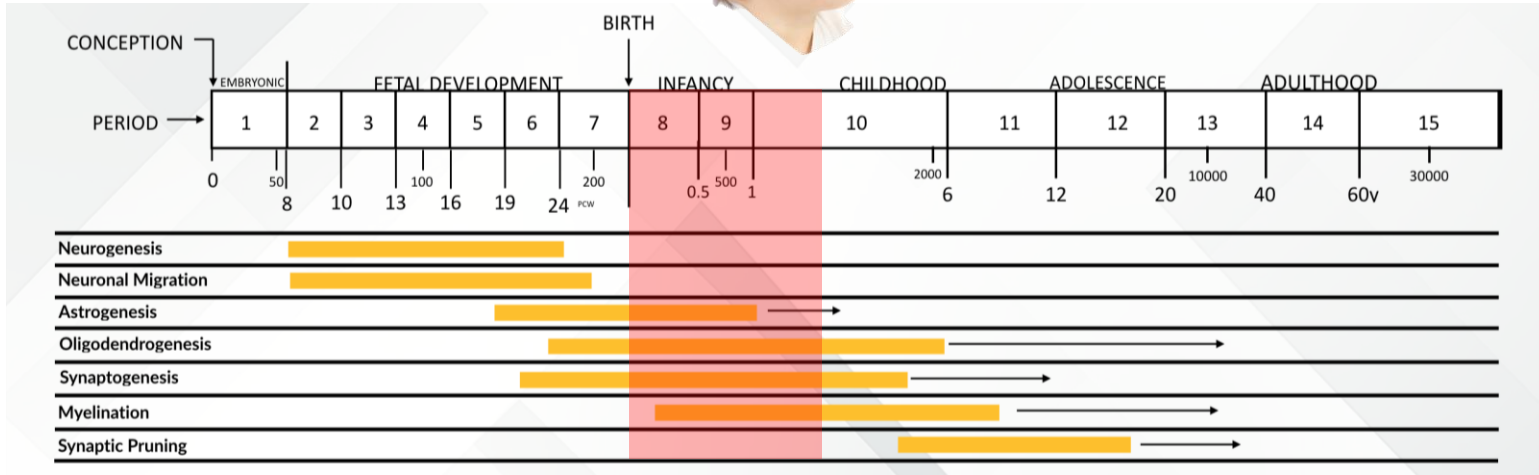
9th Annual Wyeth Nutrition Science Center Global Summit

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Presentation Outline

- Brain Development in Early Life
- Brain Structure & Function: Myelination and Cognition
- Executive Functions – building core cognitive capabilities for life

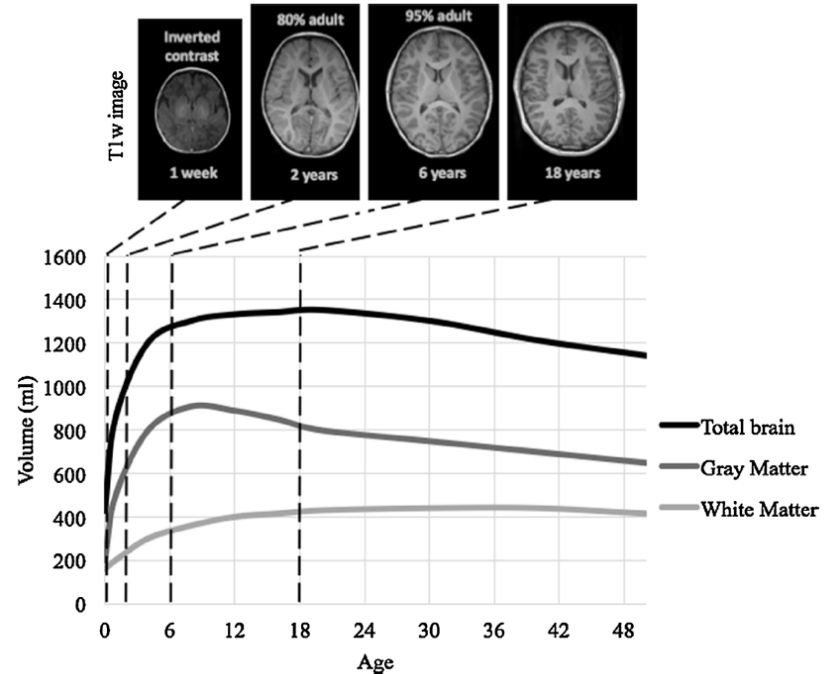
Brain Development – early life



Adapted from Silbereis et al., 2016

Brain Development – early life

- Children's brains are significantly different from the adult brain as the organization and properties of brain structures change with age¹
- The early brain develops with a level of plasticity; this facilitates both adaptive changes, representing potential **opportunity**, and malformations, reflecting potential vulnerability²



¹Vân Phan et al., 2018, ²Gao et al 2017

Influencing Factors

Gene expression (nature)

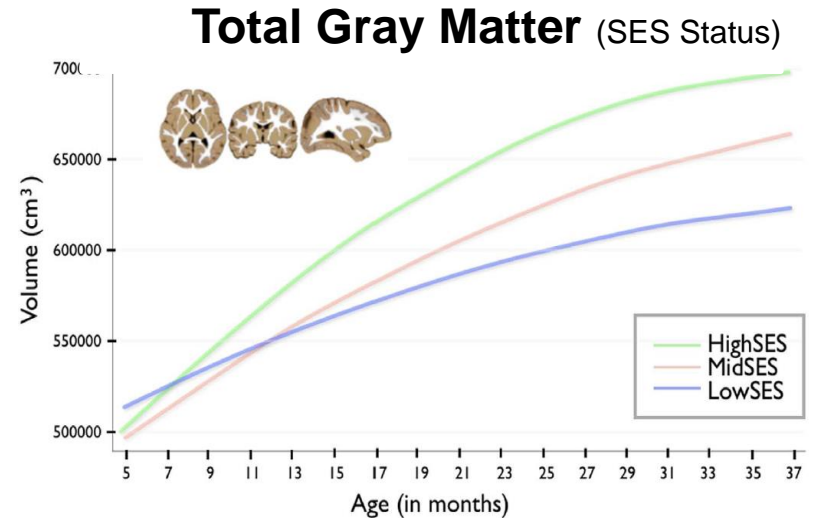
Environmental factors (nurture)

→ Molecular cues that guide development and are dependent on environment and experiences of the developing child

Influencing Factors

Environmental factors

- Socioeconomic status (SES)
- Social mobility
- Nutrition
- Social interactions
- Stress
- Urbanisation
- Pollution



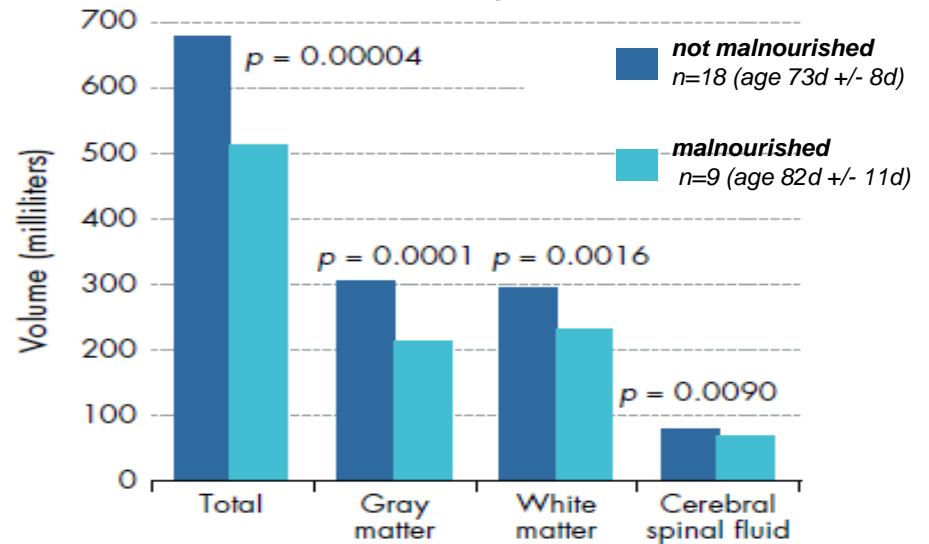
Hanson J.L. et al., PlosOne, 2013

Influencing Factors

Environmental factors

- Socioeconomic status (SES)
- Social mobility
- Nutrition
- Social interactions
- Stress
- Ubranisation
- Pollution

Total White and Gray Matter (stunting status)

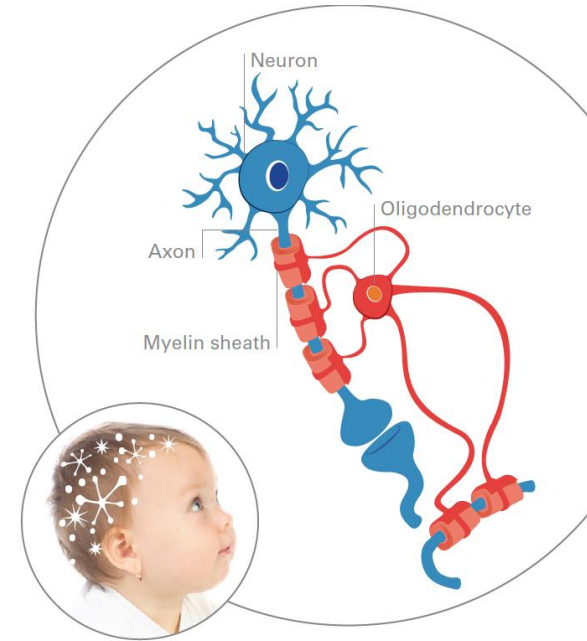


Source: worldbank.org/en/publication/WDR2018, using data from Nelson and others (2017). Data at http://bit.do/WDR2018-Fig_S2-1.

Myelination

Myelination is a key part of neurodevelopment^{1,2}

- Wrapping of nerve fibers (axons) with lipid-rich sheath
- Ensures fast, efficient & synchronized communication between cells and networks
- Protects the neuron
- Matures alongside cognitive and behavioural development

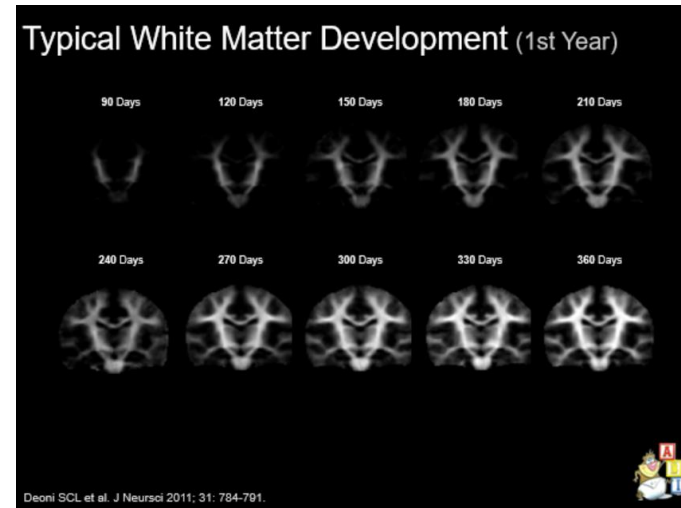


¹Silberis 2016; ²Deoni et al 2011

Myelination & Cognition

Clinical studies demonstrate the link between myelination and cognition, including:

- general cognitive ability¹
- language² & reading³
- working memory⁴
- processing speed⁵
- sensory reactivity⁶



¹Schmithorst et al., 2005; Deoni et al., 2015; ²Büchel et al., 2004; O'Muircheartaigh et al., 2013; ³Nagy et al., 2004; Beaulieu et al., 2005; ⁴Nagy et al., 2004; Short et al., 2013; ⁵Turken et al., 2008; Bartzokis et al., 2010; Lu et al., 2013; ⁶Weinstein et al., 2014

Myelination & Cognition¹

Objective

To examine the longitudinal relationship of maturing brain myelination to variability in cognitive skills.

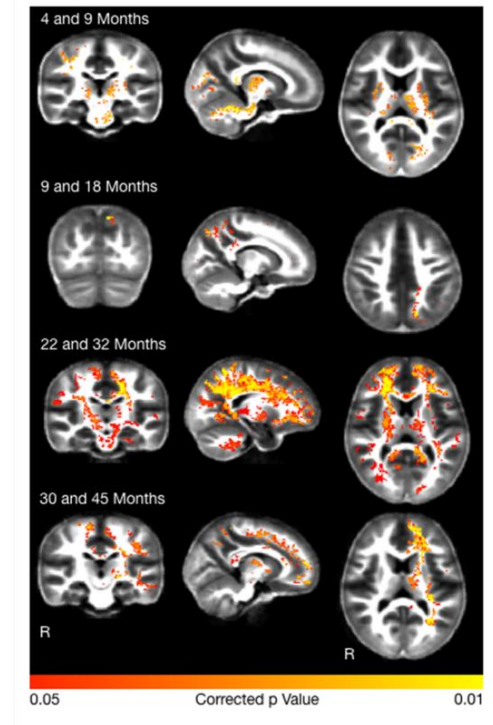
Population

N = 187 children with repeated MRI (Myelin Water Fraction) and cognitive assessments (MSEL) between 2 mo and 6 yrs of age

Results

Marked age-related variability in repeated cognitive measures that was variably related to on-going changes in myelination.

¹Deoni et al 2019

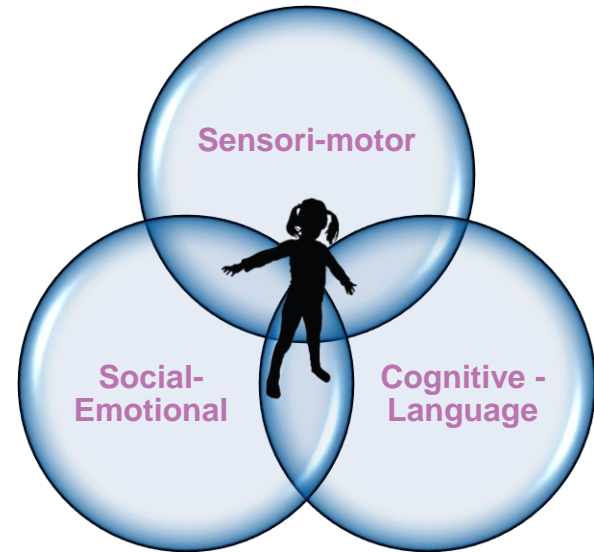


Brain & Cognitive Development

- Changes in neurobiology parallel behavioral maturation → milestones¹

Continuous learning process of gaining:

- Sensory
- Motor
- Cognitive
- Language
- Social &
- Emotional abilities



¹Silbereis et al., 2016

What are Executive Functions (EF)?

- EFs are a family of key **mental processes** that **organize**, manage **information** and control **behavior**¹
- 3 core EF:
 - Inhibition (inhibitory control, including self control (behavioural inhibition and interference control (selective attention and cognitive inhibition))
 - Working memory
 - Cognitive flexibility (also called set shifting and mental flexibility)

¹Diamond A. 2013

What are Executive Functions (EF)?

3 core Executive Functions^{1,2}



- Planning and organisation
- Flexible thinking
- Emotions & behavior controls
- Multi-tasking
- Solving complex problems
- Learning rules
- Making decisions
- Motivation
- Concentrating
- Self-awareness

¹ Diamond A. (2013); ² Collins & Koechlin (2012)

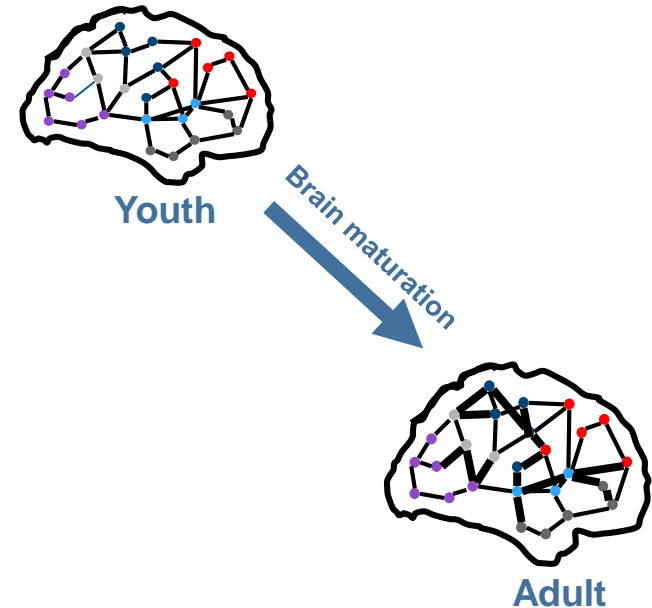
Development of EFs – related to other cognitive skills

- Children are not born with EFs skills, but with the ability to develop them¹
- EF draw on other, earlier developing cognitive skills
- Generally cognitive development proceeds from sensory, motor and early language activity to social interaction and finally higher order cognition and executive functions²

¹ <https://developingchild.harvard.edu/science/key-concepts/executive-function>;²Dajani et al 2015

Development of EFs – related to brain connectivity

- EF are orchestrated by neural activity within the prefrontal cortex (PFC)¹ that is connected with other brain regions
- Those networks of different brain regions continuously process and share information with each other, either via anatomical links (structural connectivity) or via simultaneous activation of spatially separate regions (functional connectivity)²⁻⁴ and they develop from local to wide-spread networks



¹Best & Miller (2010), ²Huang & Ding (2016), ³van den Heuvel & Hulshoff (2010), ⁴Sporns O. (2013).

EFs as core cognitive capabilities for life

- EF are skills for life and learning¹
- EFs are skills essential for success in school and in life as well as cognitive, social, and psychological development²
 - Mental & physical health
 - Quality of life
 - School readiness
 - Job success
 - Marital harmony
 - Public safety
- They can be more predictive than IQ or socioeconomic status³

¹ www.developingchild.harvard.edu/resources/; ²Diamond A (2013), ³Diamond A (2016).

Factors influencing the development of EFs¹⁻⁴



- Neural maturation
- Schooling & education
- Language
- Social environment & connections
- Positive relationships with adults
- Physical & mental activity
- Practice
- Creative play



- Factors disrupting brain architecture
- Negative stress
- Neglect & violence

¹ <https://developingchild.harvard.edu/science/key-concepts/executive-function/>; ² Matsuda et al. 2017; ³ Baum et al. 2017; ⁴Best & Miller (2010)

Wrap up & key messages

Key messages

- The first years of life are a rapid & dynamic period for brain maturation. Most brain processes during that period are primarily focused on connecting the brain, e.g. myelination
- The resulting connectivity is central to establishing cognitive functions, such as Executive Function (EF). EF development and maturation is fundamental to success in school, life as well as health & well-being.
- Many factors influence brain growth and myelination; early life nutrition is an important and modifiable factor that can shape myelination and, consequently, cognitive development including executive functions



Brain & Cognitive Development: The Role of Polar Lipids

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