

What can light show us about brain activity?: Examining early neural and behavioral predictors of childhood ADHD



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ADHD Diagnostic Criteria (DSM 5)

Listed as a Neurodevelopmental Disorder

Six or more symptoms of Inattention and/or Hyperactivity/Impulsivity

- Symptoms persist for at least 6 months
- Symptoms are present prior to age 12
- Symptoms are present in two or more settings
- Symptoms are not better explained by another mental disorder

Severity of Symptoms and Impairment

Executive Function Skills

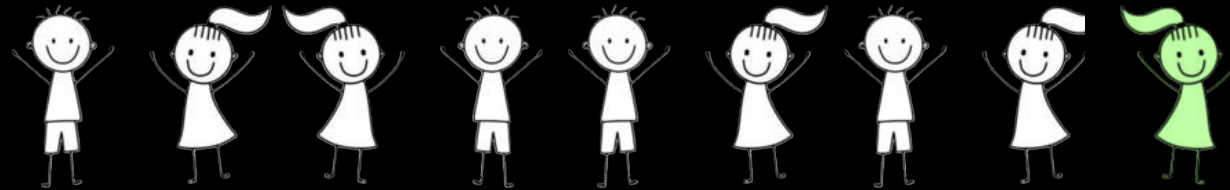
A set of cognitive processes necessary for goal executions

| Impulse Control | Working memory | Attention | Problem solving (Planning & organizing) |
|---------------------------------------|---|----------------------------|---|
| Responds without considering options | Difficulty keeping track of time | Distractible* | Difficulty forgoing current pleasure for future rewards |
| Easily excited, impatient, frustrated | Difficulty keeping track of belongings | Difficulty 'multi-tasking' | Impulsive actions (without regard for consequences) |
| Restlessness & hyperactivity | Difficulty following conversations for appropriate interactions | Fails to finish activities | Difficulty organizing tasks and activities |

* Does well with novel or stimulating activities ("hyperfocus")

Many children and adults are affected by ADHD

~1 in 9 children in the U.S.



ADHD persist into adulthood for 2 out of 3 children with ADHD

Individuals with ADHD often experience challenges in many aspects of life



What Causes ADHD?

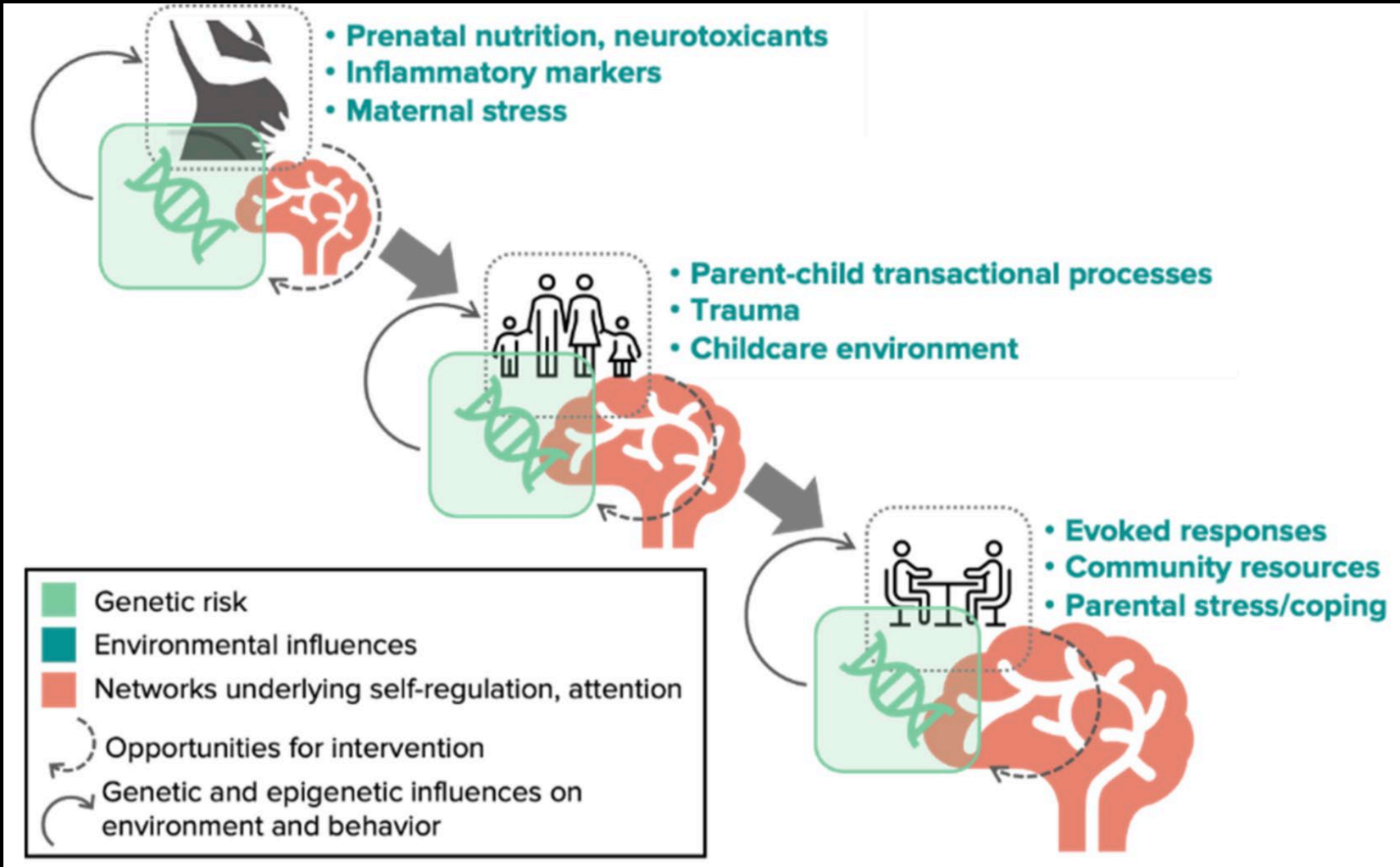


Figure adapted from
Miller et al., 2023 *JCPP Advances*

Featured on the ICIS Baby Blog

Why Study ADHD?

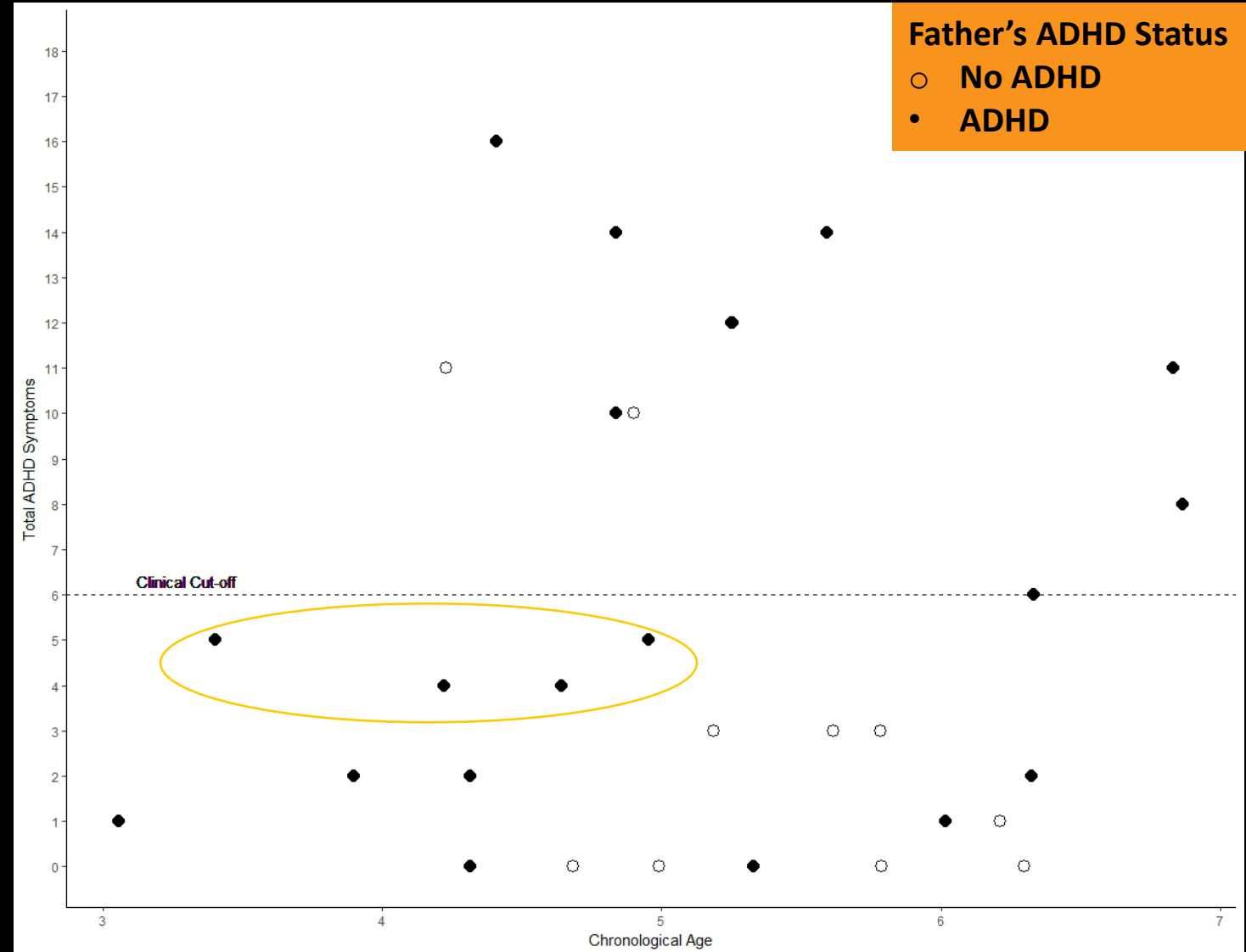
ADHD is a common reason to present for psychiatric care in childhood

Children are often referred for treatment between 7-9 years of age due to challenges occurring at school and home



Pittsburgh ADHD Longitudinal Study (PALS) Offspring Pilot Study

- Examined ADHD symptoms and behavior of 29 young children born to fathers with and without h/o ADHD
- **2.76 Odds of ADHD if father has ADHD**
- 4 Preschool aged children (3-5yrs) were subthreshold for disorder



Joseph et al., 2019 Journal of Child Psychiatry and Human Development
Funding provided by AACAP Pilot Award for Attention Disorders

Inattention, hyperactivity, and impulsivity are common among preschoolers



Less than 3% of preschool-aged children receive a diagnosis of ADHD

- 70-90% continue to meet diagnostic criteria in later childhood



*What do we know
about the brain
and ADHD?*

*Can that inform early
identification?*

Children and adults with ADHD have been found to have reduced brain volumes



Prefrontal cortex (PFC) involved in executive functioning (planning, decision making, working memory, impulse control).

Parieto-temporal region is involved in attention.

Friedman & Rapoport (2015) *Current opinion in neurobiology*;
Hoogman et al. (2019) *American Journal of Psychiatry*

Brains of children with ADHD appear younger than same age peers without ADHD

Longitudinal MRI studies have identified **delayed brain maturation**

Brains of children with ADHD appear ~3 years younger than children without ADHD



The background is a composite image with a blue-to-red gradient. It features several axial brain MRI slices. Overlaid on these are various technical labels and data points in white text. Labels include 'FoV 199', '296*512', 'Sag(1.5)', 'Tra>Cor(6.1)', 'W 128', 'C 66', 'Chilman', 'Harmon', '4VA12', 'HFS', '+LP', 'AF', 'RFP', '5cm', and 'R'. There are also some numbers like '23 / 180' and '280663'.

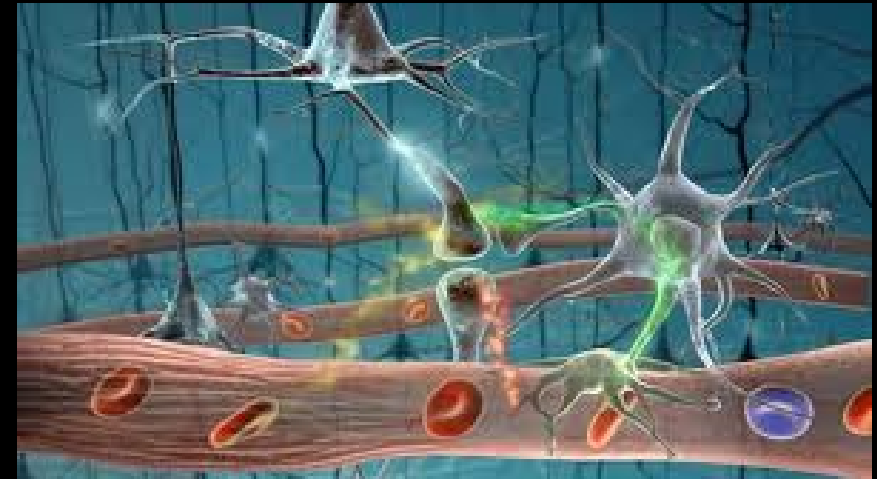
Moving from brain structure to function

How is brain activity measured?

Neurons use oxygen to produce energy needed to function.

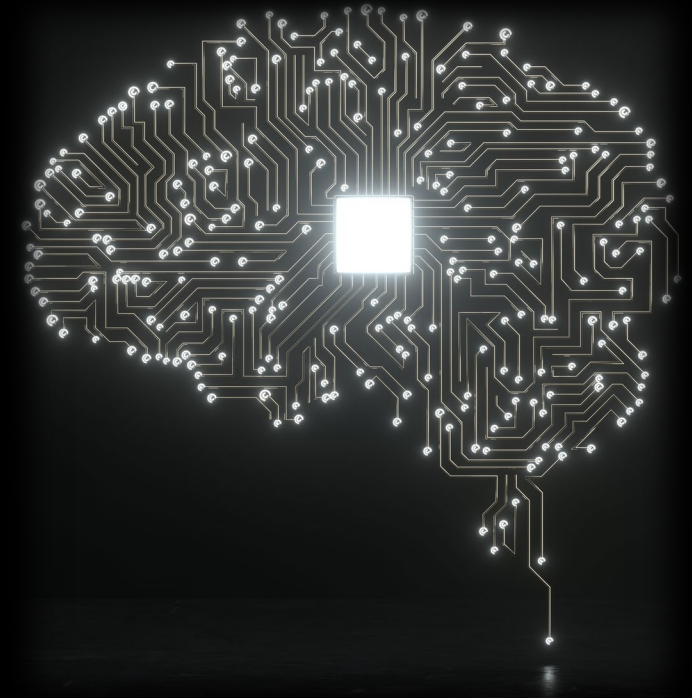
As neurons increase firing rates, the oxygen consumption also increases.

By detecting changes in oxygen levels in the brain we can identify which areas have neurons that are active.



Children and adults with ADHD have altered brain function

Individuals with ADHD have been found to have altered brain activity in areas involved in the control of attention, cognition, and inhibitory control.



What about preschool-aged children?

This is a period of rapid brain development

- Ideal for interventions

Very few MRI studies with preschool-aged children with ADHD

- Limited due to motion

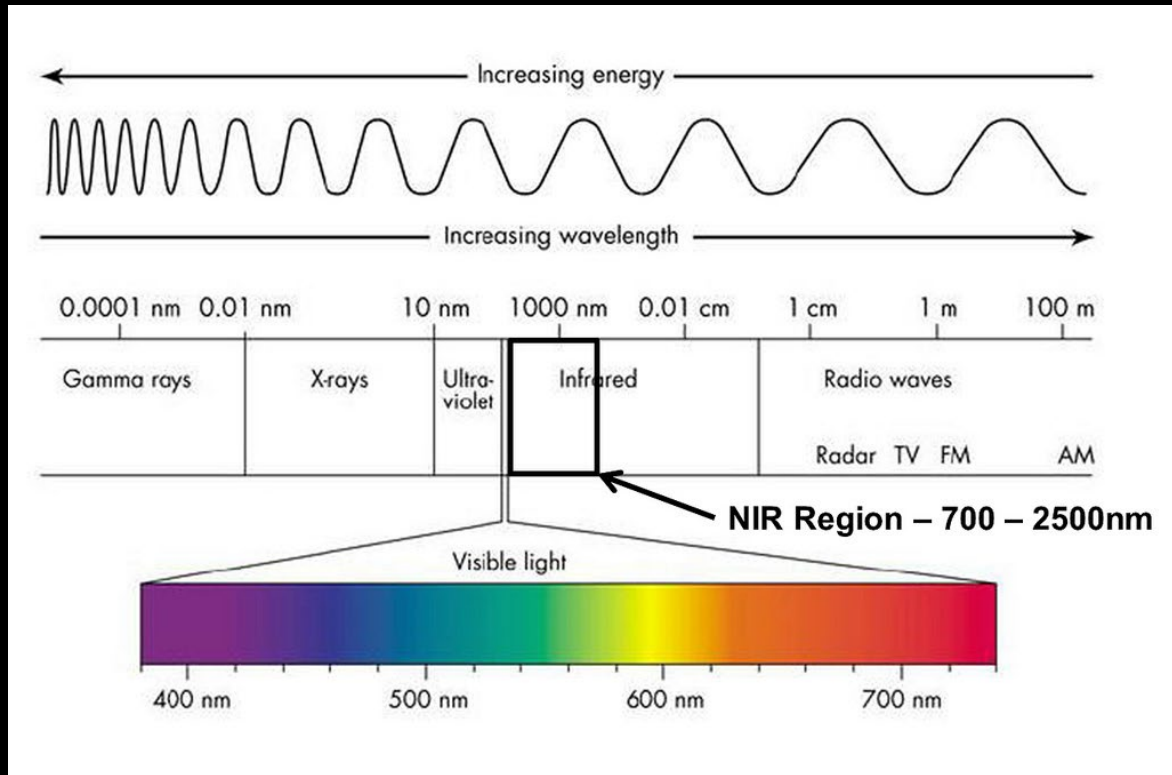
It is **not yet known** if altered neural connectivity can be detected **before** the onset of childhood ADHD



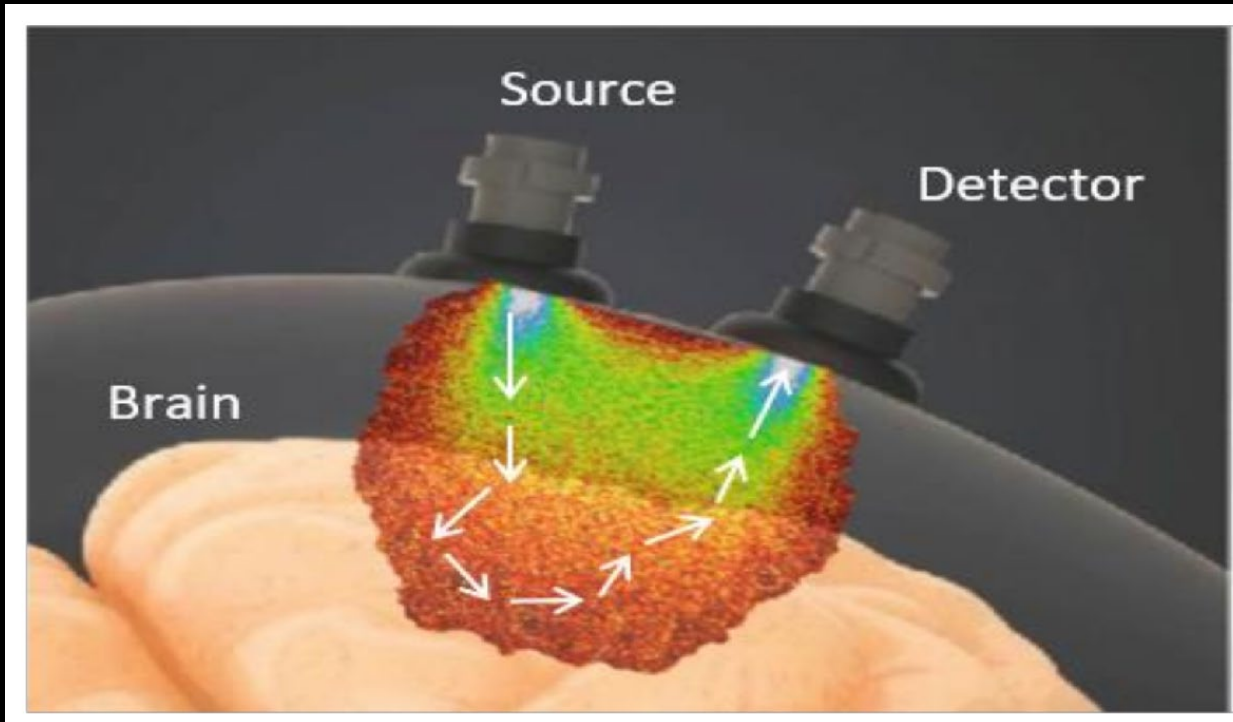


Functional Near-Infrared Spectroscopy (fNIRS)

Near-Infrared Light



Near-Infrared Spectroscopy (NIRS)



- Safe (no radiation)
- Sources emit light and detectors absorb light
- The penetration range of light in the brain is about 3 cm (limited to cortical structures)

BBRF 2021

Young Investigator Award

Examine prefrontal cortex and the dorsal attention network in preschool-aged children with and without familial risk for ADHD

fNIRS during two lab visits at 4 and 5 years old

→ Watching a computer screen with moving images and music (inscapes)



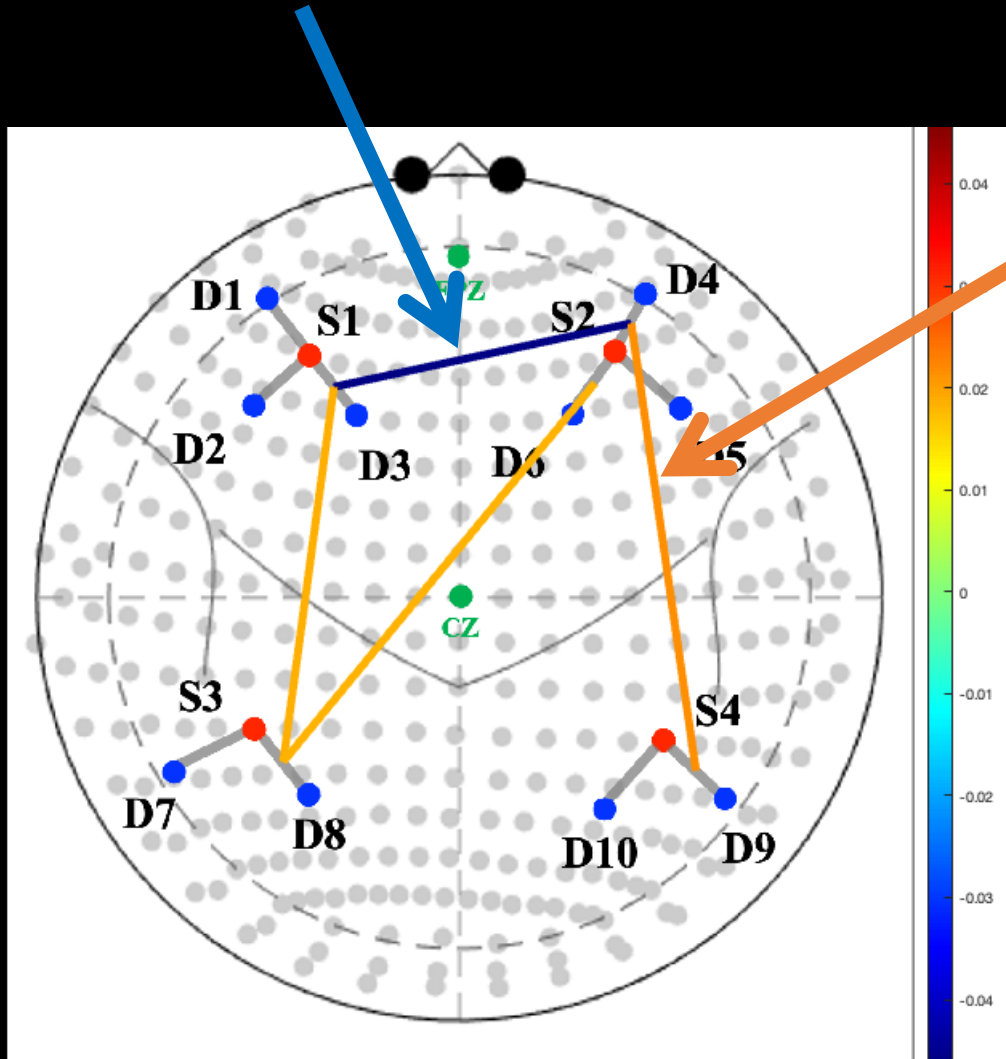
Project Aims

1

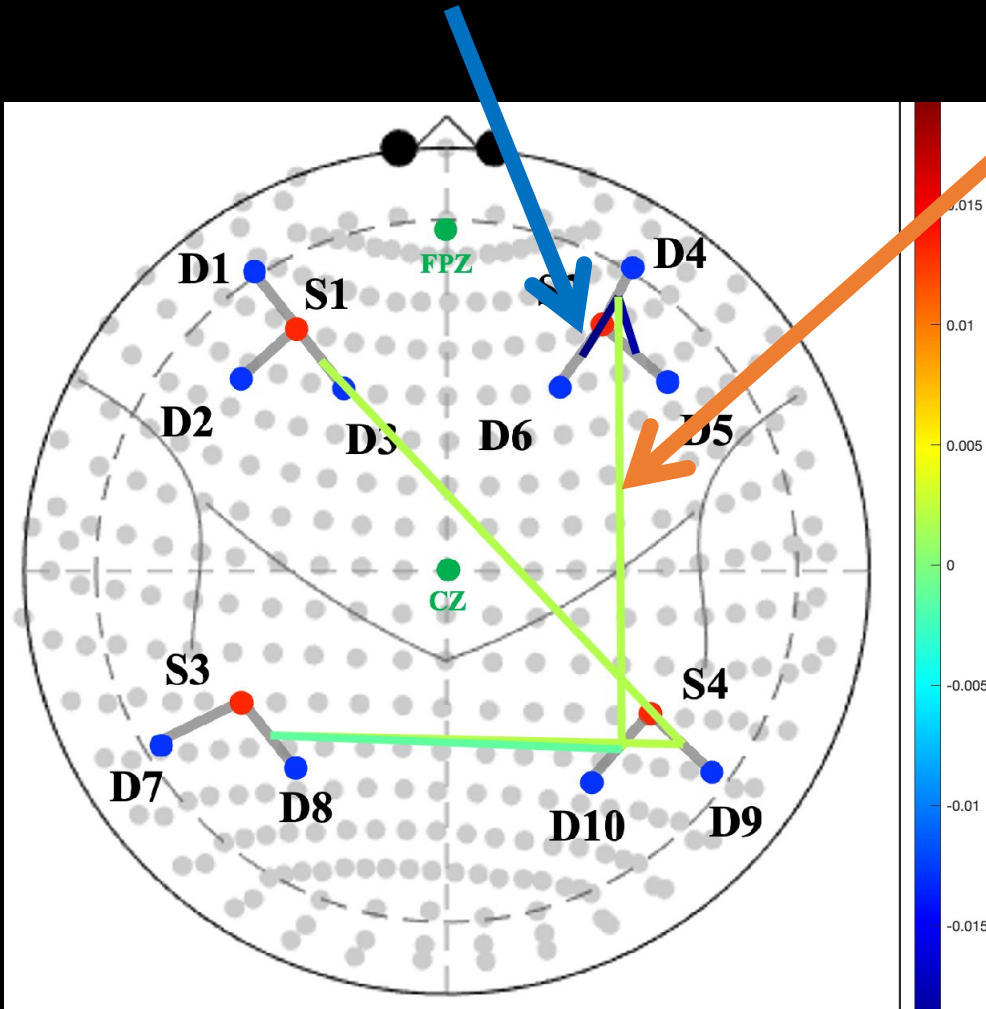
Compare brain activity of children born to parents with vs. without ADHD.

2

Examine brain activity at age 4 as a predictor of ADHD symptoms at age 5.



Offspring of parents with ADHD require greater connectivity between the dlPFC and dorsal attention network to stay engaged.



Greater connectivity between the dlPFC and parietal cortex during an engaging task at age 4 is associated with more ADHD symptoms a year later.



AthENS Study

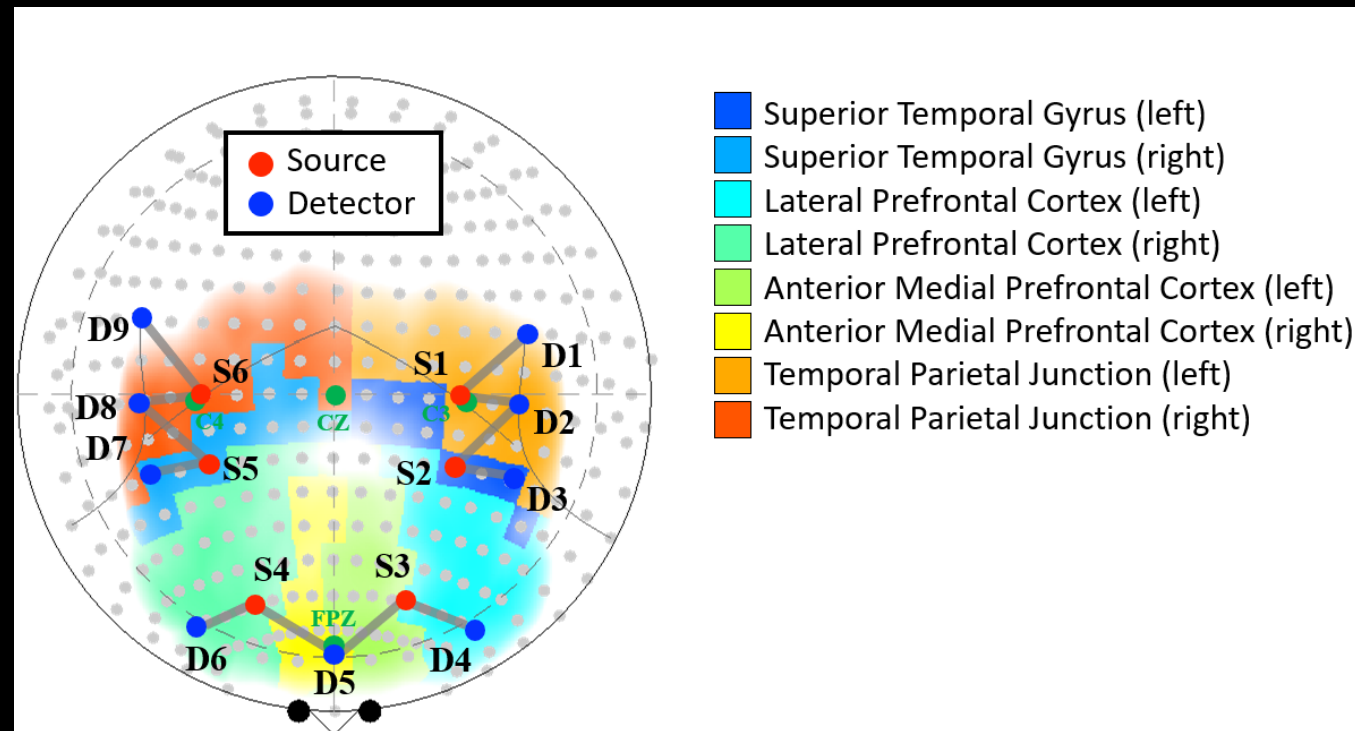
Attention, Hyperactivity, and Early Neurodevelopment Study

NIMH R01MH138380



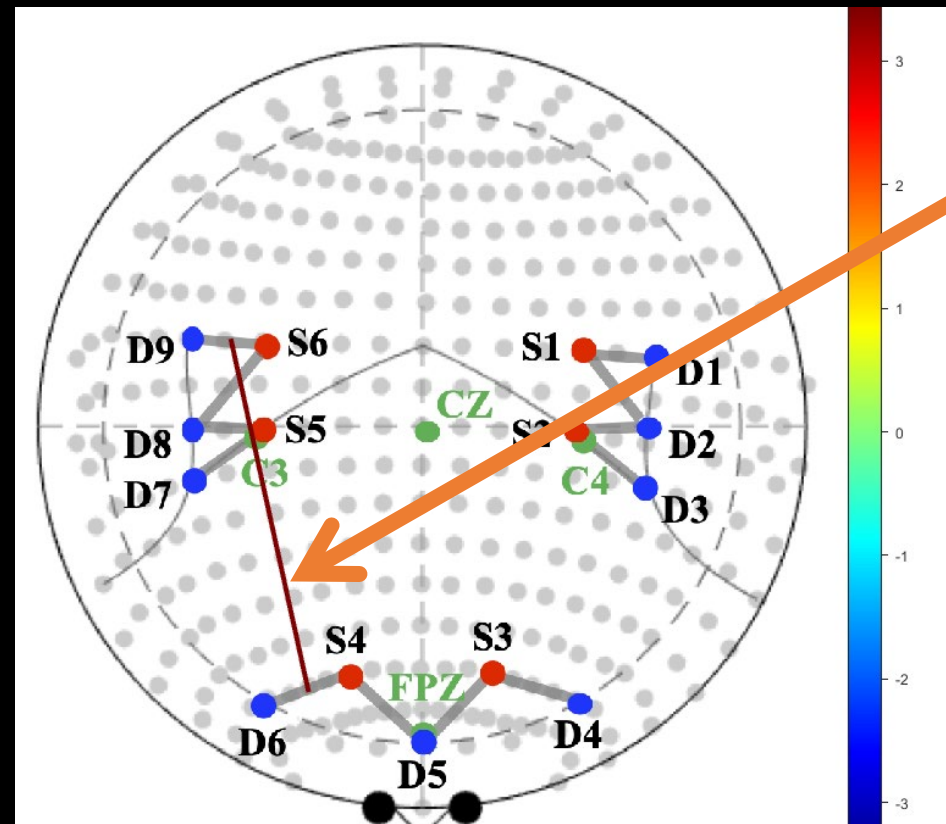
Can we identify even earlier markers of risk before the onset of ADHD symptoms?

We have found similar brain activity among toddlers with greater inattention.



NIMH R01MH113777
Principal Investigator:
Judith K. Morgan, PhD

We have found similar brain activity among toddlers with greater inattention.



NIMH R01MH113777
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A woman with long brown hair is sitting and holding a young child with blonde hair who is crying. The child is wearing a teal shirt. They are in a room with a yellow wall. To the right, there is a television showing a colorful cartoon. In front of the television, on a white surface, are a giraffe toy and a small green dinosaur toy. A black speaker is also visible on the surface. The text "Infant Attention Task" is overlaid in the center of the image.

Infant Attention Task

A photograph of a baby with dark hair and blue eyes, wearing a black t-shirt, crawling on a light-colored wooden floor. The baby is looking towards the camera. In front of the baby are several colorful alphabet blocks. Some blocks are stacked, and others are scattered. The background is a wall with a pattern of white circles and colorful stars. A large black curved shape on the right side of the image contains white and orange text.

PARIS & New(born) PARIS Studies

151 mother-father-infant triads

**Parental ADHD diagnosis as determined
by semi-structured clinical interview:**

71 High Risk for ADHD Infants

At least 1 parent with ADHD

80 Low Risk for ADHD Infants

Neither parent had ADHD



Infant Looking Time



Looking Time

=



Attention

What is Habituation?



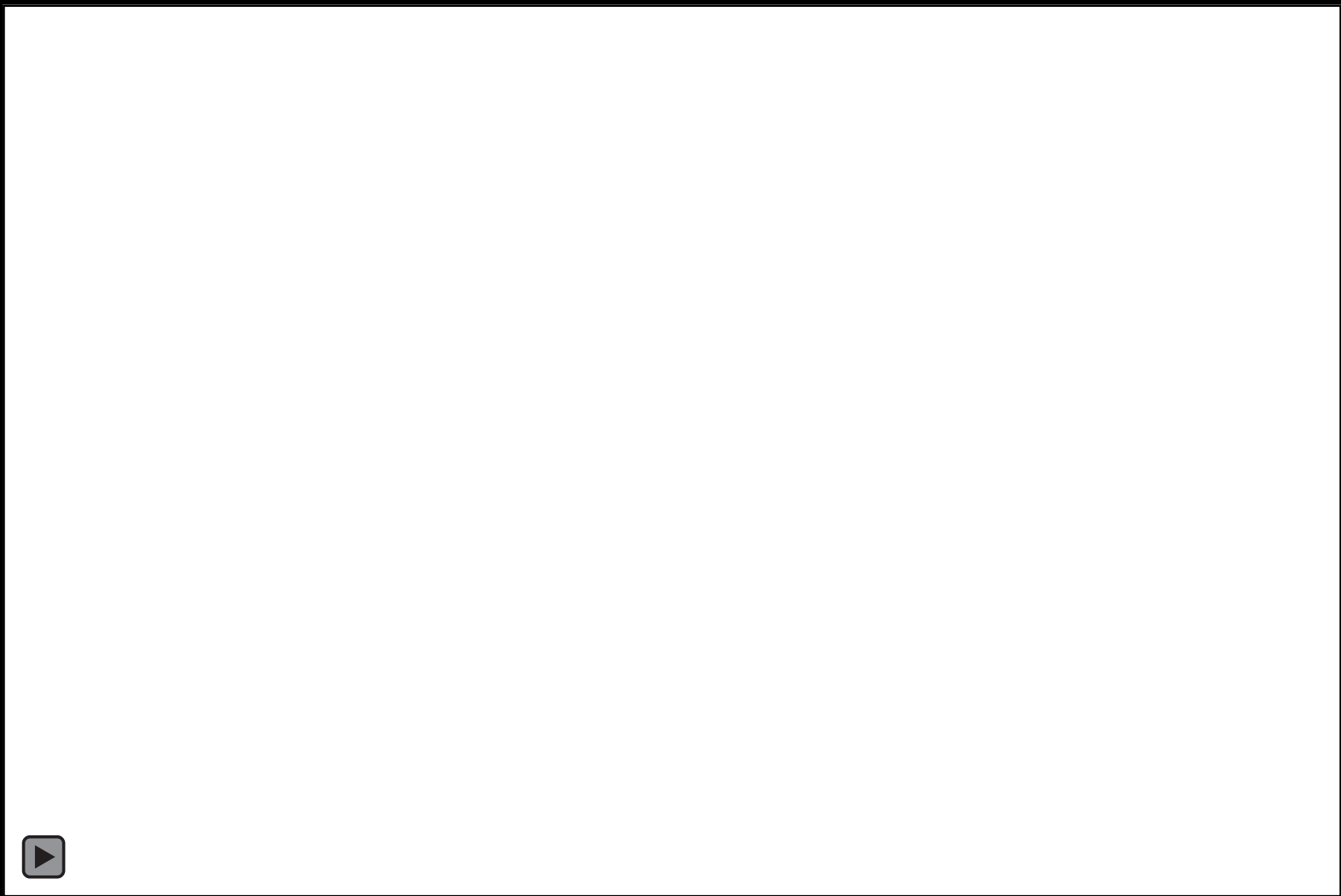
A diminished response to a frequently repeated stimulus

Faster Habituation = Greater Attention

Task Design

Up to 20 test trials (max of 20 seconds each)

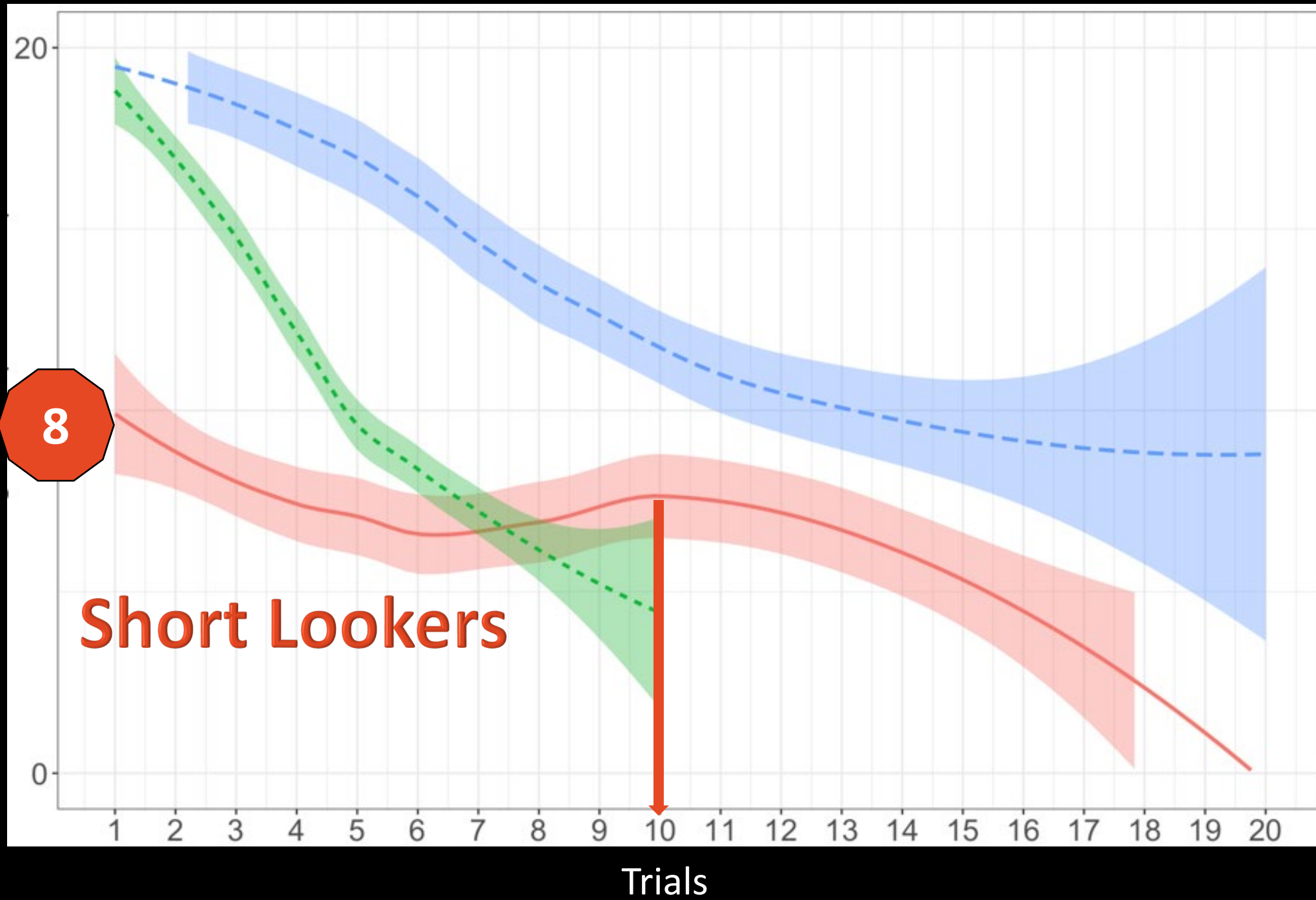
Habituation is achieved when looking time drops to less than $\frac{1}{2}$ of the first three trials averaged



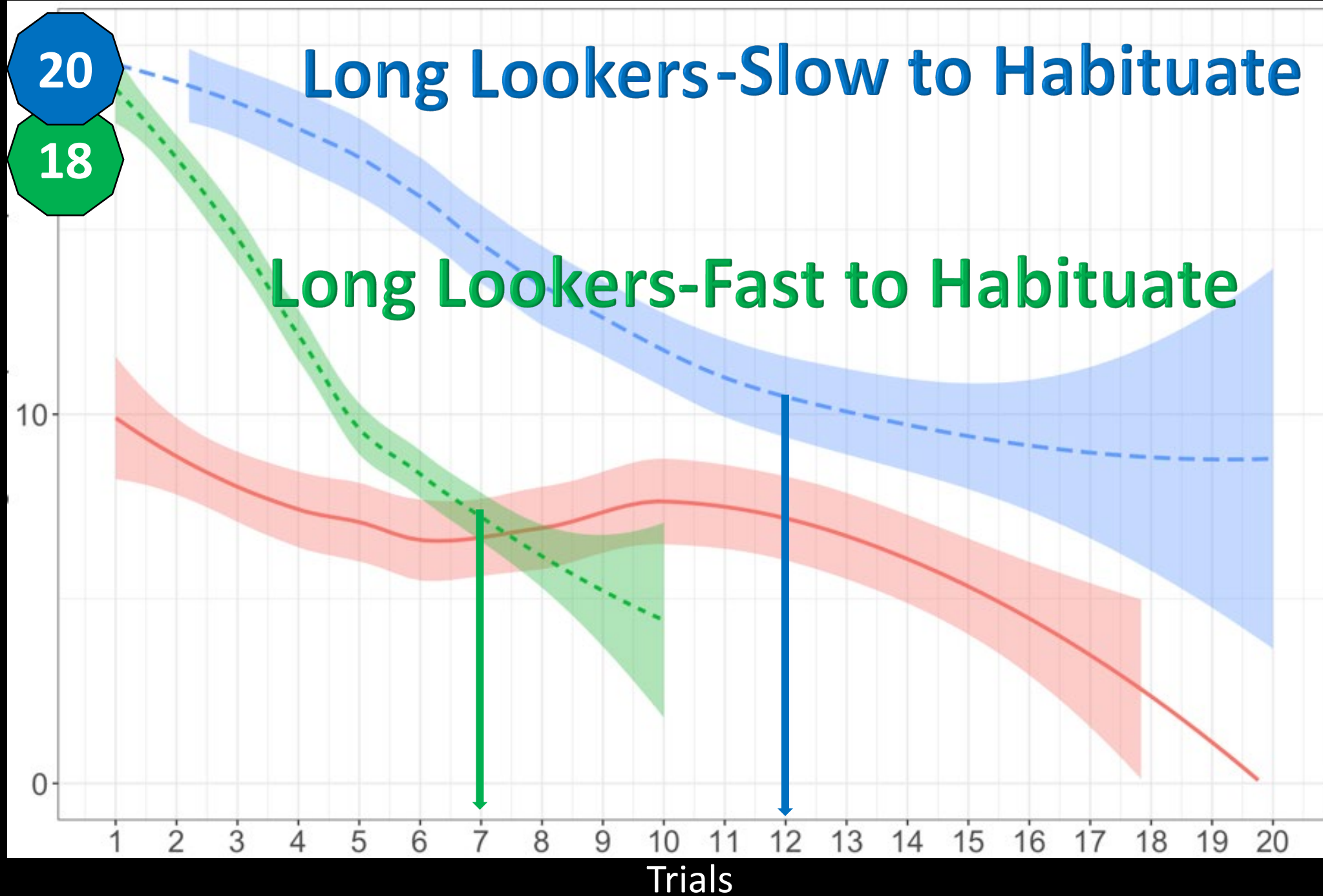
Looking Time in Seconds

8

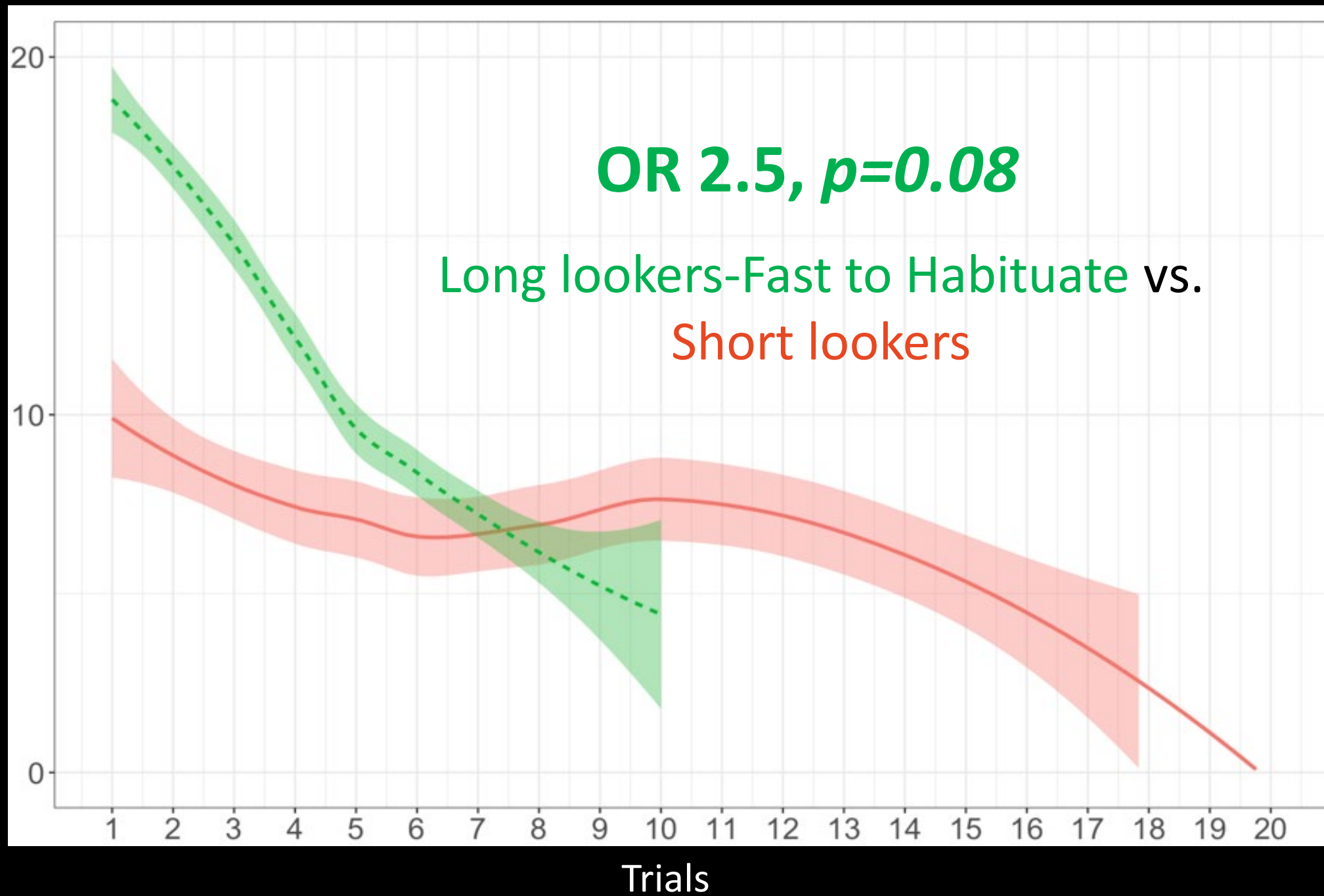
Short Lookers



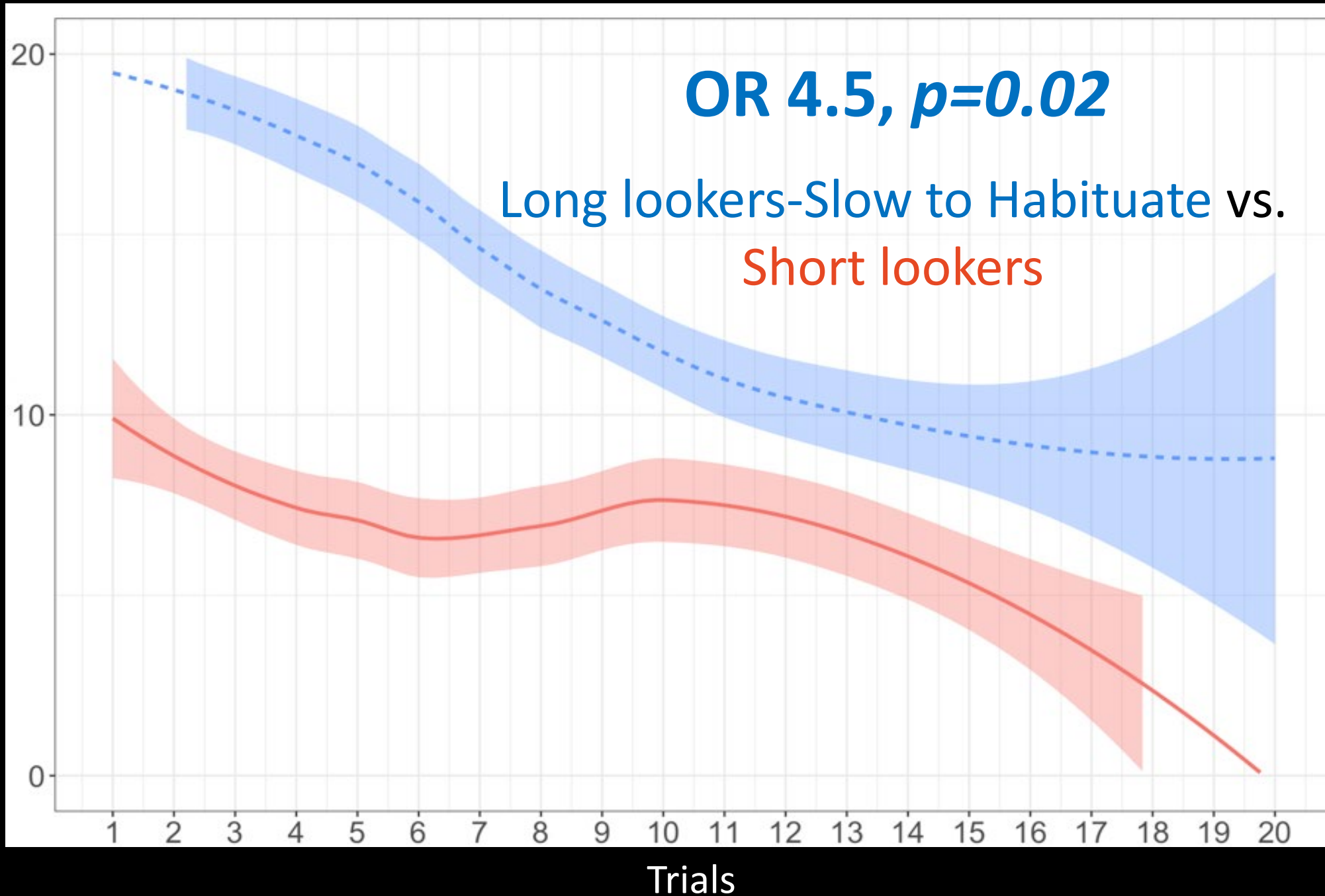
Looking Time in Seconds




Looking Time in Seconds



Looking Time in Seconds





Habituation patterns in infancy
might be an early marker of
risk for childhood ADHD





Next Steps...



A young child with curly hair is blowing bubbles. The child is looking up and to the right, with their mouth open and a small bubble wand held in their hand. Several bubbles are visible in the air around the child's head. The background is a soft, out-of-focus green, suggesting an outdoor setting. The overall mood is playful and joyful.

Longitudinal Follow Up

Infant Task Performance in relation to Brain Connectivity & Development



A group of ice hockey players in black and red uniforms are celebrating a goal on an ice rink. They are huddled together, holding their sticks, with a goal net visible on the left. The word "Teamwork" is overlaid in large yellow text.

Teamwork

Acknowledgments

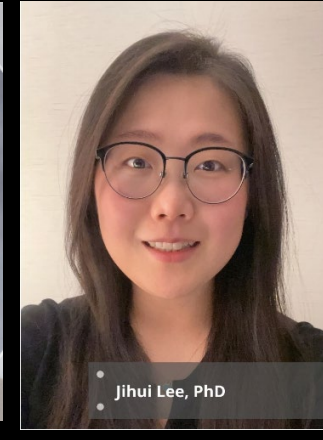


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Questions