

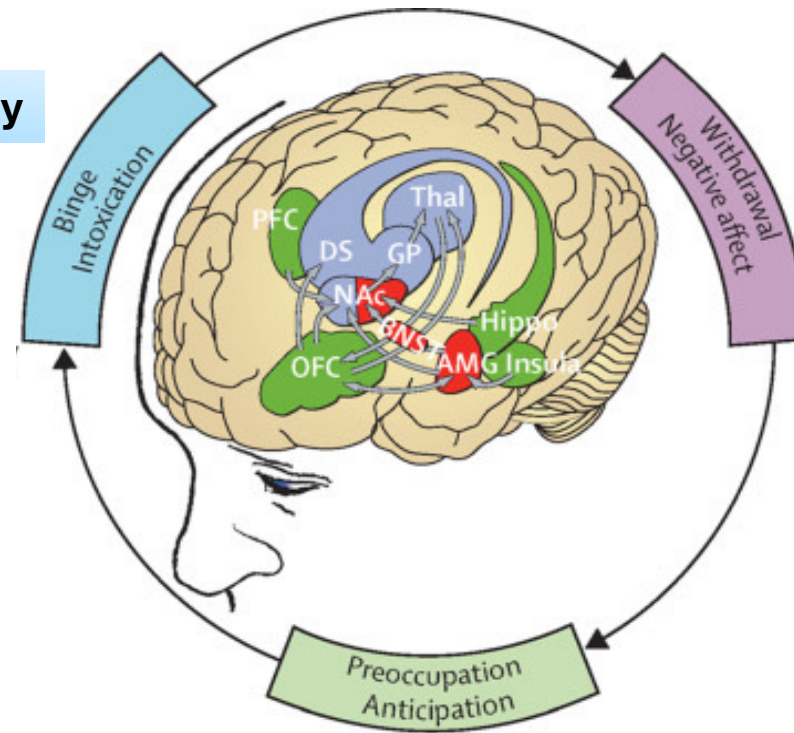
# **Albatros Conférence – ABCD & Évolution des addictions**

**Nora D. Volkow M.D.**  
**Director**

# Neurocircuitry of the Addiction Cycle

Reward Motivation Circuitry

Emotion and Fear Circuitry

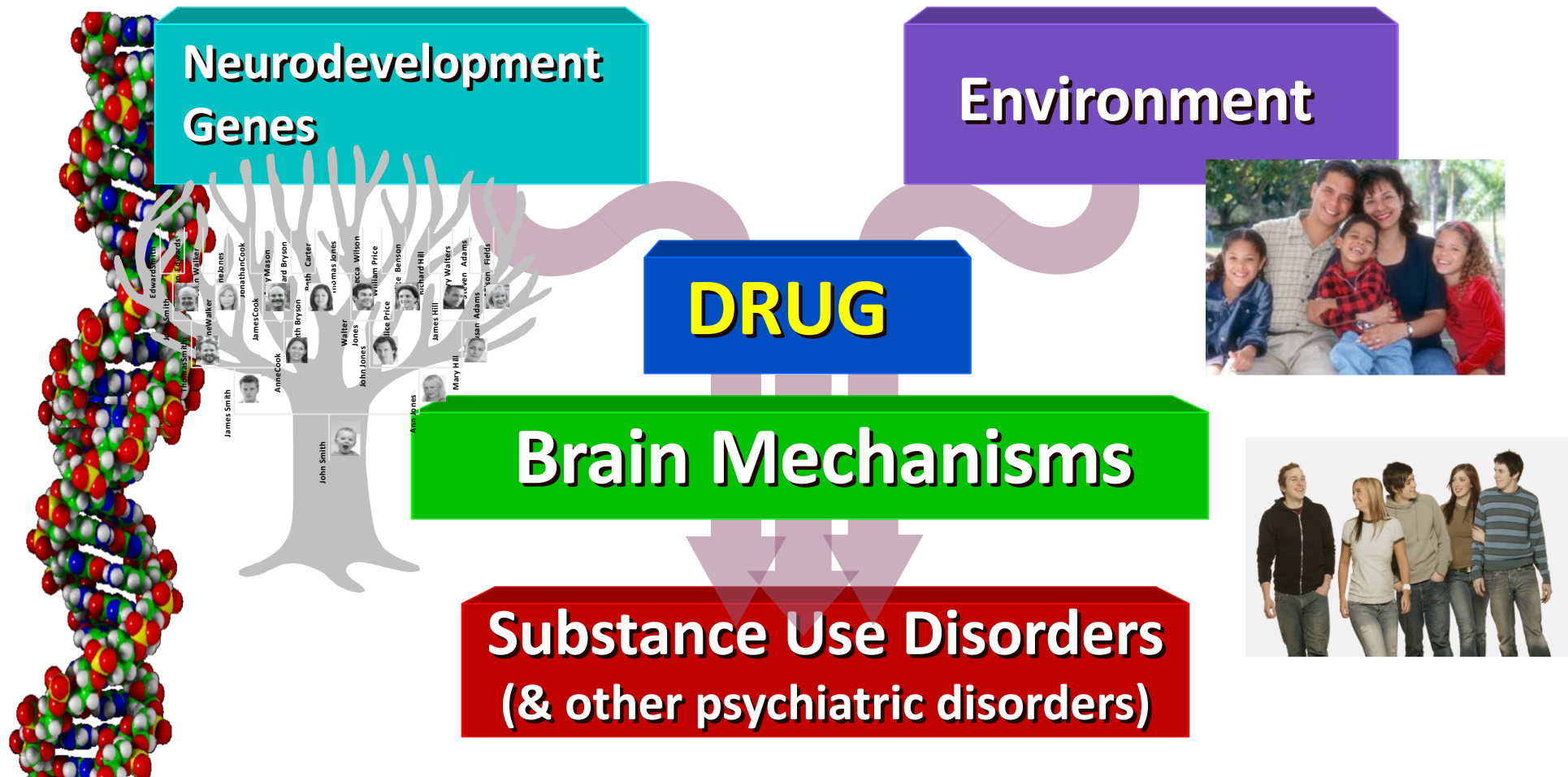


Executive Neurocircuitry

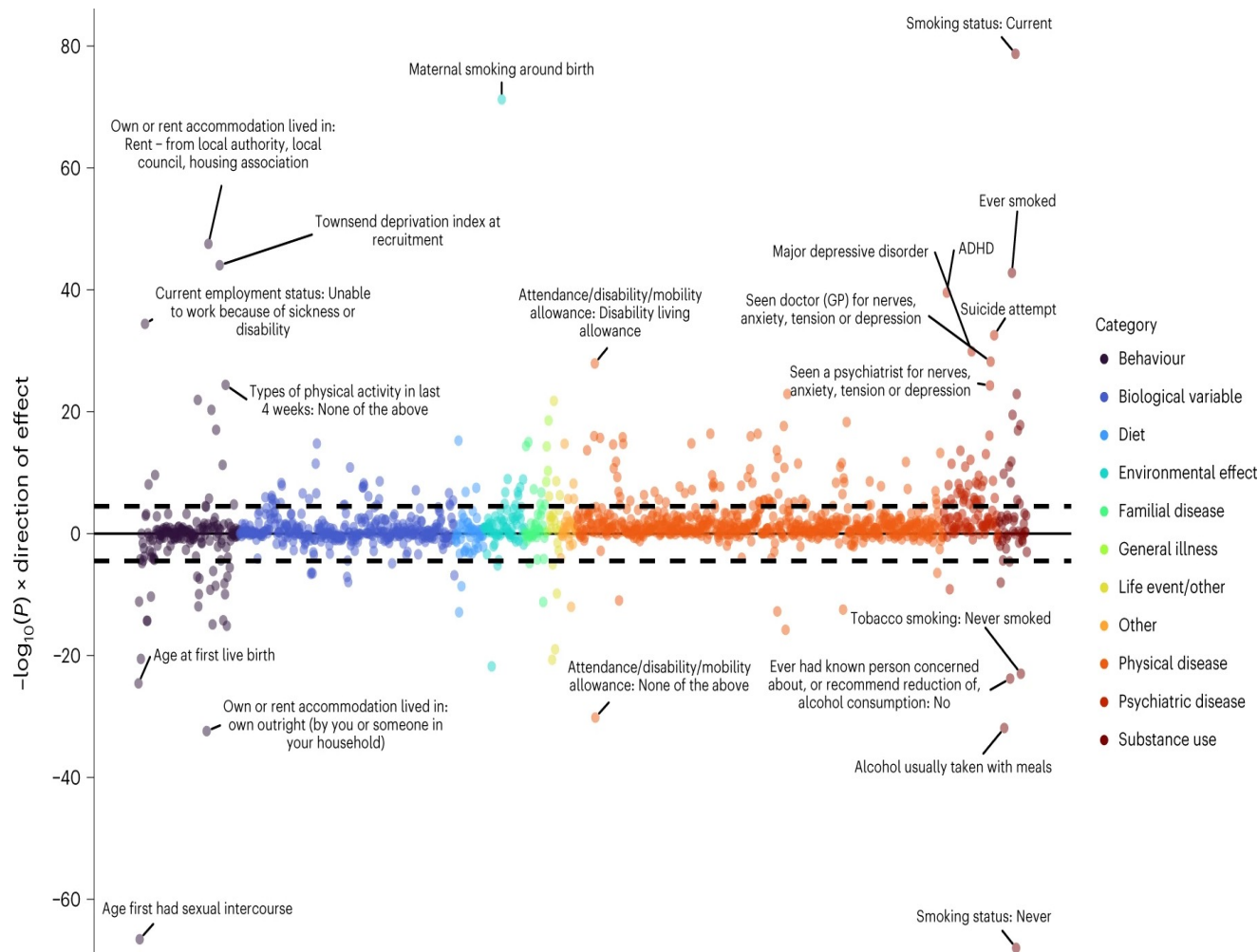
Saliency and Interoceptive Circuitry

[Koob and Volkow, Lancet Psychiatry \(2016\).](#)

# Multiple Shared Factors Influence Substance use Disorders



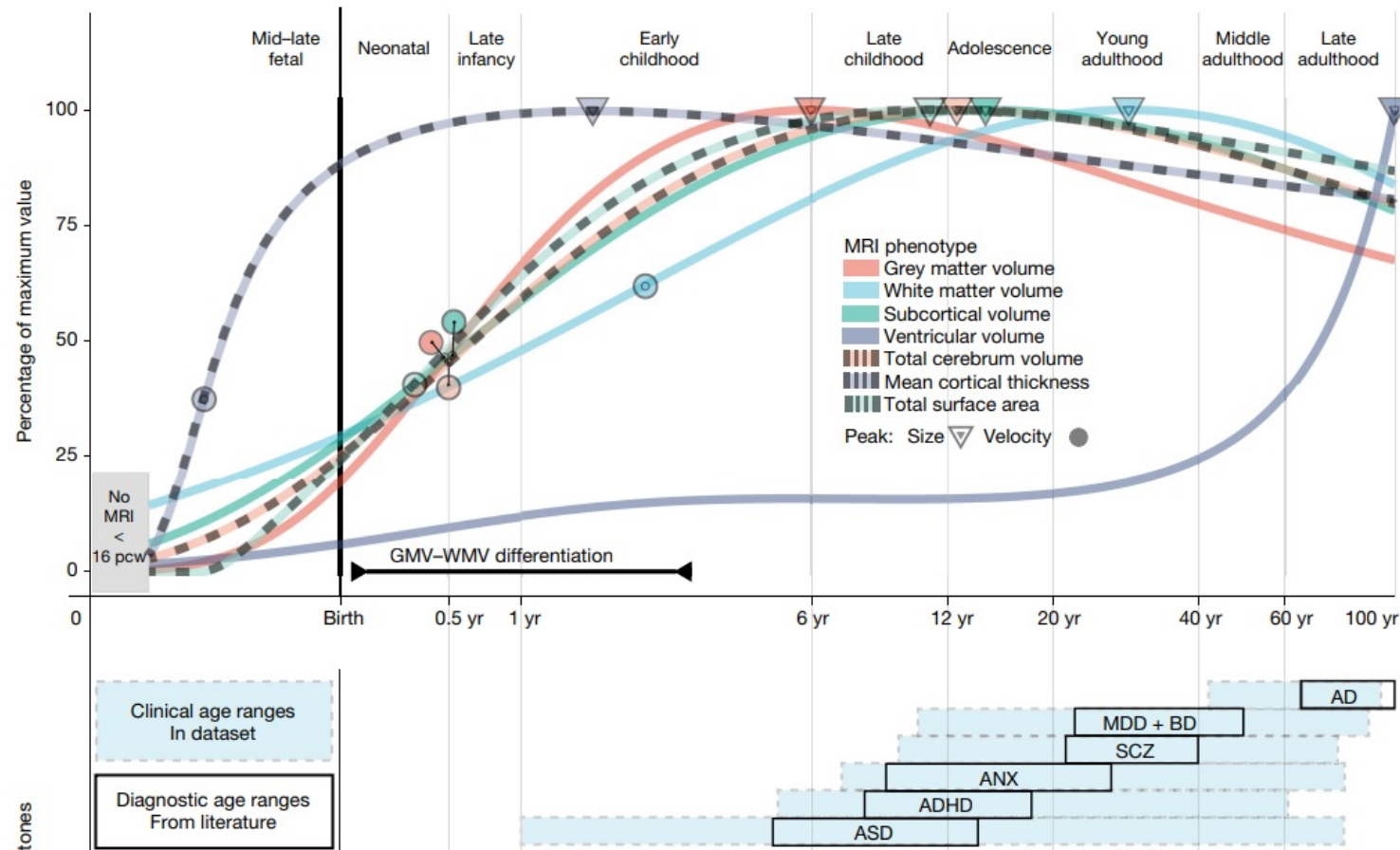
# GENETICS: GWAs >1 Million Subjects Identifies Loci Underlying Multiple SUD and other Psychiatric Phenotypes



A general addiction risk factor identified 42 genes that included **FTO**, **DRD2** and **PDE4B** and that also correlated with several psychiatric associated traits including suicidal behavior, anxiety, depression and externalizing behaviors.



# Neurodevelopmental Milestones

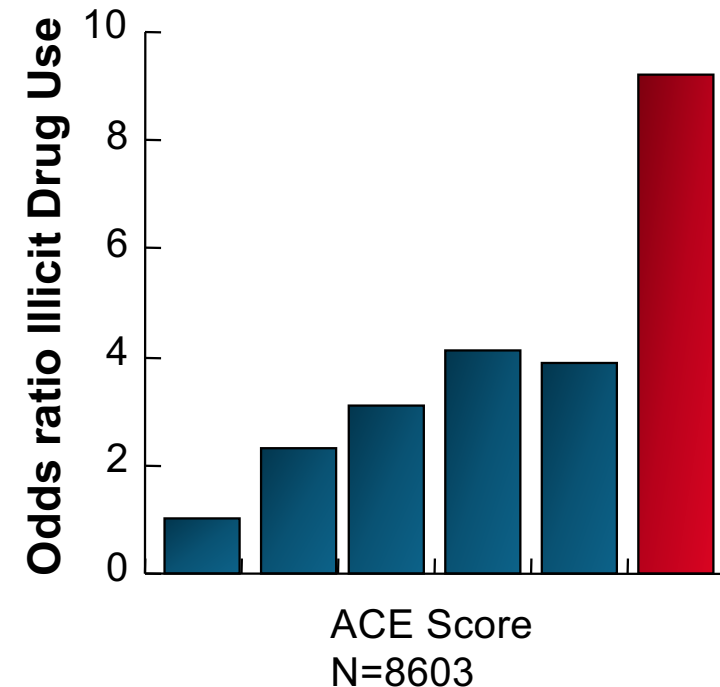
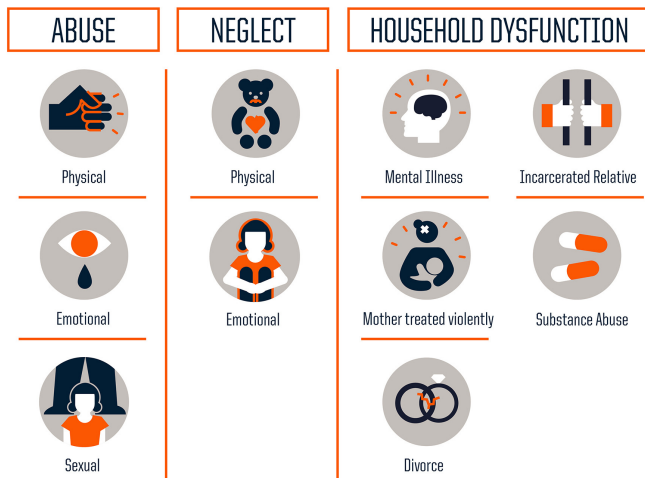


# What Drives Vulnerability for Drug Use?

## Social Determinants of Health



## Adverse Childhood Experiences (ACE)

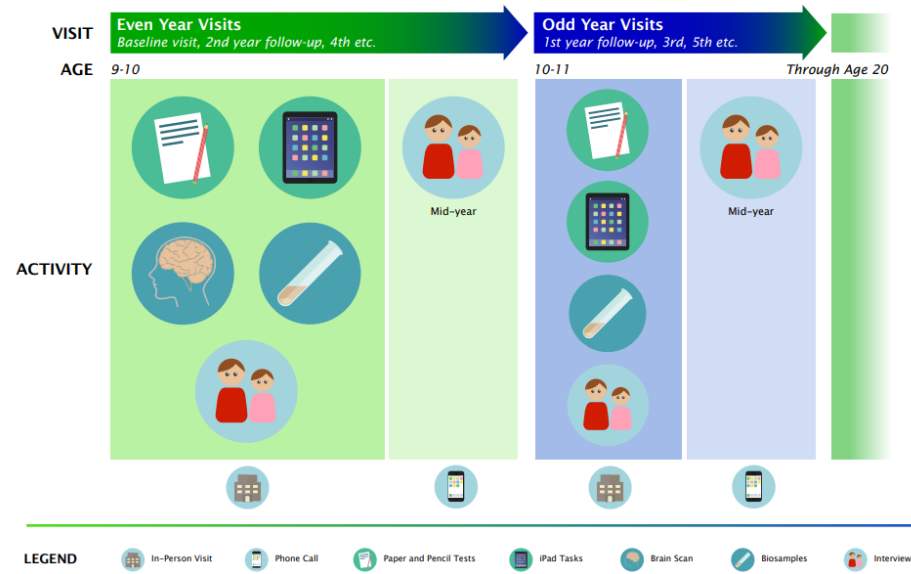


***ACE account for one half to two third of serious problems with drug use***

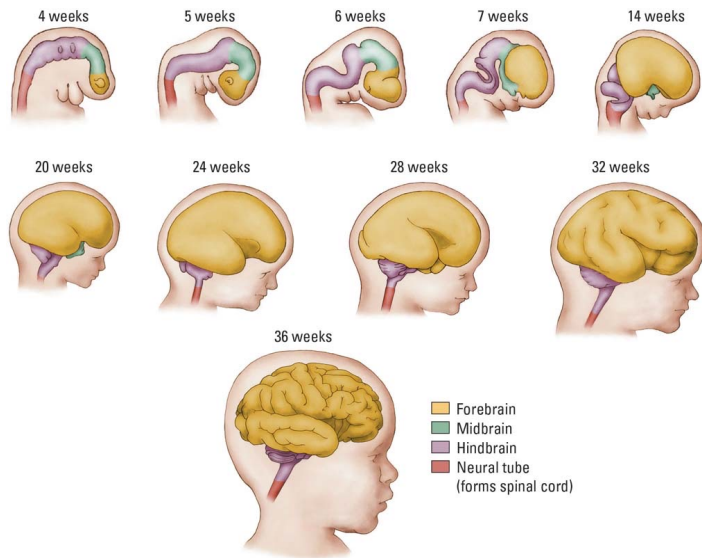
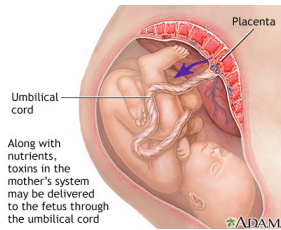


Longitudinal study ~12,000 children, including ~2,000 youth of multiple births, from ages 9-10 into early adulthood to assess factors that influence individual brain development trajectories and functional outcomes.

Approximately 95% of participants retained 8 years into the study



# Prenatal Drug Exposures: Effects on Brain Development

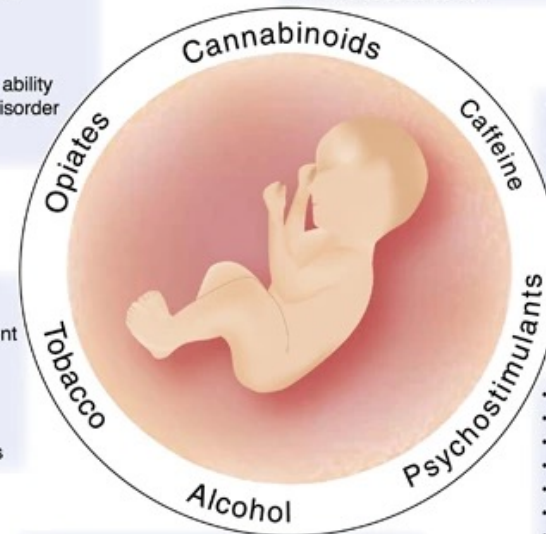


**Figure 8.7**  
Kolb/Whishaw/Teskey, *Intro to Brain & Behavior*, 5e, © 2016 Worth Publishers  
Research from Cowan, W. M. (1979). The development of the brain. *Scientific American*, 241 (3), p. 116.

- Neonatal abstinence syndrome
- Preterm birth and obstetric complications
- Attenuated myelination in infants
- Respiratory insufficiency
- Heart defects
- Reduced growth
- Deficits in cognitive and motor ability
- Attention deficit hyperactivity disorder
- Lower IQ
- Behavioral problems

- Decreased birthweight
- Altered response to stimuli
- Poorer academic achievement
- Poorer cognition
- Attention deficits and hyperactivity
- Adolescent aggression
- Oppositional defiance issues

- Decreased growth
- Deficits in attention
- Increased impulsivity
- Long-term deficits in executive function
- Depression diagnosis
- Future substance use



- Increased risk of growth restriction and prematurity (at high levels)
- Possible decrease in executive function at school age

- Preterm labor
- Short- and long-term growth deficits
- Cardiac and cardiovascular anomalies
- Cranial and brain abnormalities
- Behavior problems
- Emotional and social effects
- Deficits in attention, memory and motivation
- Anxious/depressed behaviors and symptoms
- Aggression and delinquent behavior

- Prematurity and spontaneous abortion
- Limb and facial development
- Reduced growth
- Cognitive delays and impairments
- Reduced brain volumes
- Abnormalities in the corpus callosum
- Deficits in attention, memory, verbal fluency, executive functioning, reaction times, and motor learning

# Prenatal Cannabis Effects on Developmental Trajectories Cognition and Brain Volumes

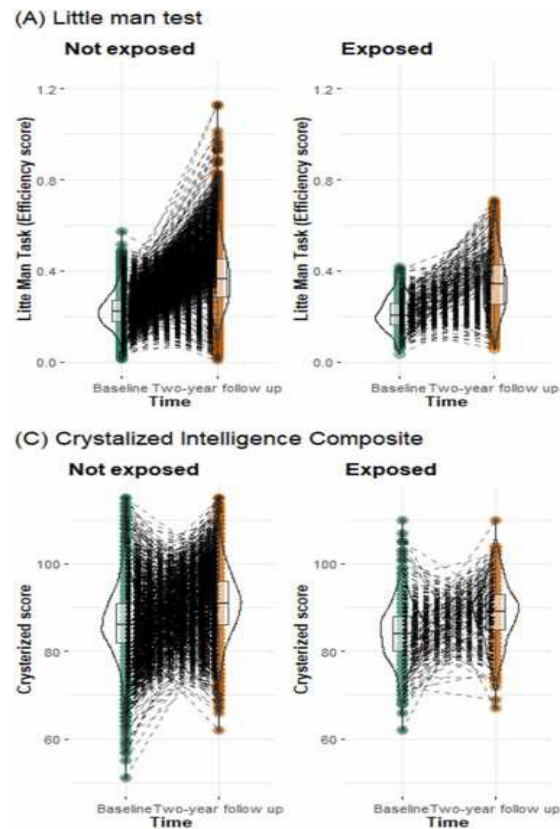
2002-2016



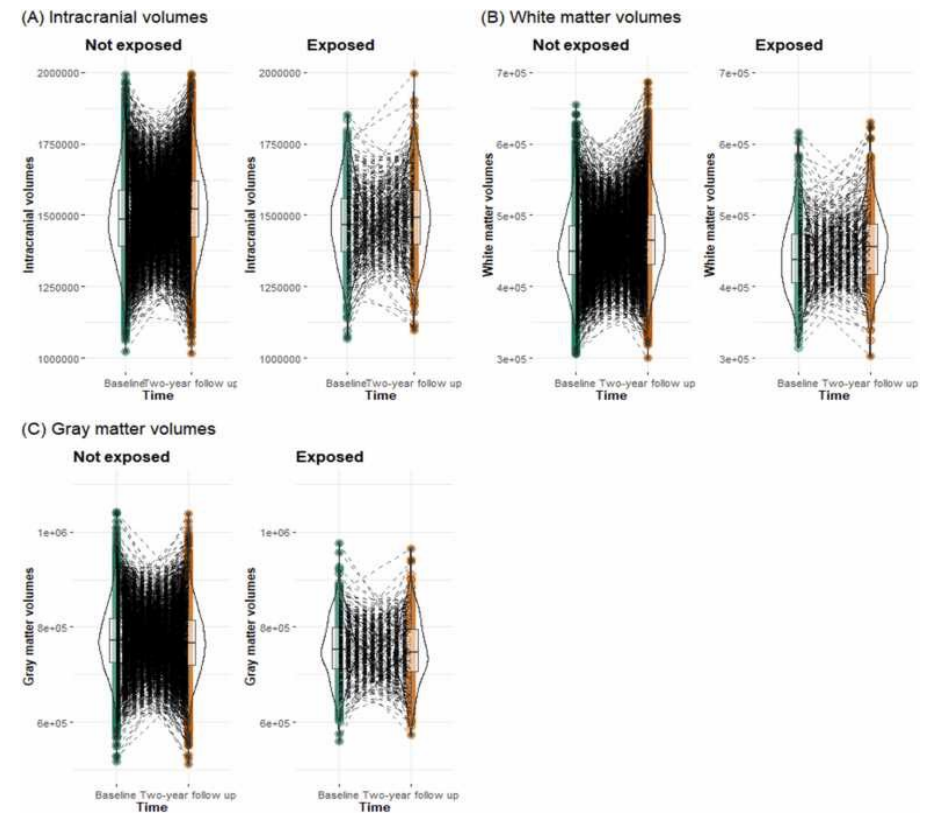
Increase in past-month  
Cannabis use in pregnant  
mothers in US

Agrawal et al., 2018

## Cognitive Abilities



## Brain Volumes

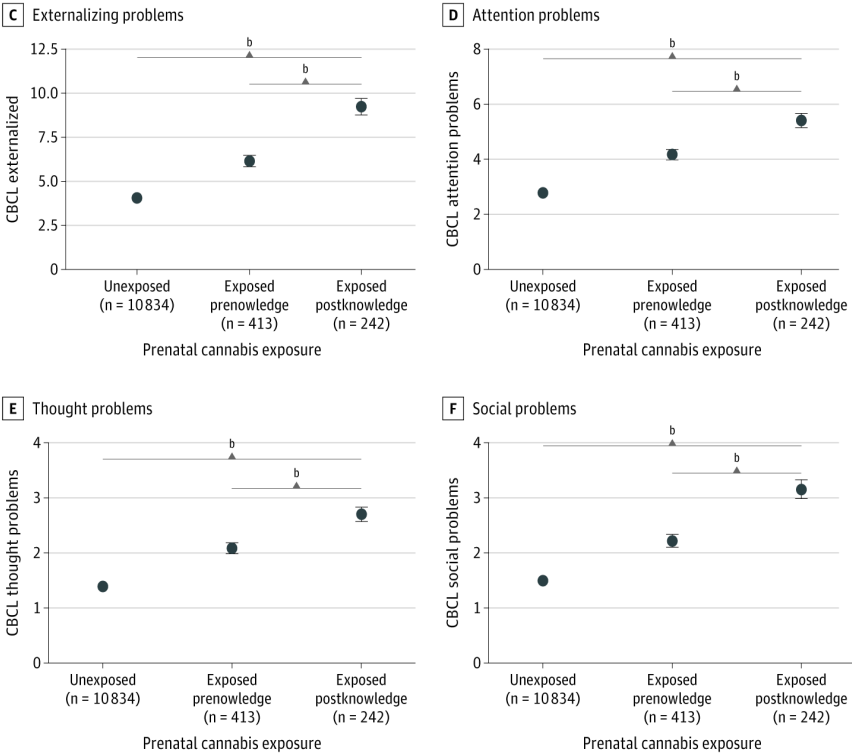


Hiraoka et al., Dev Cogn Neurosci. 2023



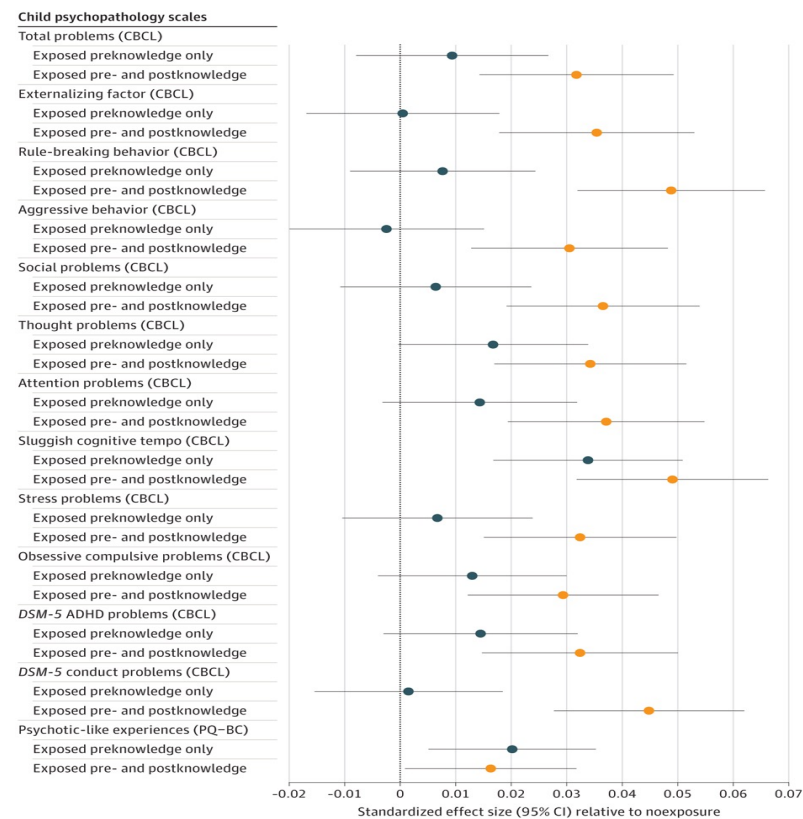
# CANNABIS Prenatal exposure after maternal knowledge and mental health problems

Baseline (9-10 years old)



Paul et al. JAMA Psychiatry 2021

Two years follow up (11-12)

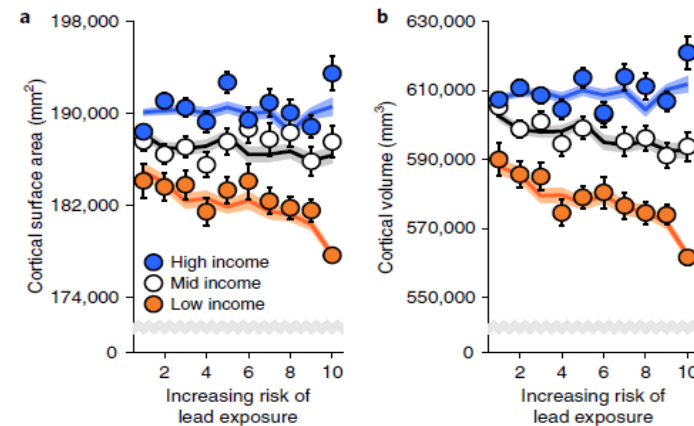
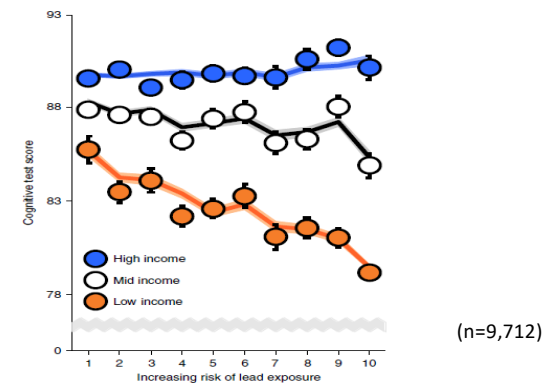
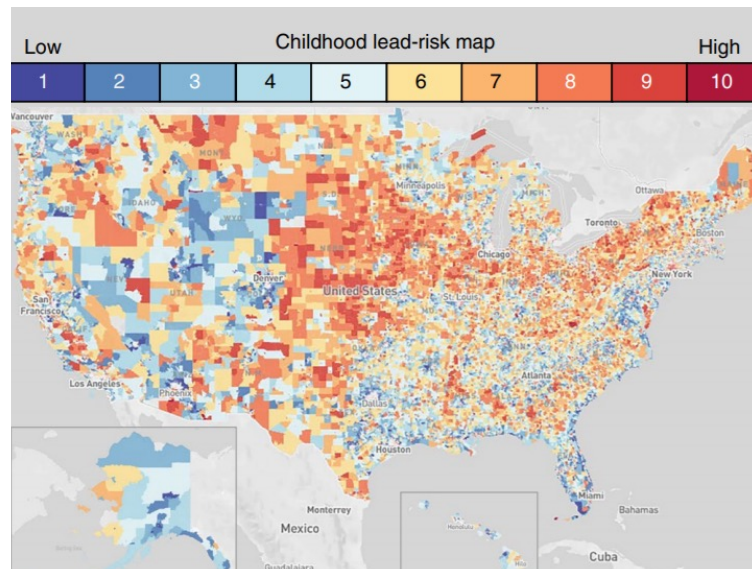


Baranger et al., JAMA Pediatr 2022

# **Social Determinants of Health: Effects on Brain Development**



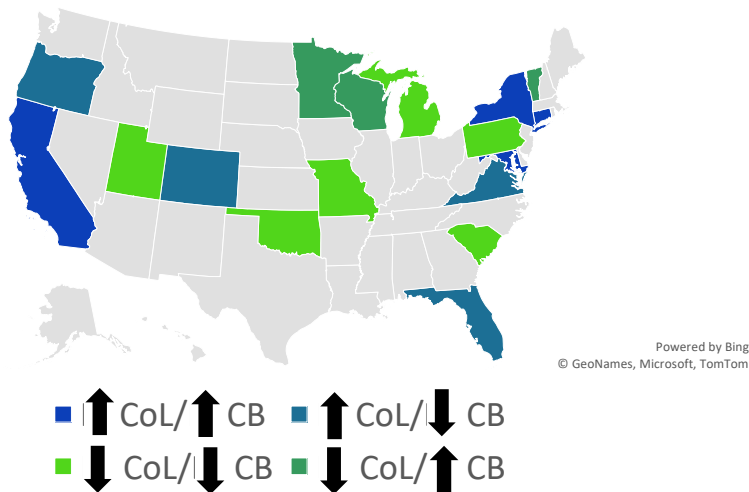
# Lead Risk, SES, Cognition, and Brain



Marshall AT et al. (2020) Nature Medicine 26: 91–97. <https://www.nature.com/articles/s41591-019-0713-y>

# Antipoverty Programs Associated with Reduced Disparities in Brain Development and Mental Health

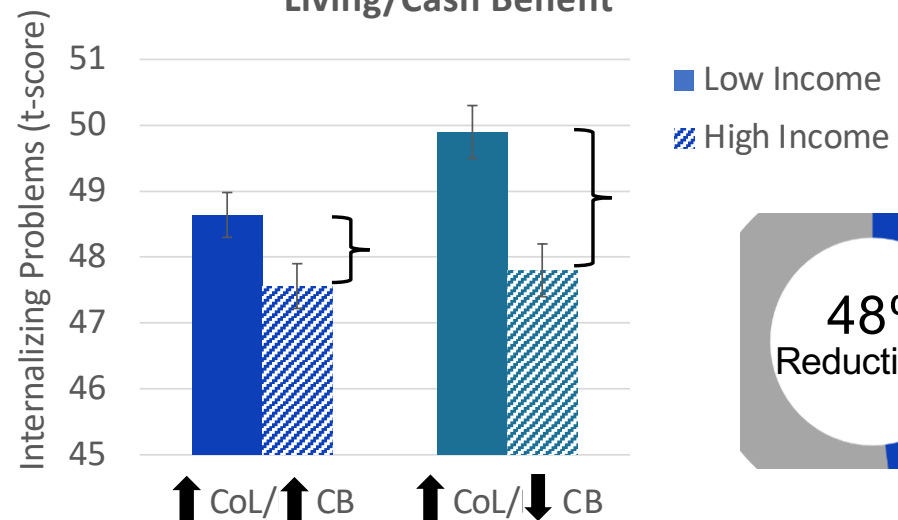
Cost of Living (CoL), Average Cash Benefit(CB)  
by State



Controlled for:

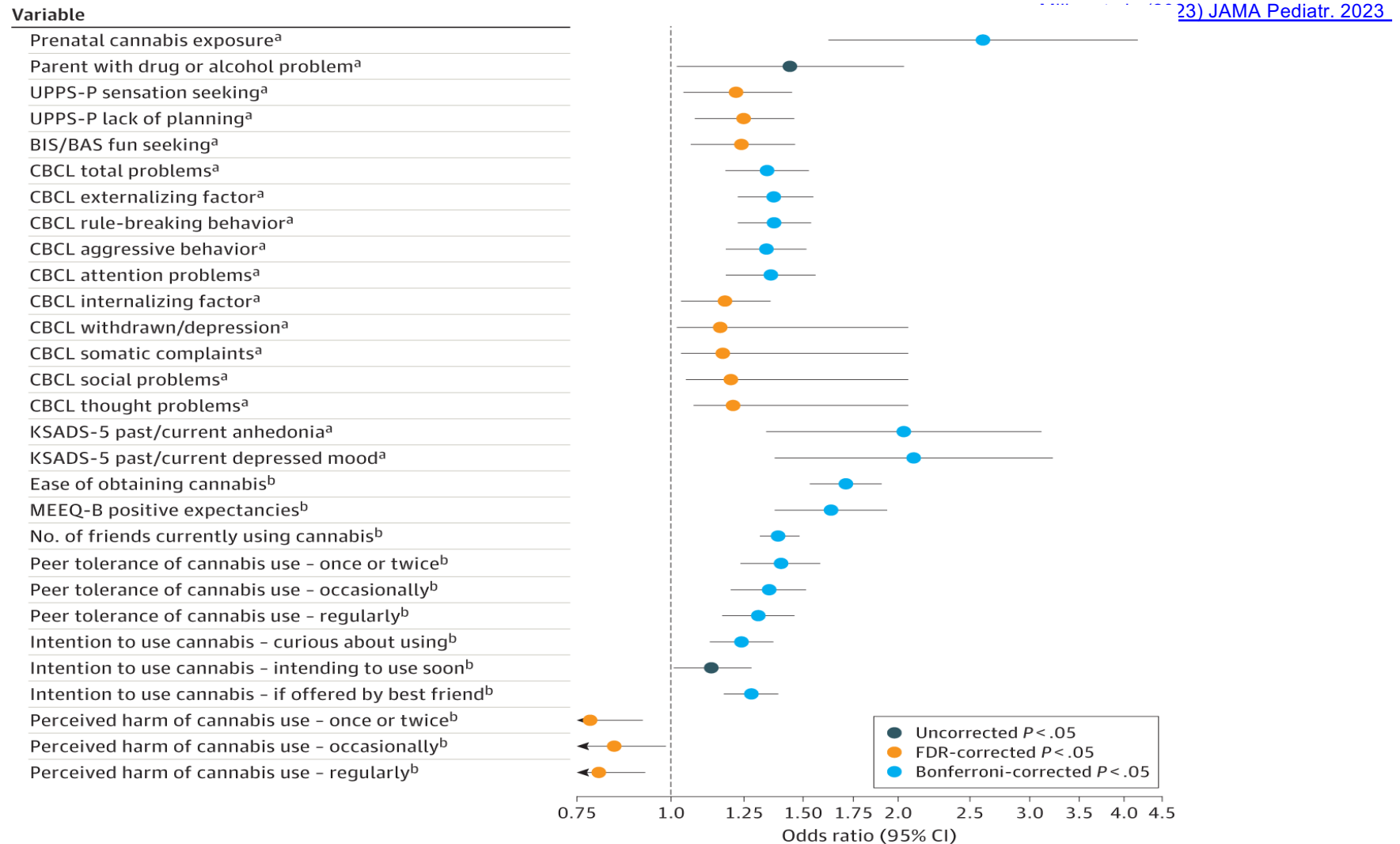
- Population density
- State economic conditions (economic inequality, unemployment rate),
- Non-economic characteristics social and political factors (political preferences, women's political participation, reproductive rights, incarceration rate, tightness/looseness—i.e., cultural differences around rule and norm adherence),
- Education system equity (state-funded preschool enrollment, reading proficiency among students from low-income backgrounds).

Internalizing Problems by Income/Cost of  
Living/Cash Benefit



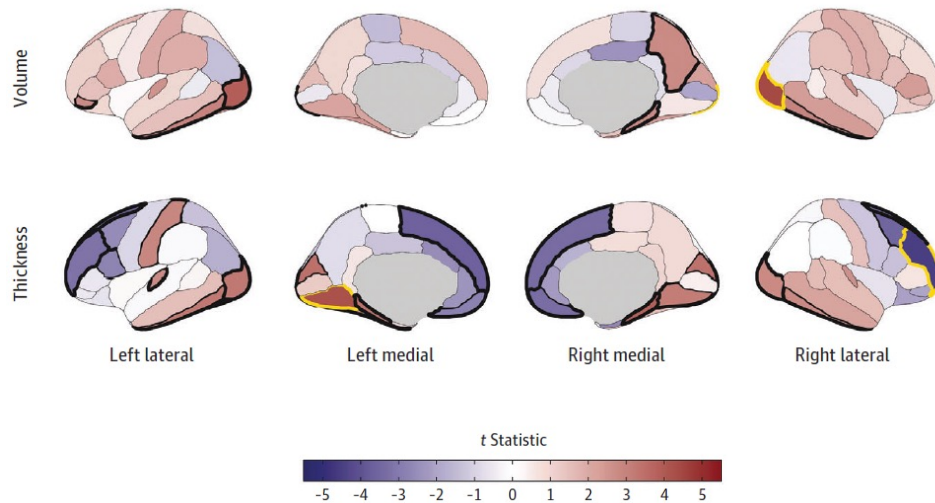
[Weissman, et al. Nature Communications. \(2023\).](#)

# Variables associated with cannabis use in early adolescence

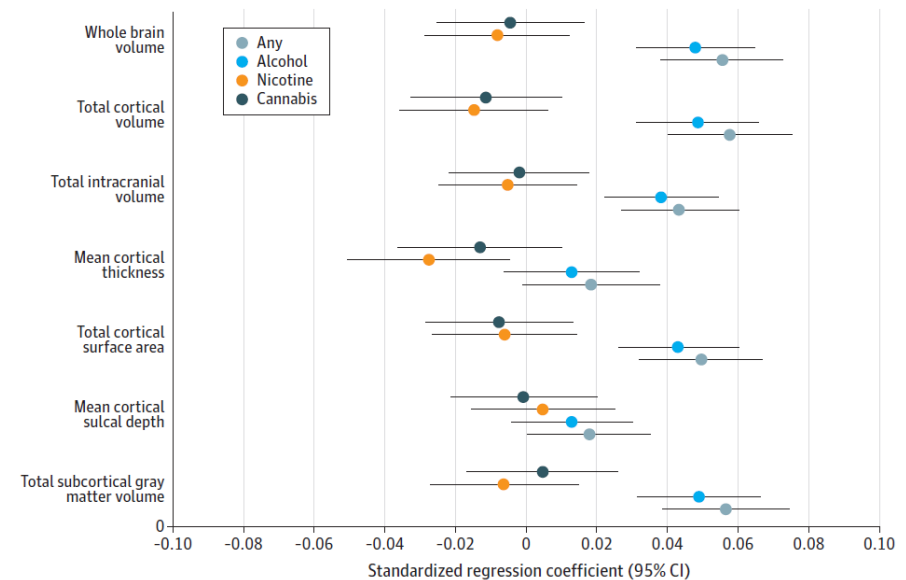


# Neuroanatomical Variability and Substance Use Initiation in Late Childhood and Early Adolescence

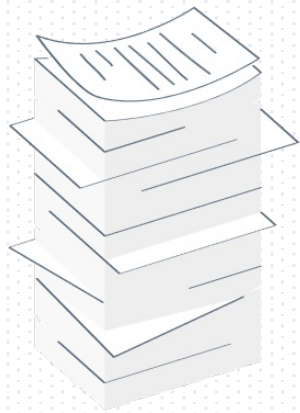
Regional Cortical Volume and Thickness Associations With Early Substance Use



Standardized regression coefficients for associations between global metrics and substance use initiation.



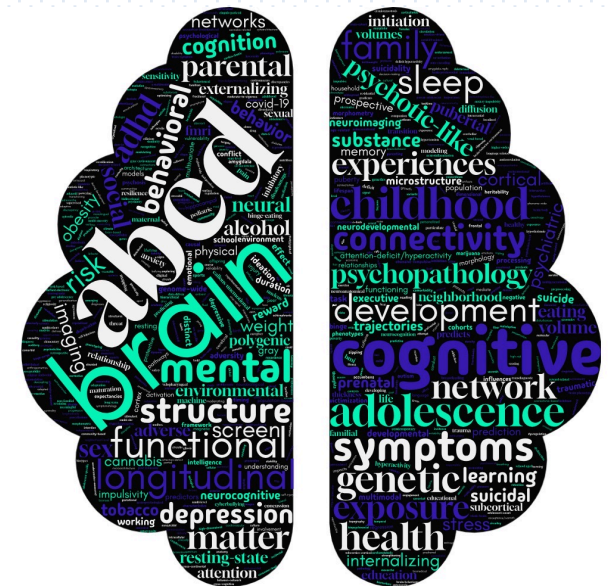
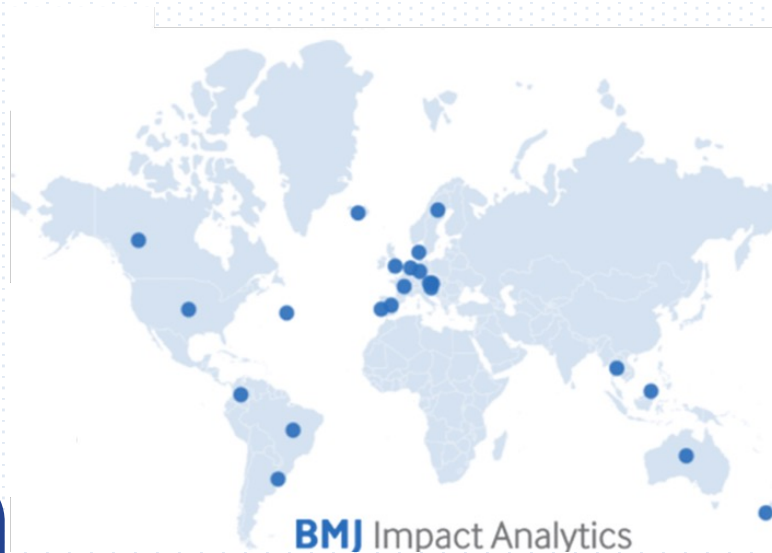
Preexisting neuroanatomical variability was associated with substance use initiation and may reflect a predispositional risk for initiating substance use earlier in life with potential cascading implications for development of later problems



**1300+**  
scientific  
publications

Cited in **83**  
clinical or other  
guidelines

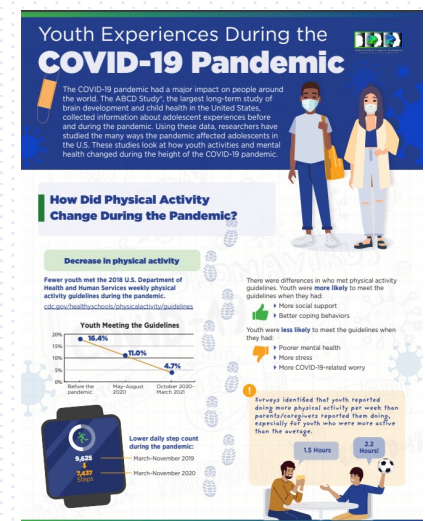
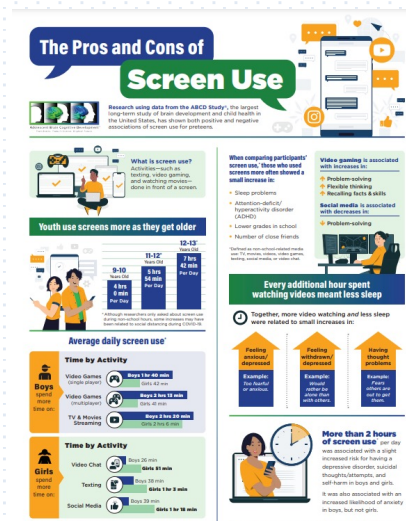
# ABCD Impact



ABCD  
publications  
available at:



Infographics and  
webinar recordings  
available at:

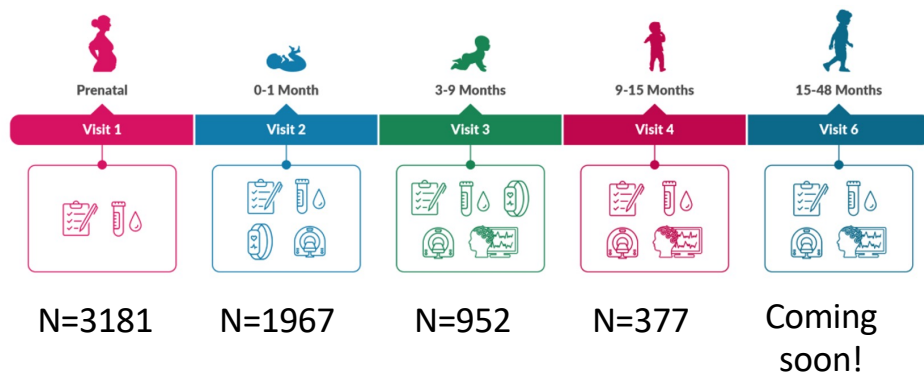




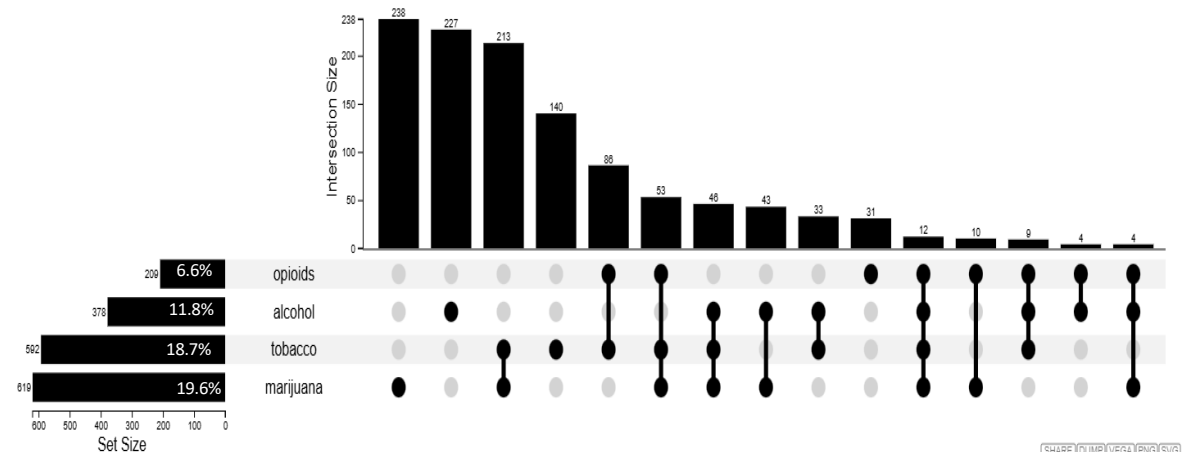
# HEALthy Brain and Child Development Study (HBCD)

Longitudinal study of >7000 children, tracking brain and behavioral development from **prenatal stages through 9-10 years of age**. It aims to understand how biological, environmental, and social factors—including substance exposure—impact child development.

Enrollment began July 2023  
3282 participants enrolled (104% of target)  
36.3% reported prenatal substance use  
1<sup>st</sup> data release Spring 2025 via NBDC platform  
Full sample enrolled by 2027



## Prenatal Substance Use: 36.3% (N=1149/3163)



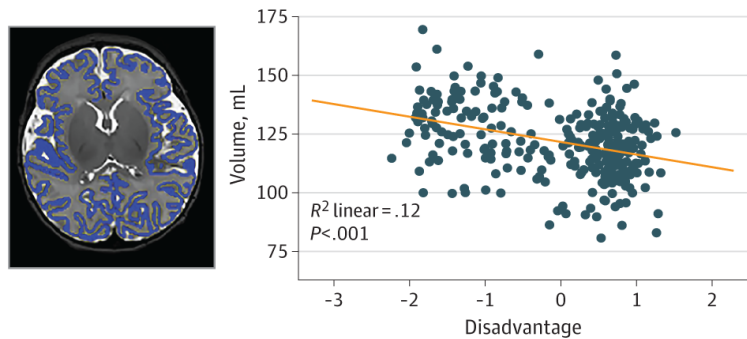
## Prenatal Substance Use Criteria

**Opioids:** ≥Weekly for ≥2 weeks  
**Tobacco/Nicotine:** ≥Weekly for ≥4 weeks  
**Marijuana/Cannabis:** ≥Weekly for ≥4 weeks  
**Alcohol:** ≥7 standard drinks/week for ≥2 weeks; or ≥3 standard drinks/ occasion on ≥2 occasions

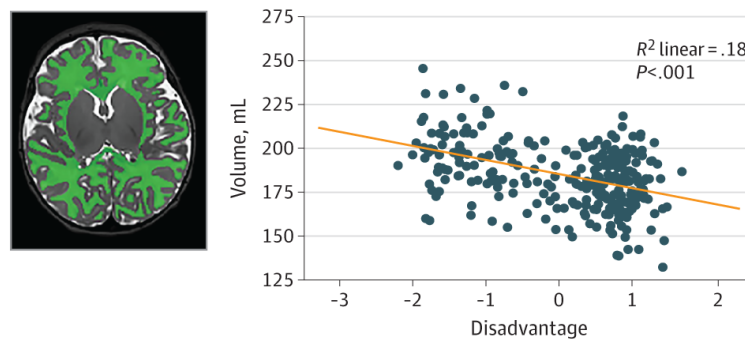
# Association of Prenatal Exposure to Early-Life Adversity With Neonatal Brain Volumes at One Year of Age

Correlation Between Total Brain Volume and Maternal Social Disadvantage Factor

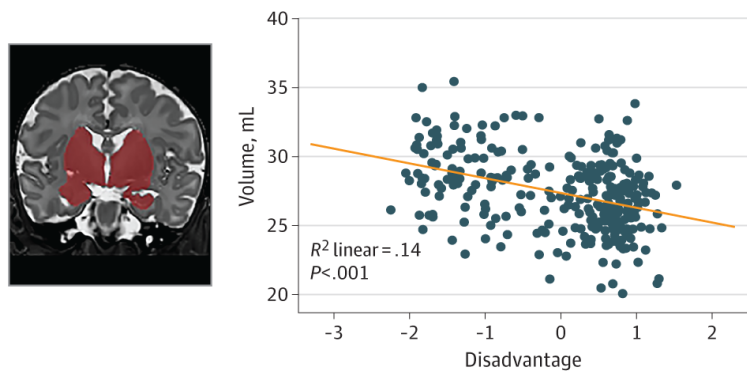
**A** Cortical gray matter



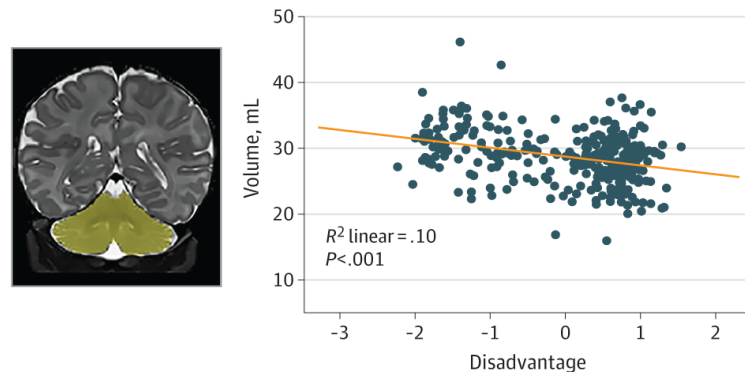
**B** White matter



**C** Subcortical gray matter

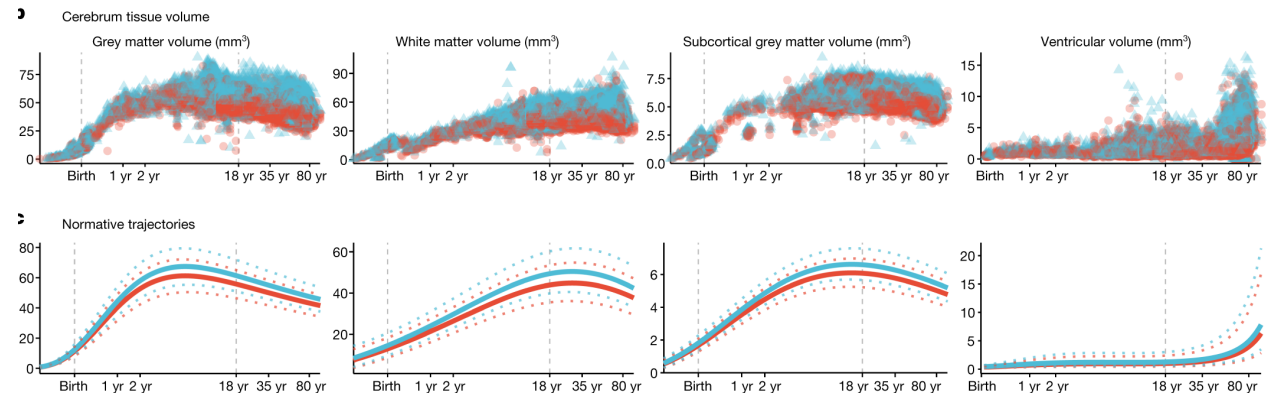
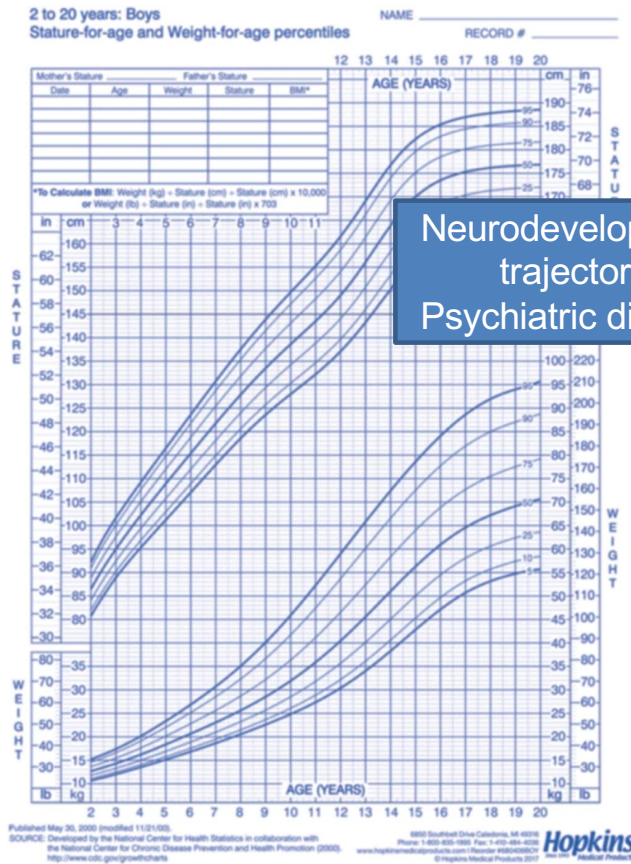


**D** Cerebellum





# No standards currently exist for brain development

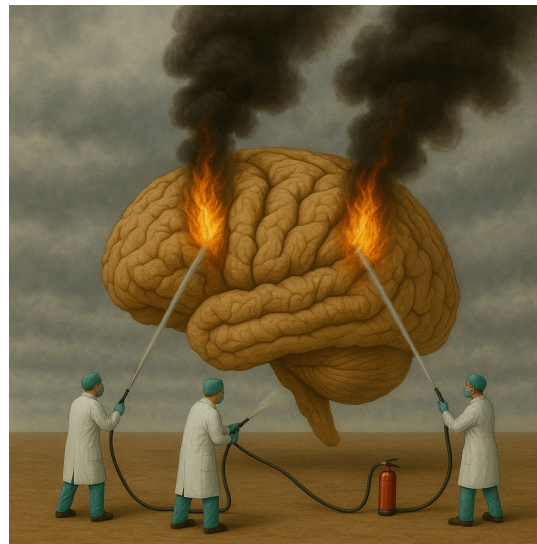


# Prevention is the Most Effective Strategy to

1. Address/alleviate the negative impact of drugs and SUD
2. promote healthy brain development
3. nip the vicious cycle of dual disorders in the bud



**Prevention**



**Treatment**



**Recovery**

**THANK YOU!**