# Marijuana: What Does Science Tell Us?

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Secretaries' Innovation Group November 17, 2015

NIH National Institute on Drug Abuse

# **State Marijuana Policies**

- State Laws related to marijuana are rapidly changing
- 24 States and DC have legalized at least some forms of MJ for medicinal use
- 4 States and DC have legalized or voted to legalize MJ for adult recreational use
- Legalization is expected to be on state ballots in 2016

### How Can Science Inform the Policy Debate?

- Drug use trends (epidemiology)
- Impact of state policies on public health outcomes
- How marijuana affects the brain (neurobiology)

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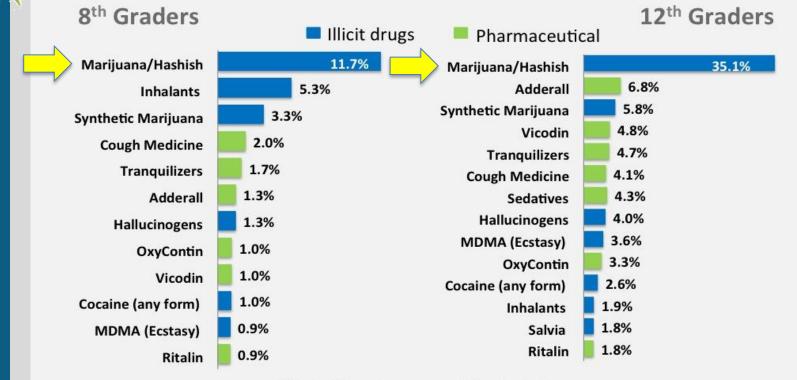
- How it affects behavior (e.g. driving)?
- How it affects life outcomes? Education?
  Jobs? Happiness? Success?
- Are the components of marijuana effective medications?



# WHAT DOES SCIENCE TELL US ABOUT MARIJUANA USE?

### Marijuana is the Most Commonly Used Illicit Drug

#### Top Drugs among 8<sup>th</sup> and 12<sup>th</sup> Graders, Past Year Use

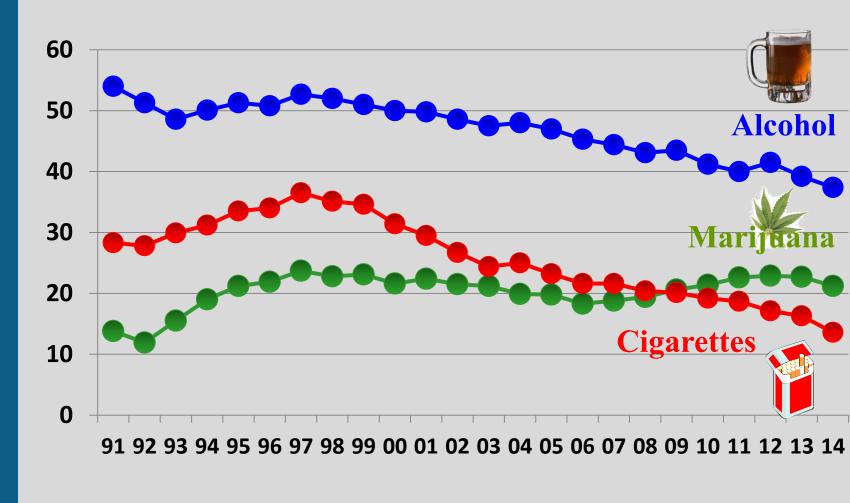


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\* Only 12<sup>th</sup> graders surveyed about sedatives use

Source: University of Michigan, 2014 Monitoring the Future Study

### How Many Youth are Using? (Past Month, %)

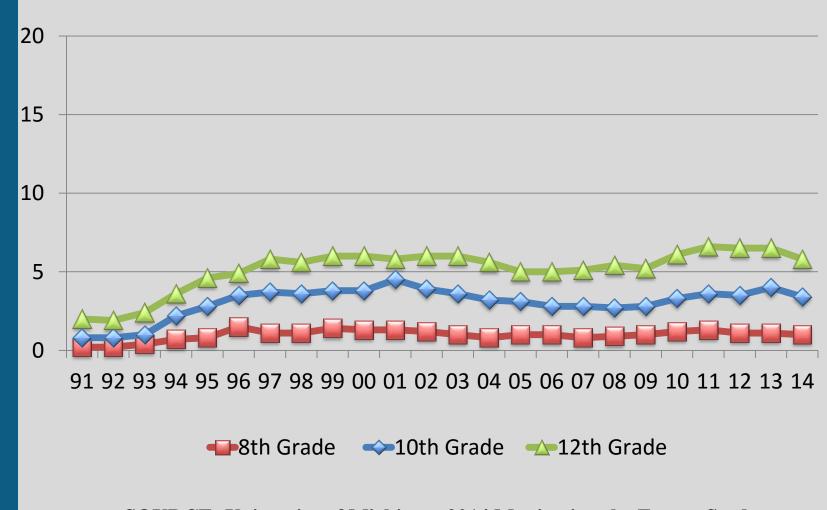


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**SOURCE:** University of Michigan, 2014 Monitoring the Future Study.

# **Percentage of High School Seniors Using Marijuana Daily**

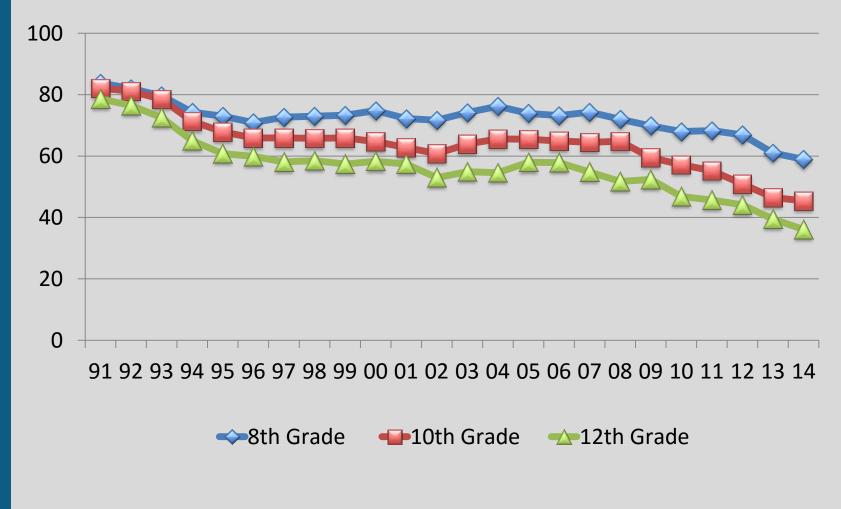


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**SOURCE:** University of Michigan, 2014 Monitoring the Future Study.

# **Decreasing Perception of Harms**

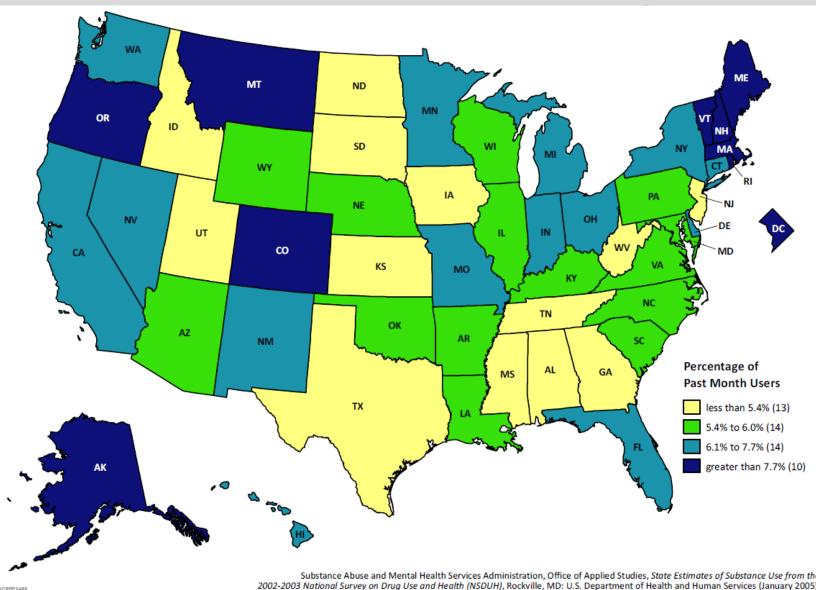


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**SOURCE:** University of Michigan, 2014 Monitoring the Future Study.

#### Percentage of Past Month Marijuana Users Aged 12 or Older: Annual Averages, 2002-2003

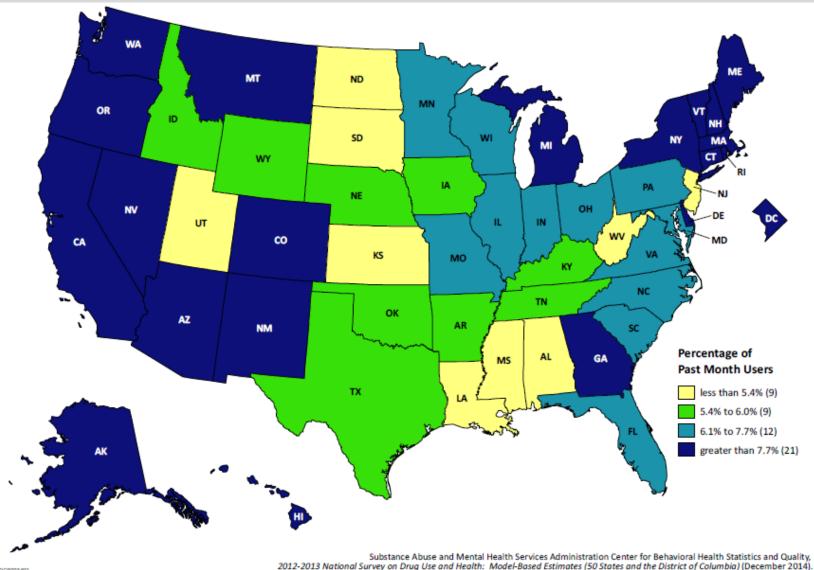


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#### Percentage of Past Month Marijuana Users Aged 12 or Older: Annual Averages, 2012-2013



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# Relationship Between Medical Marijuana Laws and Use?

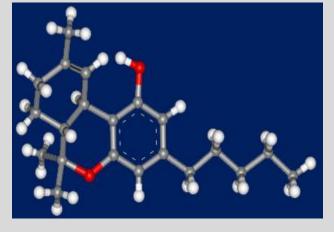
- Study results/conclusions vary
- States that legalized for medical purposes have higher rates of use
- Not necessarily causal
- Policies vary by state
- Need to consider state-level risk factors (e.g., home cultivation, dispensaries)

Source: Pacula, et al (2013), NBER Working Paper; Hasin, et al (2015)



# HOW DOES MARIJUANA IMPACT THE BRAIN?

Tetrahydrocannabinol (THC) Active Ingredient in Marijuana



### Marijuana Binds Cannabinoid Receptors Located Throughout the Brain

#### Brain Development

Memory & Cognition

 Motivational Systems & Reward

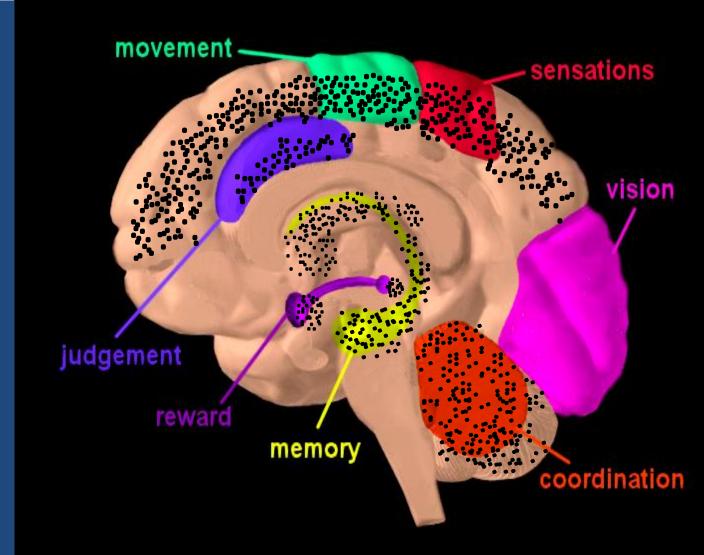
Appetite

 Immunological Function

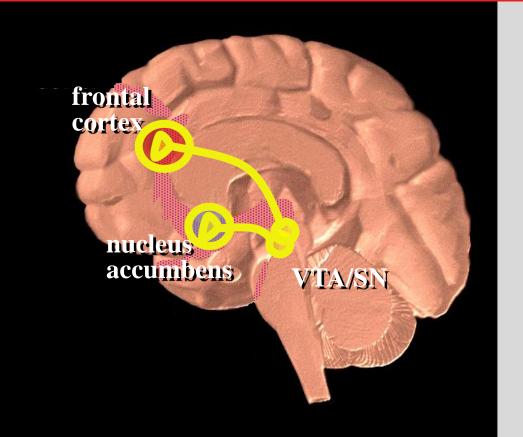
Reproduction

 Movement Coordination

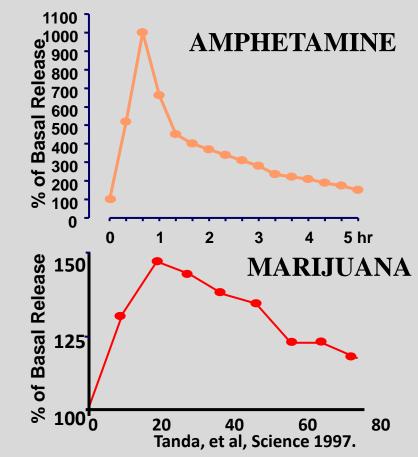
 Pain Regulation & Analgesia



# **The Reward Circuit**

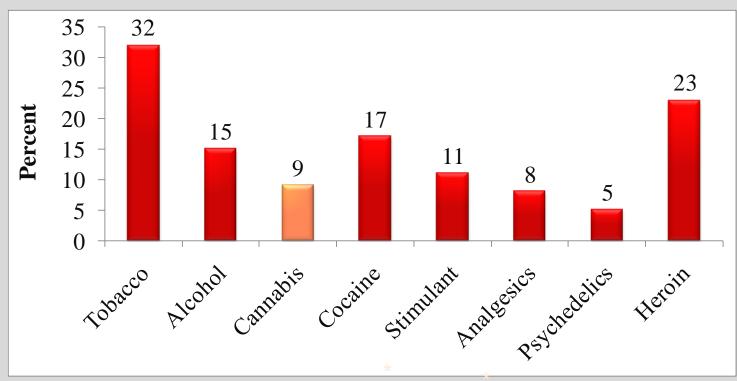


Drugs of abuse increase dopamine in the Nucleus Accumbens



# Is Marijuana Addictive?

**Estimated Prevalence of Dependence Among Users** 



#### **Risk for Addiction is Significantly Higher When Starting** in Adolescence: Approximately 16% for Marijuana

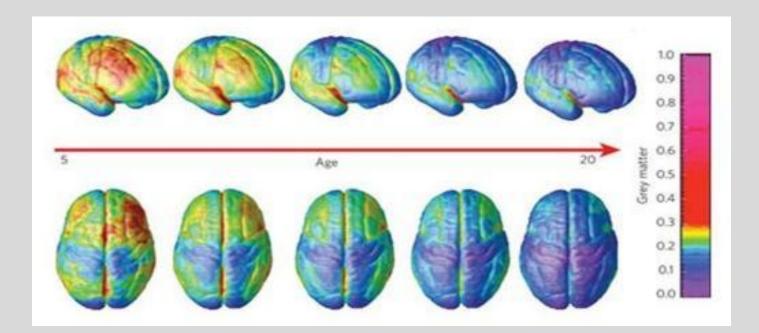
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\* Nonmedical Use; Source: Anthony JC et al., 1994

### **Marijuana and Cognitive Development**

The teen brain is **still developing** and it is especially vulnerable to drug use.

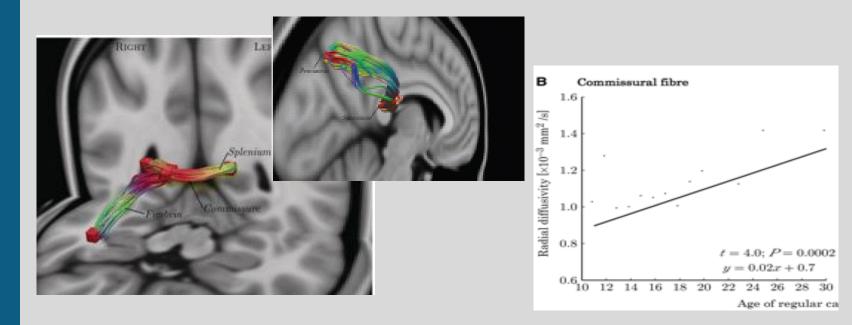


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### Marijuana and Neural Development

#### Some Studies Suggest Structural Differences in Brains of **Regular Marijuana Users versus Nonusers**

Early (<18y) Long-Term Cannabis Use Decreases Axonal Fiber Connectivity



Axonal paths with reduced connectivity (diffusion-weighted MRI) in cannabis users than in controls.

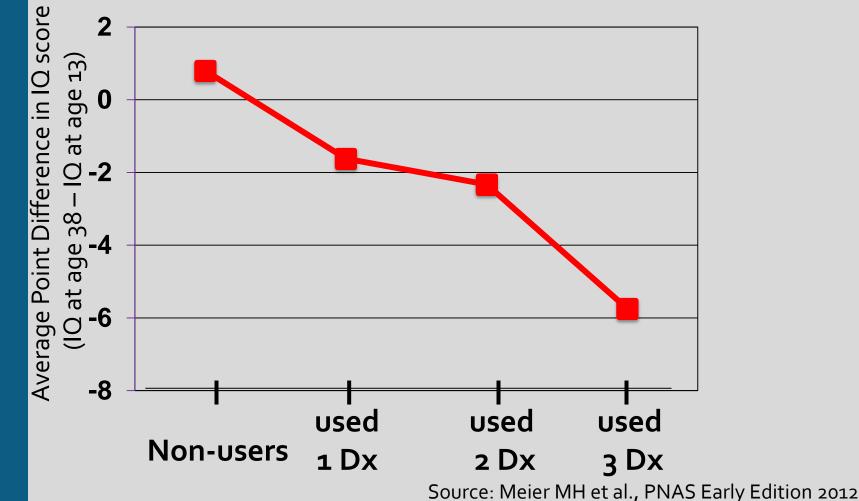
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Zalesky et al Brain 2012.

#### Persistent Marijuana Users Show A Significant IQ Drop between Childhood and Midlife

1,037 individuals (o- 38 yrs old); marijuana use reported at 18, 21, 26, 32 and 38 yrs old; IQ at 13 and 38 yrs old



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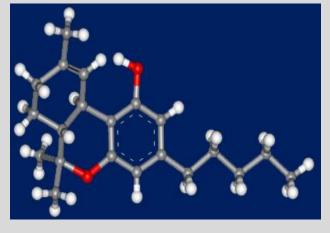
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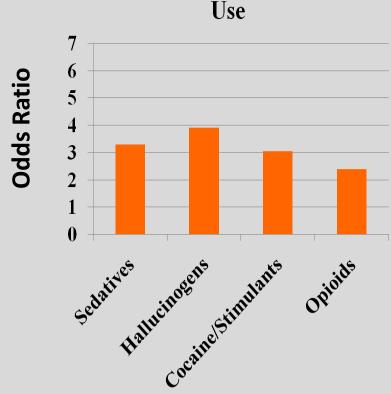
# HOW DOES MARIJUANA IMPACT LIFE OUTCOMES?

Tetrahydrocannabinol (THC) Active Ingredient in Marijuana



# Marijuana: Impact on Other Drug Use

**Similar to Early Alcohol or Tobacco Use, Early Marijuana Use Increases Likelihood of Using Other Drugs** 



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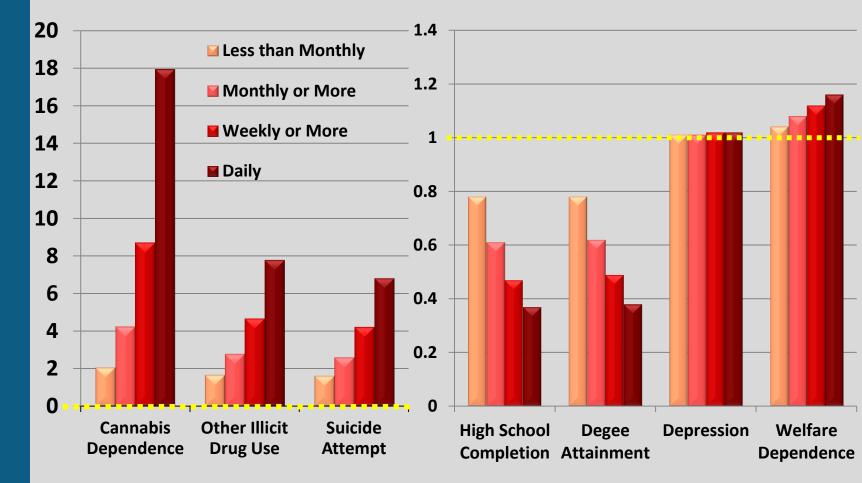
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Drug Use in Twin Pairs Discordant for Cannabis Use Before Age 17

Source: Lynskey, MT et al., JAMA, 289, pp. 427-433, 2003.

### **Marijuana: Impact on Outcomes**

More frequent use in adolescence and adverse outcomes



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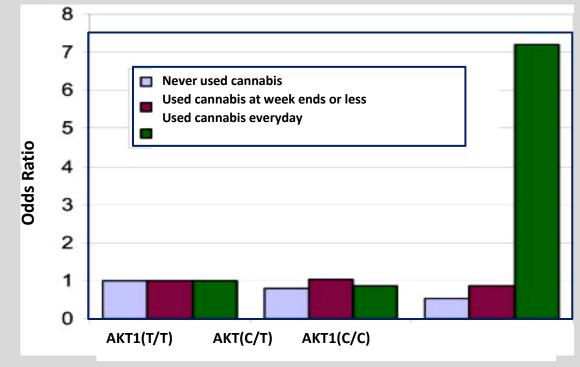
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Silins E et al., The Lancet September 2014.

# **Marijuana and Mental Illness**

Marijuana use increases risk for psychosis in people with specific genetic risk factors

Regular Cannabis Use Increases Schizophrenia Risk in those with AKT1 rs2494732 genotype



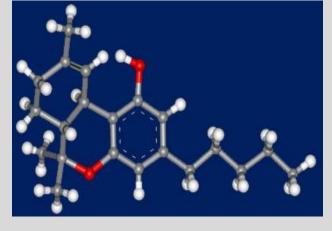
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Di Forti et al., Biological Psychiatry, 2012.



# ARE THERE THERAPEUTIC BENEFITS OF MARIJUANA?

Tetrahydrocannabinol (THC) Active Ingredient in Marijuana



# Marijuana as Medicine?

- Promise lies in purified ingredients (e.g., CBD) or synthetic compounds with more selectivity, less adverse effects
- Applications: pain, nausea, wasting, obesity, muscle spasticity, addiction, inflammatory conditions, HIV
- NIH research focuses on THC, CBD, and compounds that alter the function of the endocannabinoid system

### **Adolescent Brain Cognitive Development (ABCD) Study**

Ten year longitudinal study of 10,000 children from age 10 to 20 years to assess effects of drugs on individual brain development trajectories



# Summary

- Marijuana is the most commonly used illicit drug in US
- Use generally begins in adolescence
- Wide range of effects on brain, body, and behavior
- Policy landscape changing

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- Impact of policies need further research
- Science can/should inform policy





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