Marijuana and Cognition: A Hazy Issug



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Overview

- History
- How does marijuana work?
- Effects (some data)
 - What about college students?
- Medical marijuana
- Status and the future



What is Marijuana?

- Cannabis sativa
- Cannabis indica

Weed, Pot, Reefer Grass, Ganja Mary Jane, Hash Skunk, Blunt

Components

CANNABIS VS HEMP VS MARIJUANA

Cannabis

is a plant family that includes many species, including both hemp and marijuana.





Hemp

is a variety of cannabis grow for its tall, sturdy stalks and **low levels of THC.**

Marijuana

is a variety of cannabis grown for its **high levels of THC.**

Even though Hemp contains low levels of THC it also contains higher levels of other beneficial cannabinoids (like CBD) which may promote relaxation and calming effects in the body.

Marijuana History



PSYCHOPHARMACOLOGY 2e, Figure 14.4





Our History with Drugs

COUCH SYRUP

ALCOHOL (locathan 1%)	4/4m
ALCOHOL, HESSMan TO	4%而.
CANNABIS INDICA, F.E.,	21/200
CHLOROFORM	_C/5115
MODDUIA SIII PH	. Yagr
MORFINA, SOLFI,	
	and

SKILLFULLY COMBINED WITH A NUMBER OF OTHER INGREDIENTS

DIRECTIONS DOST-One half teaspoonful three

COCAINE TOOTHACHE DROPS Instantaneous Cure !

COBS PHARMACY

SURGICA

RUSSE

GTICRINES

FLUIDEXTRACT

Greater New fort

1950年1月1日日日

ACOBS PHARMAC

PRICE 15 CENTS. Prepared by the LLOYD MANUFACTURING CO. 219 HUDSON AVE., ALBANY, N. Y.

For sale by all Druggists. (Registered March 1885.) See other side.

JACOBI

Hey Man, What Exactly is in Marijuana?

- 1805 Morphine was isolated from opium
- 1855 Cocaine was isolated from coca leaves
- 1963 THC in marijuana
- Raphael Mechoulam, Israel
 - Got 5 kilos of confiscated Lebanese hash from the Israeli national police
 - Experimented with monkeys (normally aggressive → sedated)
 - Isolated THC then CBD







Receptor Sites



CB1

CB₂

Receptors



Acute Effects

There is no dispute that cannabis produces cognitive impairment during acute intoxication

- Impaired coordination, memory, attention and judgement
- Euphoria
- Peripheral effects -Increased HR

Frequent vs Occasional Users

Residual Effects

- Persistent effects (lasting a few days or weeks) due to:
 - Residue of cannabinoids in the brain
 - •Memory and learning skills
 - •Withdrawal symptoms
- Long-term effects due to toxicity to the brain
 - Cumulative effects of chronic abuse



Common Effects of Marijuana with Regard to Students



- Affects several mental and physical activities:
 - Short-term memory
 - Coordination
 - Learning
 - Problem solving





The Adolescent Brain



The adolescent brain is undergoing significant neurodevelopment well into the 20s, and the regions that are last to develop are most populated by cannabis receptors and are critical to cognitive functioning



PSYCHOPHARMACOLOGY 2e, Figure 14.17 © 2013 Sinauer Associates, Inc.

Brain Maturation

Nucleus Accumbens

(highly active in adolescence)

- Pleasure center
- Active during pleasurable activities, including feeding, sex, and drug use
- Impulsive behavior
- Plays a major role in addiction



Prefrontal Cortex (matures late adolescence or early adulthood)

- Impulse control
- Decision-making
- Organizing and planning
- Abstract thought, rational thinking
- Attention, focus
- Working memory





Drug Use and College Students

- Annual use of illicit drugs other than marijuana - 18% in 2017 for college and non-college youth
- The 2017 annual prevalence of nonmedical use of prescription narcotic drugs (other than heroin), such as OxyContin and Vicodin, was 3.1% for college students and 4.1% for non-college youth



Reasons Young Adults Use Marijuana — Experimentation vs Motivation

- Peer pressure
- Belief That Marijuana Is Harmless
- Availability and Opportunity
- To feel good
- Ease boredom
- Relieve tension or frustration
- Seek deeper insights
- Escape problems
- To increase (or decrease) the effects of other drugs



Marijuana and College Students

- The more students report using marijuana, the more they skip classes, have lower GPAs, have enrollment gaps, and do not graduate on time
 - Discontinuous enrollment during college: Associations with substance use and mental health Amelia M. Arria, Ph.D et al, Psychiatr Serv, 2013
- College students who use marijuana have lower GPAs than those who do not and are more likely to drop out and less likely to graduate on time
 - Marijuana use trajectories and academic outcomes among college students Suerken CK et al, Drug and Alcohol Dependence, 2016
- A significant decline in GPAs over two years when students used both marijuana and alcohol at moderate to high levels
 - Longitudinal influence of alcohol and marijuana use on academic performance in college students, Meda SA et al, PLOS one, 2017

What do the data show?

STUDIES	Cognitive	Brain Structure	Brain Function Effects
Meier et al., 2012	↓IQ		
Pope et al., 2003	↓ IQ		
Ehrenreich et al., 1999	↓ attention		
Huestegge et al., 2002	\downarrow visual search		
Fontes et al., 2011	↓ executive functioning		
Solowij et al., 2012	↓ executive functioning		
Churchwell et al., 2010		↓ prefrontal cortex volume	
Gruber et al., 2011	↑ impulsivity	\downarrow white matter integrity in prefrontal cortex	
Lopez-Larson et al., 2011		↓ prefrontal cortex thickness	
Wilson et al., 2000		\downarrow total gray matter,	
Becker et al., 2010a			↑ prefrontal cortex activity during working memory task
Gruber et al., 2012			\downarrow anterior cingulate activity during inhibition task
Jager et al., 2010			↑ prefrontal cortex activity during in working memory

Stroop Test fMRI Activation

PURPLE YELLOW RED BLACK RED GREEN RED YELLOW ORANGE BLUE PURPLE BLACK RED GREEN ORANGE



Sagar, K. A., Dahlgren, M. K., Gönenç, A., Racine, M. T., Dreman, M. W., & Gruber, S. A. (2015). The impact of initiation: Early onset marijuana smokers demonstrate altered Stroop performance and brain activation. Developmental Cognitive Neuroscience, 16, 84-92.

Memory



N. Solowij; K.A. Jones; M.E. Rozman; S.M. Davis; J. Ciarrochi; P.C.L. Heaven; D.I. Lubman & M. Yücel; Verbal learning and memory in adolescent cannabis users, alcohol users and non-users; Psychopharmacology (2011) 216:131–144

Persistent cannabis users show neuropsychological decline from childhood to midlife

Madeline H. Meier^{a,b,1}, Avshalom Caspi^{a,b,c,d,e}, Antony Ambler^{e,f}, HonaLee Harrington^{b,c,d}, Renate Houts^{b,c,d}, Richard S. E. Keefe^d, Kay McDonald^f, Aimee Ward^f, Richie Poulton^f, and Terrie E. Moffitt^{a,b,c,d,e}

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

Followed 1,037 individuals from birth to age 38. Tested marijuana use at 18, 21, 26, 32 and 38. Tested for IQ at ages 13 and 38



Marijuana Use and IQ



Persistent cannabis users show neuropsychological decline from childhood to midlife; M.H. Meier, A. Caspi, A. Ambler, H. Harrington, R. Houts, R.S.E. Keefe, K. McDonald, A. Ward, R. Poulton, and T.E. Moffitt; PNAS (2012) 109 (40)

Does Cessation of MJ use Restore Neuropsychological Functioning?



Adolescent-Onset (Used Cannabis Weekly Before Age 18)

Adult-Onset (Did Not Use Cannabis Weekly Before Age 18)

Persistent cannabis users show neuropsychological decline from childhood to midlife; M.H. Meier, A. Caspi, A. Ambler, H. Harrington, R. Houts, R.S.E. Keefe, K. McDonald, A. Ward, R. Poulton, and T.E. Moffitt; PNAS (2012) 109 (40)

Harvard Study Shows Smoking Marijuana Improves Cognitive Function

Scientific findings indicate that pot use improves cognitive performance.

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Medical Marijuana Improves Executive Function in the Brain



Unlike recreational marijuana, medical marijuana has benefits that haven't been fully tested out — until now.

Wait.... What??

THC * CBD



Medical Marijuana

THC

 Helps with pain, nausea, sleep apnea, stress disorders

CBD

- Helps with inflammation, stress, depression, anxiety, cancer, epilepsy, schizophrenia, multiple sclerosis, migraines, arthritis, the side effects of cancer treatments
- Does not have psychoactive effects like THC

Medical Marijuana







Federal Government Stance on Marijuana

Legal



Legal



Legal 100 mL * Epidiolex® 100 mg/mL Oral Solution Epidiolex oral administration of

GREENWICH

Rx On

100 mg/mL Onal Solution

Illegal



Marijuana Cigarettes, Medium Potency RTI Log No. <u>12792-0109-120</u> Approximately 300 cigarettes per can lite Weight = <u>20.02 g</u> Average weight per cigarette = <u>6.8%</u> ± <u>6.06 g</u> Manufactured January 2009 Expiration Date: To Be Determined Expiration Date: To Be Determined Can No. = 5537 earch Triangle Institute







Since 1976

Since 1985





HealTHCare

- In chemotherapy induced nausea and vomiting, oral cannabinoids are effective antiemetics
- In chronic pain, patients who were treated with cannabis or cannabinoids are more likely to experience a clinically significant reduction in pain symptoms
- In multiple sclerosis (MS) related spasticity, short-term use of oral cannabinoids improves patient-reported spasticity symptoms
- Evidence suggests that CBD may be an effective treatment for patients with drug-resistant forms of epilepsy





Key Points

- Recreational marijuana use
 relates to poorer cognition across
 numerous domains (particularly
 executive function and memory),
 as well as alterations in brain
 structure and function
- Those with the highest neurodevelopmental vulnerability (adolescents) are using MJ more frequently than in previous years
 - Earlier age of MJ onset (adolescence), as well as higher frequency and magnitude of use, relate to further impairment

Key Points

Medical marijuana may confer a unique impact on the brain, given that its consumers are often adults who are beyond critical neurodevelopmental periods and who may also seek products based on therapeutic potential, rather than for mood altering effects

Thank You!



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