

Blunt Truths: Debunking Myths about Cannabis in the Perinatal and Postpartum Period

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Disclosure

- I, Luke Peterson, hereby declare that the content for this activity, including any presentation on therapeutic options, is well balanced, unbiased, and to the extend evidenced based.
- My partner/spouse and I have no financial relationships with commercial entities producing, marketing, re-selling, or distributing health care goods or services consumed by, or used on, patients relevant to the content I am planning, developing, presenting, or evaluating.
- Off label discussion of medication for cannabis use disorder will be discussed and noted in this presentation.



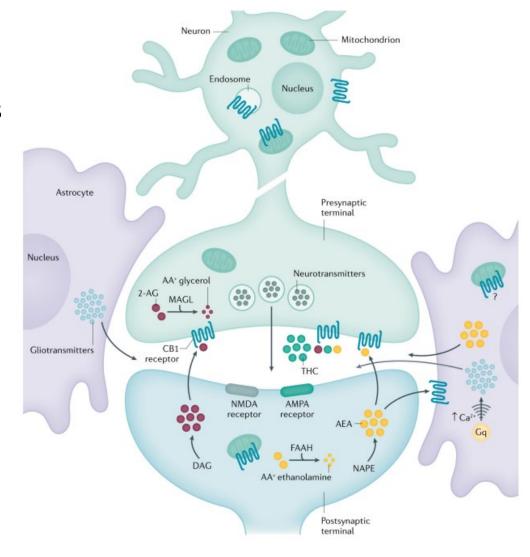
Objectives

- Review risks of cannabis use in the perinatal and postpartum period
- Provide anticipatory guidance for patients using cannabis while breastfeeding
- Discuss biopsychosocial treatment for patients with cannabis use disorder



Cannabis Pharmacology

- Marijuana contains over 500 chemical compounds
- Δ9 -tetrahydrocannabinol (THC) is major psychoactive compound
 - Partial agonism of presynaptic cannabinoid receptor
 CB1 euphoric effect
 - Inhibition of adenylyl cyclase activity through activation of Gi/o protein
 - Reduces presynaptic neurotransmitter release
 - Highest expression of CB1 is in glutamate and GABAergic neurons
- Reward pathway
 - THC inhibits GABA in the ventral tegmental presynaptic neuron which increases dopamine release
- Metabolized by liver various P450 enzymes





Cannabis Pharmacology



INHALED

- · Smoking, vaping or dabbing
- · Quickest method
- Less chance of overconsumption

Onset (Min): 1-3 Duration (hrs): 1-3

ORAL ABSORPTION (GI)

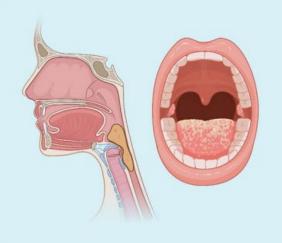
- Oral: Drops, tinctures, sprays, lollipops, strips
- GI: Edibles, candies, drinks, snacks, capsules
- Delay in onset symptoms/ Higher likelihood of overconsumption

Onset (min)

- Oral: 10-25
- Ingested: 30-90

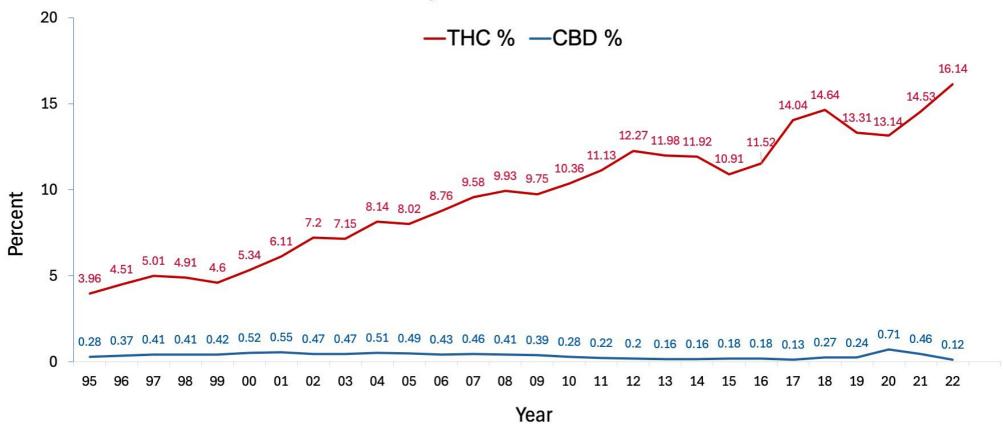
Duration (hrs):

- Oral: 1-3
- Ingested: 6-8





Percentage of THC and CBD in Cannabis Samples Seized by the DEA, 1995-2022

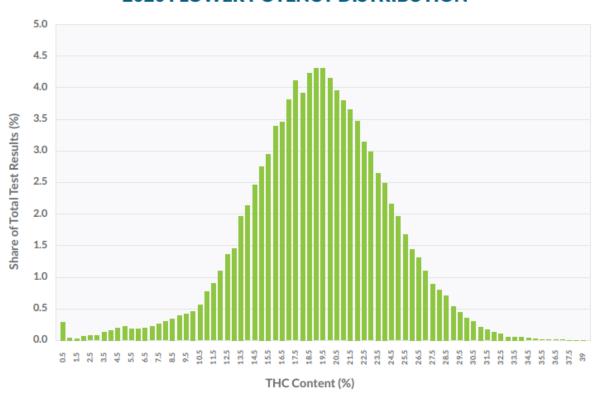


SOURCE: U Miss, Potency Monitoring Project

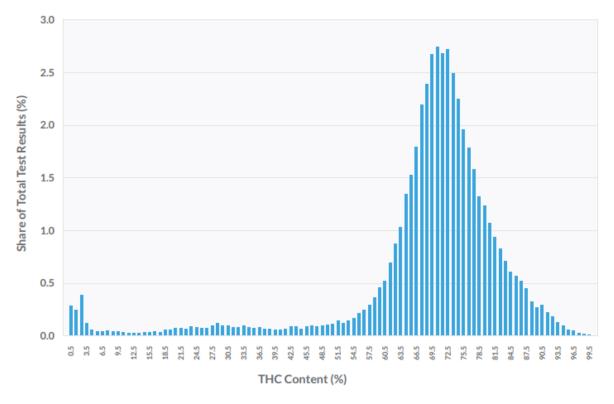


Potency Distribution for Flower & Concentrates

2020 FLOWER POTENCY DISTRIBUTION



2020 CONCENTRATE POTENCY DISTRIBUTION





Effects of Cannabis





EFFECTS OF DELTA-9 THC

Short-Term

Effects on the Brain and Nervous System

- Altered senses
- · Distorted perception of time
- Changes in mood
- Delusions
- · Difficulty thinking
- · Impaired movement
- Impaired memory
- Hallucinations
- Psychosis

Long-Term

Effects on the Brain and Nervous System

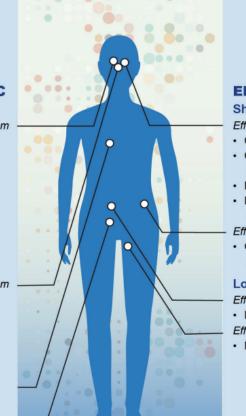
- Impacts on child development (if used when pregnant)
- Impaired brain development
- Impaired learning, memory, and thinking
- Mental illness (e.g., depression, anxiety, paranoia)

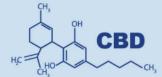
Effects on the Respiratory System

· Breathing problems

Effects on the Digestive System

· Nausea and vomiting





EFFECTS OF CBD

Short-Term

Effects on the Brain and Nervous System

- Changes in alertness
- Changes in mood (e.g., irritability, agitation)
- · Drowsiness and/or sedation
- Drug interactions that may cause serious side effects

Effects on the Digestive System

 Gastrointestinal distress (e.g., nausea, diarrhea)

Long-Term

Effects on the Digestive System

Liver injury

Effects on the Reproductive System

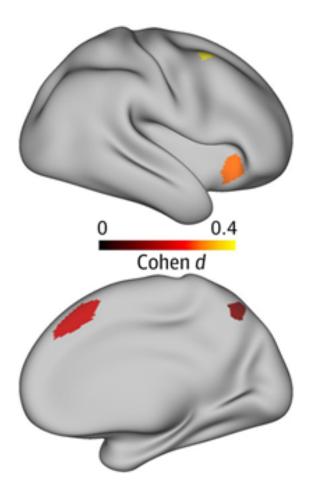
· Male reproductive toxicity

Effects of Cannabis

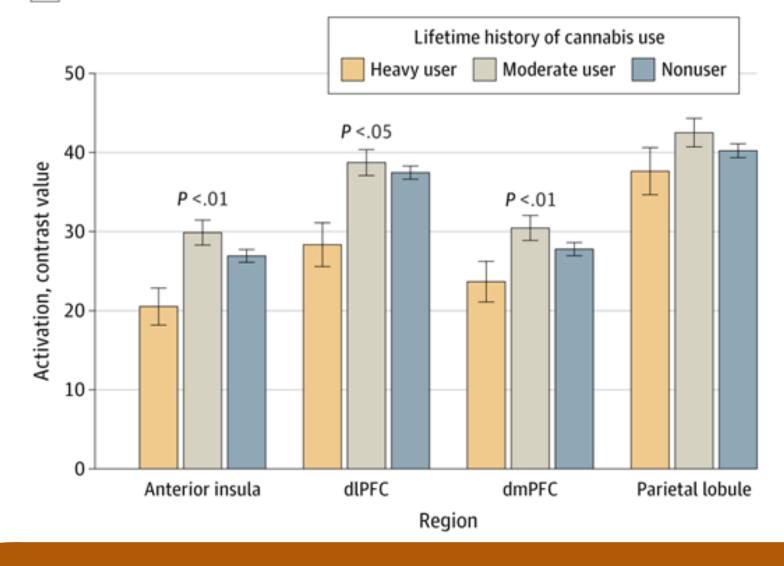
- Decreases working memory
- Temporary trends in motor vehicle accidents
- Increase in ER presentations due to cannabis use
- Increase in cannabis use disorder prevalence after medicinal and recreational legalization
 - Larger increase in patients with chronic pain
- Increase in cardiovascular disease
- Psychosis/paranoia occurs in 1 in 10 people who use daily/regularly
- Increase of anxiety (OR 1.04-1.15)



A Region and effect size



B Working memory task (2-back – 0-back)





 Cannabis use disorder prevalence by state status for legalization of cannabis

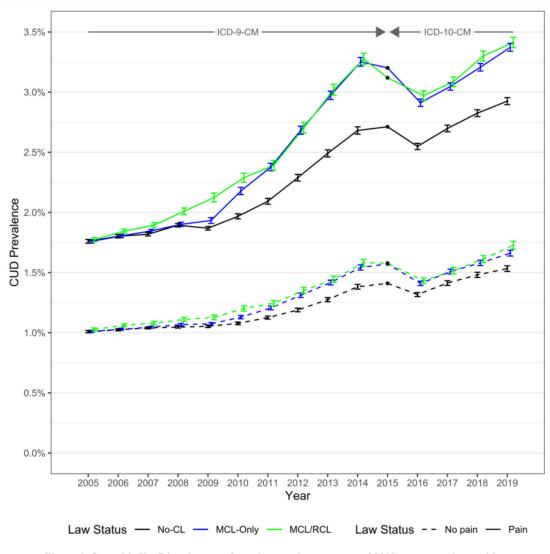
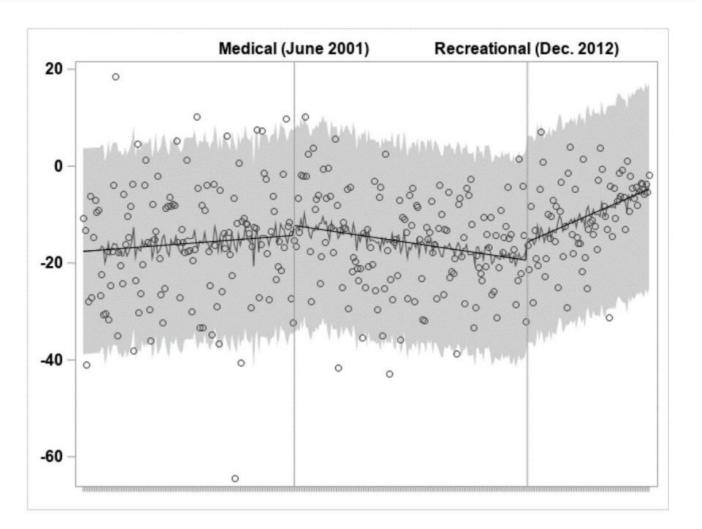


Figure 1. Cannabis Use Disorder prevalence by state law status as of 2019, among patients with and without chronic pain



 Multiple studies show increase in non-fatal and fatal car crashes after passing of marijuana legislation, in particular recreational laws.

Difference in monthly fatal motor vehicle per million population crashes





Cannabis associated with heart disease

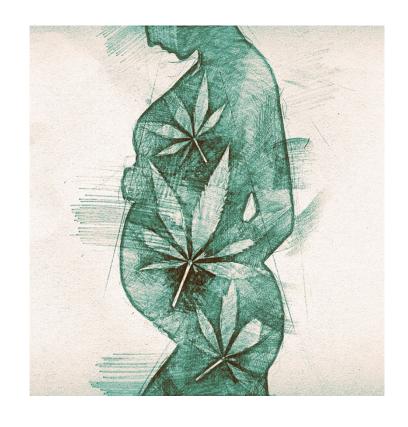
- Jeffers et al, 2024
 - Controlled for tobacco smoking status, age, sex, race and ethnicity, body mass index, diabetes, alcohol use, educational attainment, and physical activity
 - Past 30 day cannabis use associated with
 - Myocardial infarction
 - Stroke
 - Composite cardiovascular disease
 - More prevalent in people who use more frequently

Cardiovascular disease	aOdds Ratio	95% CI
Coronary heart disease	1.16	0.98-1.38
Myocardial infarction	1.25	1.07-1.46
Stroke	1.42	1.20-1.68
Composite (CHD, MI, CVA)	1.28	1.13-1.44



Cannabis use

- In the last decade, cannabis use among child bearing aged people has increased from 10.4% to 18%
- Cannabis use in pregnancy has increased from 3.4% (2002) to 7% (2017)
 - COVID19 pandemic 2020 reported 25%
 - Highest prevalence among 18-25 years and during 1st trimester



Cannabis use in prenatal period



Serious maternal complications linked with use of marijuana before and early in pregnancy, study says

By Sandee LaMotte, CNN

① 6 minute read · Updated 4:39 PM EDT, Mon July 22, 2024





"While it is sensible to advise women to abstain from all drugs during pregnancy, the weight of scientific evidence suggests that marijuana dose not directly harm the human fetus" - 420 Magazine, March 25, 2013, Use of Marijuana during pregnancy









As is the case for many young women, my indulgence in recreational drugs, including alcohol and caffeine, came to an abrupt halt when my husband and I discovered we were pregnant with our first child. To say we were ecstatic is an understatement. Doctors had





Perception of Cannabis use in Pregnancy

- Perception that cannabis has no risk has risen three-fold from 2005 to 2015
- 70% of pregnant and non-pregnant women believe there is little or no risk of using cannabis one to two times per week
- Most commonly cited source of information for cannabis use in pregnancy is the internet and anecdotal evidence



Recommendations From Cannabis Dispensaries About First-Trimester Cannabis Use

Dickson, Betsy MD; Mansfield, Chanel MPH; Guiahi, Maryam MD, MSc; Allshouse, Amanda A. MS; Borgelt, Laura M. PharmD; Sheeder, Jeanelle PhD; Silver, Robert M. MD; Metz, Torri D. MD, MS

Author Information ⊗

Obstetrics & Gynecology 131(6):p 1031-1038, June 2018. | DOI: 10.1097/AOG.0000000000002619

Findings:

- 69% recommended cannabis for morning sickness
- 36% safe in pregnancy
- 81% recommended discussion with healthcare provider



A word of caution....

- Limited and poor studies have led to fear mongering and finger pointing
- Many studies associating drug use with risk in pregnancy don't account for other confounding issues that people with substance use face:
 - Nicotine and alcohol use, prenatal care, insured, mental health, medical history, psychopharmacology, domestic violence, trauma, SDoH, unstable housing, STD, nutrtion, blood born pathogens, etc.
- Lesson's learned...
 - "Yet, after almost 20 years of research, not one single condition or disorder that could be labeled "crack baby" has been identified. Nor is there evidence of the extent of harm that was predicted by physicians and trumpeted by the media." (Lewis, 2005)
- Recommendation: Be skeptical, and dig in to the evidence



Cannabis effect on pregnancy fetus and neonate

- Increased risk of NICU admission
- Lower birth weight
- Small of gestational age
- Preterm birth
- Higher still birth rates
- Lower Apgar scores
- Growth of infant while breastfeeding
- Decreased childhood and adolescent memory and problems with impulse control, problem solving, quantatative reasoning, verbal development, visual analysis tests





Supplement 4: Forest Plot 1: Odds of Maternal Anemia among Cannabis Users

				Odds Ratio	Odds Ratio
Study or Subgroup	log[Odds Ratio]	SE	Weight	IV, Fixed, 95% CI	IV, Fixed, 95% CI
Berenson 1996	-0.35	0.99	1.2%	0.70 [0.10, 4.91]	-
Gargari 2012	0.1773	1.147	0.9%	1.19 [0.13, 11.31]	
Greenland 1982	0.3502	0.80434	1.8%	1.42 [0.29, 6.87]	- •
Greenland 1983	-0.4081	0.503	4.6%	0.66 [0.25, 1.78]	
Quinlivan 2002	-0.05	0.275	15.4%	0.95 [0.55, 1.63]	
Witter 1990	0.4381	0.1236	76.1%	1.55 [1.22, 1.97]	
Total (95% CI)			100.0%	1.36 [1.10, 1.69]	◆
Heterogeneity: Chi²=	5.29, df = 5 (P = $0.$	38); l² = 59	%		1 10 100
Test for overall effect:	Z= 2.88 (P = 0.004	4)			0.01 0.1 1 10 100 Normal Hemoglobin Anemia

Exclusions/Adjustments

(Witter) 1990, recruitment in 1980's, only crude data



Supplement 3: Forest Plot 3: Odds of Low Birth Weight among Infants Exposed to Cannabis

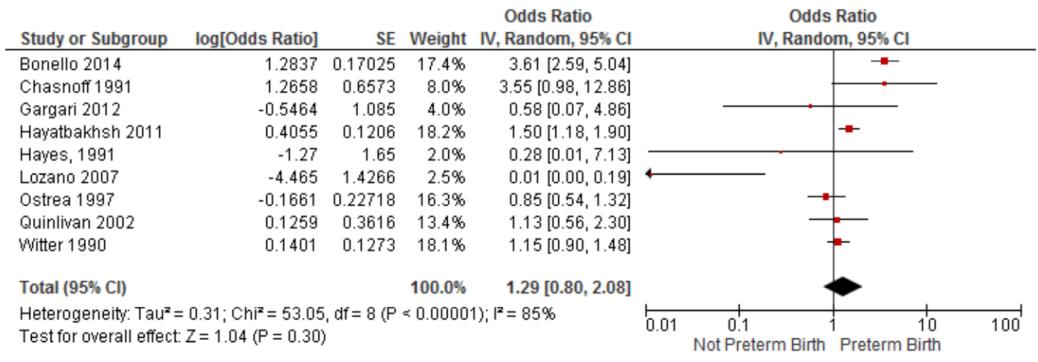
				Odds Ratio	Odds Ratio
Study or Subgroup	log[Odds Ratio]	SE	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Berenson 1996	0.1823	0.4322	13.2%	1.20 [0.51, 2.80]	- -
Bonello 2014	1.579	0.159	18.7%	4.85 [3.55, 6.62]	
Greenland 1982	0.4661	0.94661	5.8%	1.59 [0.25, 10.19]	- •
Greenland 1983	1.0074	0.85404	6.7%	2.74 [0.51, 14.60]	
Hayatbakhsh 2011	0.53	0.13	19.1%	1.70 [1.32, 2.19]	- - -
Ostrea 1997	0.1875	0.2296	17.5%	1.21 [0.77, 1.89]	 -
Witter 1990	0.13235	0.1318	19.1%	1.14 [0.88, 1.48]	_
Total (95% CI)			100.0%	1.77 [1.04, 3.01]	•
Heterogeneity: Tau² =	0.36; Chi ² = 55.18	, df = 6 (P	< 0.0000°	1); I²= 89%	0.01 0.1 1 10 100
Test for overall effect:	Z = 2.12 (P = 0.03)				0.01 0.1 1 10 100 Normal Birth Weight Low Birth Weight

Exclusions/Adjustments

(Berenson)	Adjusted for race, #cigarettes, alcohol
(Bonello)	Pre-existing DM/HTN, PreE, GDM, smoking, SES, delivery, infant gender, fetal death
(Greenland)	Matched based off age, ethnicity (laxed), gravida – tobacco and alcohol allowed
(Greenland)	Crude numbers
(Hayatbakhsh)	Mothers age, arity, ethnicity, weight, cigarette, alcohol, other substances
(Ostrea)	Crude numbers
(Witter)	1990, recruitment in 1980's, only crude data



Supplement 3: Forest Plot 7: Odds of Preterm Birth Among Infants Exposed to Cannabis in-utero



Exclusions/Adjustments

Exclusions/ Auje	astinents
(Bonello)	Pre-existing DM/HTN, PreE, GDM, smoking, SES, delivery, infant gender, fetal
(Chasnoff)	Age, race, SES
(Gargari)	Crude numbers
(Hayatbakhsh)	Mothers age, arity, ethnicity, weight, cigarette, alcohol, other substances
(Hayes)	Not confounded by drugs, alcohol, tobacco, low birth weight
(Loranzo)	Cigarette smoking, sex, maternal age and gestation weeks
(Ostrea)	Crude numbers
(Quinlivan)	Height, weight, age, race, GA, gender, smoking and alcohol use
(Witter)	1990, recruitment in 1980's, only crude data



Supplement 3: Forest Plot 8: Odds of Admission to NICU/ICU among Infants Exposed to Cannabis inutero

Study or Subgroup	log[Odds Ratio]	SE	Weight	Odds Ratio IV, Random, 95% CI	Odds Ratio IV, Random, 95% CI	
Berenson 1996	-0.69	0.525	13.9%	0.50 [0.18, 1.40]		
Bonello 2014	1.0438	0.13219	38.5%	2.84 [2.19, 3.68]		
Gargari 2012	1.6796	0.8419	6.7%	5.36 [1.03, 27.93]	•	
Hayatbakhsh 2011	0.6931	0.093	40.9%	2.00 [1.67, 2.40]	-	
Total (95% CI)			100.0%	2.02 [1.27, 3.21]	•	
Heterogeneity: Tau² = Test for overall effect:	-	-	= 0.003);	I ^z = 78%	0.01 0.1 1 10 10 No NICU/ICU Placement NICU/ICU Placement)O

Exclusions/Adjustments

(Berenson) Adjusted for race, #cigarettes, alcohol

(Bonello) Pre-existing DM/HTN, PreE, GDM, smoking, SES, delivery, infant gender, fetal death

(Gargari) Crude numbers

(Hayatbakhsh) Mothers age, arity, ethnicity, weight, cigarette, alcohol, other substances



Cannabis effect on fetus and neonate

- Gunn et al, 2024
 - Systematic Review/Meta analysis
 - Cannabis exposure increases:
 - NICU Admission
 - Low birth rate
 - Anemia*
 - No statistical association with:
 - Gestational age
 - NICU/ICU length
 - Head circumference
 - Preterm birth*

- Avalos et al, 2024
 - Retrospective population cohort (Kaiser)
 - Adjusted for SES, non-cannabis substance use, medical and mental health comorbidities, and adequacy of prenatal care
 - Cannabis exposure increases (OR [95% CI]):

 Low birthweight 	1.2 (1.12-1.28)
• SGA	1.24 (1.18-1.30)
Preterm birth <37w	1.06 (1.00-1.13)
 NICU admission 	1.06 (1.01-1.11)



Cannabis effect on development (limited data)

- Maternal Health Practices and Child Development (MHPCD) study (N=763)
 - Inconsistent findings for timing of developmental delays
 - Lower performance on Stanford-Binet Intelligence Score @ 36 mo
 - Deficits in short term memory and verbal reasoning @ 6 years
 - Reading and spelling deficits @ 10-14 years
 - Lower teacher's ratings on memory, impulsivity, hyperactivity, and delinquency
- Ottawa Prenatal Prospective Study (OPPS) (N=583)
 - Increased behavioral problems
 - Decreased language comprehension, memory, and attention
 - Increased impulsivity at age 9-12



Cannabis effect on development (limited data)

- Generation R study (N=7,452) ongoing
 - Behavioral problems but not emotional problems
 - No change in global brain volume but had thicker frontal corticies
- IQ OPPS and other studies show normal IQ
- Autism Spectrum Disorder Retrospective Canadian study showed association with cannabis exposure (aHR 1.51, 95% CI 1.17–1.96)
- Sleep Very small cohort study (N=38) showed disturbed nocturnal sleep
- Mental health conflicting evidence about MH conditions such as psychosis and other psychopathology including ADHD and SUD.
- No congenital abnormalities in most studies, may be associated with anencephaly with early first trimester exposure (note THC users are less likely to supplement folic acid during pregnancy).





Cannabis and pregnancy risk



ACOG COMMITTEE OPINION

Number 722 • October 2017

(Replaces Committee Opinion No. 637, July 2015)

Committee on Obstetric Practice

This document reflects emerging clinical and scientific advances as of the date issued and is subject to change. The information should not be construed as dictating an exclusive course of treatment or procedure to be followed.

INTERIM UPDATE: This Committee Opinion is updated as highlighted to reflect a limited, focused change in the language and supporting evidence regarding marijuana use and neonatal outcomes.

Marijuana Use During Pregnancy and Lactation

- No association with maternal or perinatal mortality
- Increase risk of still birth
- Lower birth weight (<2500g) appears dose dependent (use at least weekly)
- Preterm birth association appears to be confounded by tobacco use

"Because the effects of marijuana use may be as serious as those of cigarette smoking or alcohol consumption, marijuana also should be avoided during pregnancy"



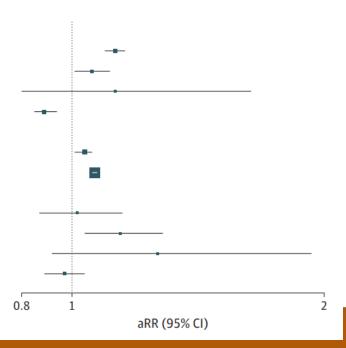
Cannabis and pregnancy risk

- Young-Wolff et al 2024 (N=316,722, 6.3% +THC)
 - Adjusted for SES, parity, birth year, prenatal care initiation, prepregnancy BMI, noncannabis prenatal substance use, and comorbidities.

Figure. Adjusted Risk Ratios (aRRs)^a of Maternal Outcomes in Pregnancies With Any Prenatal Cannabis Use vs None^b

*Gestational hypertension showed dose response increasing in risk from reported monthly, weekly, and daily THC use.

aRR (95% CI)
1.17 (1.13-1.21) ^c
1.08 (1.01-1.15) ^c
1.17 (0.80-1.71)
0.89 (0.85-0.94) ^c
1.05 (1.01-1.08) ^c
1.09 (1.08-1.10) ^c
1.02 (0.87-1.20)
1.19 (1.05-1.36) ^c
1.34 (0.92-1.95)
0.97 (0.89-1.05)





Cannabis use and breastfeeding

- Postnatal THC exposure is poorly understood/studied
- THC is most common substance in lactating individuals (5-15%)
- THC is 8-times higher in milk v serum
- Infants exposed to 0.8-2.5% of the mother's dose
- Small studies with conflicting findings regarding infant development
 - Often confounded by prenatal exposure



"In the absence of definitive data on infant safety, professional guidelines encourage reducing or eliminating cannabis use while breastfeeding. The ABM suggest that breastfeeding mothers be counseled to reduce or eliminate use to avoid possible neurobehavioral effects on the infant."



Screening for cannabis use disorder in pregnancy

 ACOG Opinion Statement 722: "Before pregnancy and in early pregnancy, all women should be asked about their use of tobacco, alcohol, and other drugs, including marijuana and other medications used for nonmedical reasons."

Screening tool	Sensitivity %	Specificity %
4P's	90.2	29.6
NIDA Quick Screen	79.7	82.8
SURP-P	92.4	21.8

Reference: hair and urine tests combined, positive on either



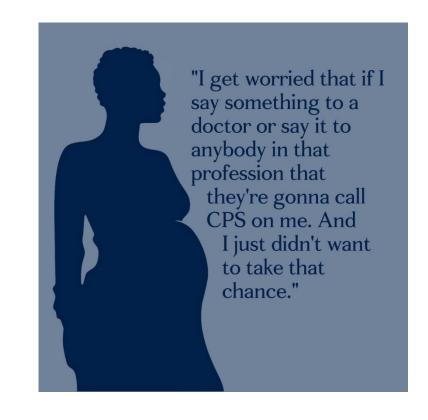
Screening Tool	Questions
NIDA Quick Screen- ASSIST	
Quick Screen [†]	In the past year, how often have you used the following? a. Five or more alcohol drinks in a day for men or 4 or more alcohol drinks in a day for women b. Tobacco products c. Prescription drugs for nonmedical reasons d. Illegal drugs
ASSIST*	 In your lifetime, which of the following substances have you used? (response options of yes or no) In the past 3 mo, how often have you used the substances you mentioned? (response options of never, once or twice, monthly, weekly, and daily or almost daily for items 2–5) In the past 3 mo, how often have you had a strong desire or urge to use (each substance)? (During the past 3 mo, how often has your use of (each substance) led to health, social, legal or financial problems?
	5. During the past 3 mo, how often have you failed to do what was normally expected of you because of your use of (each substance)?
	6. Has a friend or relative or anyone else ever expressed concern about your use of (each substance)? 7. Have you ever tried to control, cut down or stop using (each substance)? 8. Have you ever used any drug by injection?
SURP-P [§]	Have you ever used marijuana? How many alcoholic drinks have you consumed in the month before knowing you were pregnant? Do you feel the need to cut down on your alcohol or drug use?

4Ps+: Parents, Peers, (+Partner), Past, Present



Reporting in AZ

- AZ Appeals court ruled 2023 Arizona
 Department of Child Safety will no longer investigate reports of newborns testing positive for THC as long as the parent has a medical marijuana card
- Two thirds of study participants did not disclose cannabis use to their provider
 - Main fear was reporting to child protection services
- Call to revise DCS reporting policies and practices and improve provider education. Move away from punitive to treatment and support.





Cannabis use disorder treatment

WEED: Practice Mnemonic for Addressing Cannabis Use in Pregnancy

- Welcome questions about cannabis use
- Explore alternatives to cannabis for common pregnancy ailments, such as anxiety and nausea
- Explain the potential risks of cannabis use
- Deliver a harm-reduction message by recommending a decrease in the dose and frequency for patients who are not able or willing to remain abstinent during pregnancy



Cannabis use disorder (DSM-V)

1. Using more of a substance or longer than intended	Impaired control
2. Wanting to limit or stop using but not being able to	
3. Spending a lot of time using or recovering from use	
4. Craving	
5. Failure to fulfil duties in social roles (work, housekeeping)	Social impairments
6. Continue using despite substance-related social problems	
7. Giving up activities because of substance use	
8. Using in situations where using is dangerous to oneself or others	Risky use
9. Continue using despite substance-related mental or physical problems	
10. Tolerance	Pharmacological
11. Withdrawal*	criteria



Cannabis withdrawal

- Supportive care
- Behavioral approaches
 - Skills training, normalizing time line, coping skills, exercise, hot baths, avoiding caffeine
- Non-FDA comfort meds, small short term studies
 - Gabapentin
 - Quetiapine
 - Mirtazapine
 - Sedatives
 - Dronabinol, nabilone or nabiximol

TABLE 1 DSM-5 Diagnostic Criteria for Cannabis Withdrawal Syndrome (e6) Cessation of cannabis use that has been heavy and prolonged (i.e., usually daily or almost daily use over a period of a least a few months) 3 or more of the following signs and symptoms develop within approximately B 1 week of Criterion A - Irritability, anger or aggression Nervousness or anxiety - Sleep difficulty (insomnia, disturbing dreams) - Decreased appetite or weight loss - Restlessness Depressed mood - At least one of the following physical symptoms causing significant discomfort: abdominal pain, shakiness/tremors, sweating, fever, chills, or headache The signs or symptoms from criterion B cause clinically significant distress C. or impairment in social, occupational or other important areas of functioning. The symptoms are not due to another medical condition and are not better D. explained by another mental disorder.

DSM-5, Diagnostic and Statistical Manual of Mental Disorders



Cannabis use disorder treatment

- Psychosocial interventions (first line, moderate effect size, mod-high confidence in evidence)
 - Cognitive behavioral therapy
 - Motivational enhancement therapy
 - Contingency management
- Other modalities (unable to assess effect size, low confidence in evidence)
 - Social support counseling
 - Drug education counseling
 - Relapse prevention
 - Mindfulness
 - Mutual support programs (12 step, SMART, Refuge, Women for Sobriety, The Luckiest Club)



Cannabis use disorder treatment

- Biologic interventions (Non-FDA)
- Reduce cravings and relapse (limited small studies)
 - Gabapentin reduces cravings and relapse
 - N-acetylcysteine reduces relapse in adolescence population, effect not seen in adults
 - Naltrexone reduction in use



Conclusions

- Cannabis use is increasing in the US; As are cannabis and THC related medical, pregnancy, and psychiatric conditions
- Cannabis is associated with pregnancy risks, fetal complications, and neonatal risks
- Screen for cannabis in all pregnancies and provide education about risks
- Pregnant and lactating individuals should receive council to stop or reduce use to avoid pregnancy and neonatal complications and possible developmental problems
- Cannabis use disorder is managed with psychosocial support. There are no current FDA approved medications; however, some medications have shown promise in treating withdrawal, craving and preventing relapse.





Thank you! Questions?

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Sources

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