## Understanding Fetal, Neonatal and Infant Impacts of Parental Cannabis Use

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## Disclosure Information (Required)

### **Cannabis Use in Pregnancy: Person-Centered Perspectives**

Friday, April 1, 2022 10:30 AM – 11:30 AM

#### Davida Schiff

- Receives funding from the American Academy of Pediatrics for participation in the Maternal Infant Health and Opioid Use team
- \* Receives research funding and programmatic support from NIDA, SAMHSA, DOJ





#### **Outline**

- Understanding data (and limitations to existing data) on fetal and neonatal impacts of cannabis exposure in utero
- \*Lactation and cannabis
- \*Postpartum exposures and safety of use while parenting



## Cannabis use in pregnancy rarely occurs in isolation

- The data surrounding cannabis use and pregnancy is littered with confounders
- Cannabis has similar substances to tobacco smoke
- It is difficult to know the specific effects of cannabis on pregnancy and the fetus





**Open Access** Research

BMJ Open Prenatal exposure to cannabis and maternal and child health outcomes: a systematic review and meta-analysis

> J K L Gunn. 1 C B Rosales. 2 K E Center. 3 A Nuñez. 4 S J Gibson. 5 C Christ. 6 J E Ehiri5

Systematic reviews and meta-analyses assessing neonatal outcomes following prenatal cannabis exposure

Review

#### Maternal Marijuana Use and Adverse **Neonatal Outcomes**

A Systematic Review and Meta-analysis

Shayna N. Conner, MD, MSCI, Victoria Bedell, MD, Kim Lipsey, MLIS, George A. Macones, MD, MSCE, Alison G. Cahill, MD, MSCI, and Methodius G. Tuuli, MD, MPH

> DOI: 10.1111/1471-0528.15946 www.bjog.org

Systematic review



recreational cannabis use in high-income countries: a scoping review

S Singh, KB Filion, A,b,c HA Abenhaim, A,d MJ Eisenberg C,c,e

Prevalence and outcomes of prenatal

- Major confounders
  - \* Alcohol, tobacco, other substances
- Methodological issues
  - Freq and dose of exposures uncertain

Gunn, 2016; Singh, 2020; Conner 2016.

#ASAMAnnual2022

### Recent observational studies from Canada

JAMA | Original Investigation

JAMA. 2019;322(2):145-152. doi:10.1001/jama.2019.8734 Published online June 18, 2019.

## Association Between Self-reported Prenatal Cannabis Use and Maternal, Perinatal, and Neonatal Outcomes

Daniel J. Corsi, PhD; Laura Walsh, MSc; Deborah Weiss, PhD; Helen Hsu, MD; Darine El-Chaar, MD;

Steven Hawken, PhD; Deshayne B. Fell, PhD; Mark Walker, MD

**\***Corsi: Inc preterm birth

\*Koto: Lower BW







Maternal cannabis use during pregnancy and maternal and neonatal outcomes: A retrospective cohort study

Prosper Koto, Victoria M. Allen, John Fahey, Stefan Kuhle X

First published: 04 February 2022

https://doi-org.ezp-prod1.hul.harvard.edu/10.1111/1471-0528.17114



## Risks of exposure to cannabis



## Pregnancy

- Increased risk of preterm birth
- Most studies are confounded by concurrent tobacco use
- Maternal anemia



# Perinatal/Neonata

- Structural anomalies- mixed
- Small for gestational age mixed
- No increased risk of perinatal death
  - Possible increased risk of stillbirth



## Childhood

- Childhood behavior association
- Conflicting data on school performance
- Key Confounders
- Neonatal/childhood
   Prenatal vs. postnatal exposure
- Nutrition
- SES/social determinants
- Secondhand smoke exposure



## Putting prenatal cannabis findings into historical context

**EDITORIAL** 

Commentary \_\_\_\_

Cannabis Use in Pregnancy
A Tale of 2 Concerns

Michael Silverstein, MD, MPH; Elizabeth A. Howell, MD, MPH; Barry Zuckerman, MD

## The Problem of Prenatal Cocaine Exposure

A Rush to Judgment

Linda C. Mayes, MD Richard H. Granger, MD Marc H. Bornstein, PhD Barry Zuckerman, MD Silverstein JAMA, 2019

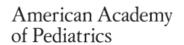


## Silverstein et al, 2019

- "It is impossible to separate data from the values that individuals bring to those data, no group is immune to the judgment of others, and women and minority groups (particularly pregnant women of color) tend to bear the greatest burden of many of these judgments."
- "More epidemiology is unlikely to completely resolve the complex issue of potentially safe moderate use or to completely remove the tendency to imbue data interpretation with implicit biases about groups of people."
- "Furthermore, the dialogue on cocaine was defined by exaggeration; so far, the dialogue on cannabis has largely been defined by a false perception of safety"



CLINICAL REPORT Guidance for the Clinician in Rendering Pediatric Care





DEDICATED TO THE HEALTH OF ALL CHILDREN"

Marijuana Use During Pregnancy and Breastfeeding: Implications for Neonatal and Childhood Outcomes

Sheryl A. Ryan, MD, FAAP, a Seth D. Ammerman, MD, FAAP, FSAHM, DABAM, Mary E. O'Connor, MD, MPH, FAAP, a COMMITTEE ON SUBSTANCE USE AND PREVENTION. SECTION ON BREASTFEEDING

- # "In summary, the evidence for independent, adverse effects of marijuana on human neonatal outcomes and prenatal development is limited, and inconsistency in findings may be the result of the potential confounding caused by the high correlation between marijuana use and use of other substances such as cigarettes and alcohol, as well as sociodemographic risk factors.
- \*However, the evidence from the available research studies indicate reason for concern, particularly in fetal growth and early neonatal behaviors.



## Neonatal withdrawal syndrome?

substance	symptoms	duration
cannabis	CNS symptoms (tone, startle, cry), decreased sleep, less regulated, altered NNNS scales	Self-resolve within 1 week (Ryan SA 2018)
tobacco	CNS (tone, excitability, asymmetrical reflexes, irritability, tremor) Altered GI/visual NNNS subscales	Begin at ~12 hours of life, last <48 hours (Law 2003, Garcia-Algar 2004)
SSRI/SNRIs	CNS symptoms, restlessness, feeding difficulties, fever, respiratory distress (rare: seizure, cyanosis)	Generally mild, self-resolve by 2 weeks (Leibovitch 2013, Moses-Kolko 2005)



## Is cannabis use a contraindication to breastfeeding?

Just how much cannabis is transferred in breastmilk?

\*What are the effects of ingested cannabis on newborns?

Do risks outweigh benefits?





## Official Recommendations

"There are insufficient data to evaluate the effects of marijuana use or infant data.



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[data] should prompt extremely careful consideration of the risks versus benefits of breastfeeding in the setting of moderate or chronic marijuana use."



## **AAP 2018 statement**

regarding the infant's long-term neurobehavioral development and thus are contraindicated."85 Although this has been interpreted by some professional organizations to indicate that in the parent using marijuana, the choice to breastfeed is "contraindicated," this was not the intent of that statement. It is suggested instead that the mother be encouraged to breastfeed while, at the same time, it is strongly encouraged that she abstain completely from using marijuana as well as other drugs, alcohol, and tobacco. This position has been supported by several other professional organizations and resources. For example, LactMed (a free



## Risks vs. well established benefits of breastfeeding

TABLE 2 Dose-Response Benefits of Breastfeeding<sup>a</sup>

Condition	% Lower Risk <sup>b</sup>	Breastfeeding
Otitis media <sup>13</sup>	23	Any
Otitis media <sup>13</sup>	50	≥3 or 6 mo
Recurrent otitis media <sup>15</sup>	77	Exclusive BF ≥6 mo <sup>d</sup>
Upper respiratory tract infection <sup>17</sup>	63	>6 mo
Lower respiratory tract infection <sup>13</sup>	72	≥4 mo
Lower respiratory tract infection <sup>15</sup>	77	Exclusive BF ≥6 mo <sup>d</sup>
Asthma <sup>13</sup>	40	≥3 mo
Asthma <sup>13</sup>	26	≥3 mo

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Condition	% Lower Risk <sup>b</sup>	Breastfeeding	Comments	0Rc	95% CI
			history		
RSV bronchiolitis <sup>16</sup>	74	>4 mo	_	0.26	0.074-0.9
NEC <sup>19</sup>	77	NICU stay	Preterm infants	0.23	0.51 - 0.94
			Exclusive HM		
Atopic dermatitis <sup>27</sup>	27	>3 mo	Exclusive BFnegative	0.84	0.59 - 1.19
			family history		
Atopic dermatitis <sup>27</sup>	42	>3 mo	Exclusive BFpositive	0.58	0.41 - 0.92
			family history		
Gastroenteritis13,14	64	Any	_	0.36	0.32 - 0.40
Inflammatory bowel	31	Any	_	0.69	0.51 - 0.94
disease <sup>32</sup>					
Obesity <sup>13</sup>	24	Any	_	0.76	0.67 - 0.86
Celiac disease <sup>31</sup>	52	>2 mo	Gluten exposure	0.48	0.40 - 0.89
			when BF		
Type 1 diabetes <sup>13-42</sup>	30	>3 mo	Exclusive BF	0.71	0.54 - 0.93
Type 2 diabetes <sup>13,43</sup>	40	Any	_	0.61	0.44 - 0.85
Leukemia (ALL) <sup>13,46</sup>	20	>6 mo	_	0.80	0.71-0.91
Leukemia (AML) 13.45	15	>6 mo	_	0.85	0.73 - 0.98
SIDS <sup>13</sup>	36	Any >1 mo	_	0.64	0.57-0.81



# individuals disproportionately due to cannabis use at five Massachusetts hospitals

	Nativ Am	Asian	Black NH	Hisp/ LatinX	Other	Unavailable	White NH
Any SUD related indication, (excluding cannabis)	30.0%	16.3%	12.9%	17.7%	25.0%	16.9%	45.8%
Cannabis	60.0%	34.9%	42.7%	40.2%	33.3%	28.2%	29.2%
Other indication (late PNC, maternal or infant clinical indication, rx monitoring, unknown)	10.0%	48.8%	44.4%	42.1%	41.7%	54.9%	25.0%



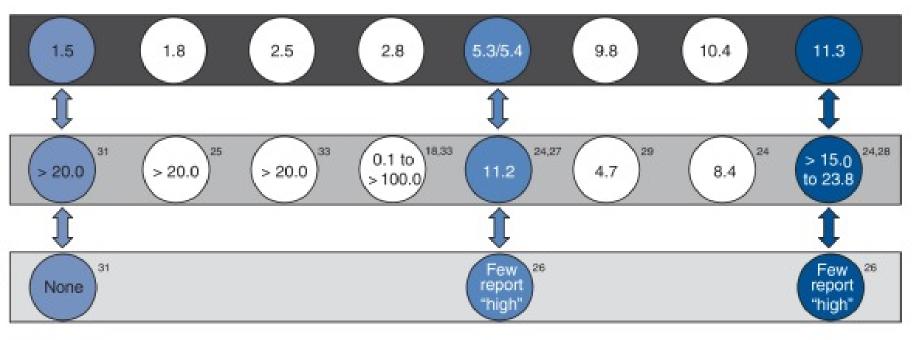
## Secondhand exposure

\*Ventilation, number of smokers, cannabis potency, air volume

Amount of THC in smoked cannabis, %

Amount of THC and metabolites in urine of participants passively exposed to cannabis smoke, ng/mL

Self-reported effects of cannabis in participants passively exposed to cannabis smoke





## Other considerations

- Safety/caregiver impairment
- Toxicology testing for cannabis



## What we know (and what we don't)

- Cannabis use in pregnancy is not uncommon
  - Legalization, conflicting messages
- Cannabis can be harmful to the fetus
  - But many confounders
- Cannabis is excreted in breastmilk and may confer harms to the infant/newborn
  - # But how much, for how long, and what harms are still unclear
- Equity concerns
  - Who uses cannabis? Who is being reported to DCF? Should cannabis use raise our suspicion for child abuse/neglect?



## Acknowledgements

\*Slides adapted from presentation prepared by Dr. Megan Miller and Dr. Leela Sarathy for MA PNQIN Spring Summit in 2021.



## References

- \* Conner, et al. Maternal marijuana use and adverse neonatal outcomes: a systematic review and meta-analysis. <a href="Obstet Gynecol">Obstet Gynecol</a> 2016;128:713–23.
- de Moraes Barros MC, Guinsburg R, de Araújo Peres C, Mitsuhiro S, Chalem E, Laranjeira RR. Exposure to marijuana during pregnancy alters neurobehavior in the early neonatal period. J Pediatr. 2006 Dec;149(6):781-7.
- Holitzki H, Dowsett LE, Spackman E et al. "Health effects of exposure to second- and third-hand marijuana smoke: a systematic review." CMAJ Open. 2017 Oct-Dec; 5(4): E814–E822.
- Ryan SA, Ammerman SD, O'Connor ME; AAP Committee on Substance Use and Prevention; AAP Section on Breastfeeding. Marijuana Use During Pregnancy and Breastfeeding: Implications for Neonatal and Childhood Outcomes. *Pediatrics*. 2018;142(3):e20181889
- \* Section on Breastfeeding. Breastfeeding and the use of human milk. Pediatrics. 2012;129(3).

