

Marijuana use in pregnancy and lactation: a review of the evidence

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Marijuana is the most frequently used illicit drug in Western countries.¹ In 2013, 19.8 million, or 7.5% of the US population, reported its use within the last month, an increase from 2007 when only 5.8% of the population had used marijuana within the past month.^{2,3}

Reported prevalence rates of marijuana use in pregnancy vary from as low as 3% to as high as 34%.^{4,5} We anticipate an increase in marijuana use in pregnancy as legalization of marijuana increases throughout the United States. This review is intended to provide practicing clinicians with an understanding of existing literature and recommendations for managing women who use marijuana during pregnancy because this will be an increasingly encountered clinical scenario.

The term marijuana is used throughout this article to represent cannabis use globally. Technically the active psychogenic component of marijuana is a cannabinoid called delta-9-tetrahydrocannabinol (THC).⁶

Search methodology

Ovid Medline (PubMed) and Embase were searched on Dec. 11, 2014, for relevant articles. A focused search was conducted with the search terms marijuana and marihuana or cannabinoids and pregnancy, lactation, and outcomes

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With the legalization of recreational marijuana in many states, we anticipate more women will be using and self-reporting marijuana use in pregnancy. Marijuana is the most common illicit drug used in pregnancy, with a prevalence of use ranging from 3% to 30% in various populations. Marijuana freely crosses the placenta and is found in breast milk. It may have adverse effects on both perinatal outcomes and fetal neurodevelopment. Specifically, marijuana may be associated with fetal growth restriction, stillbirth, and preterm birth. However, data are far from uniform regarding adverse perinatal outcomes. Existing studies are plagued by confounding by tobacco and other drug exposures as well as sociodemographic factors. In addition, there is a lack of quantification of marijuana exposure by the trimester of use and a lack of corroboration of maternal self-report with biological sampling, which contributes to the heterogeneity of study results. There is an emerging body of evidence indicating that marijuana may cause problems with neurological development, resulting in hyperactivity, poor cognitive function, and changes in dopaminergic receptors. In addition, contemporary marijuana products have higher quantities of delta-9-tetrahydrocannabinol than in the 1980s when much of the marijuana research was completed. The effects on the pregnancy and fetus may therefore be different than those previously seen. Further research is needed to provide evidence-based counseling of women regarding the anticipated outcomes of marijuana use in pregnancy. In the meantime, women should be advised not to use marijuana in pregnancy or while lactating.

Key words: cannabis, lactation, marijuana, pregnancy

including adverse perinatal outcomes and neurodevelopment. A search without any language or year limits yielded 615 unique citations. Abstracts were reviewed by the authors, and all pertinent articles were obtained and reviewed individually. In addition, references of pertinent articles were reviewed to find any additional articles that were not identified with the initial search ($n = 43$). All pertinent articles are summarized here. Our goal was not to provide a systematic review of a specific research question but rather to provide practicing clinicians with a comprehensive overview of the existing marijuana in pregnancy and lactation literature.

Legalization of marijuana

Currently both recreational and medical marijuana remain illegal by federal law in the United States. However, the legalization of medical and recreational marijuana at the state level is increasing

throughout the United States. At this point, 23 states have legalized the use of medical marijuana, and 4 states have legalized both medical and recreational marijuana (Figure).

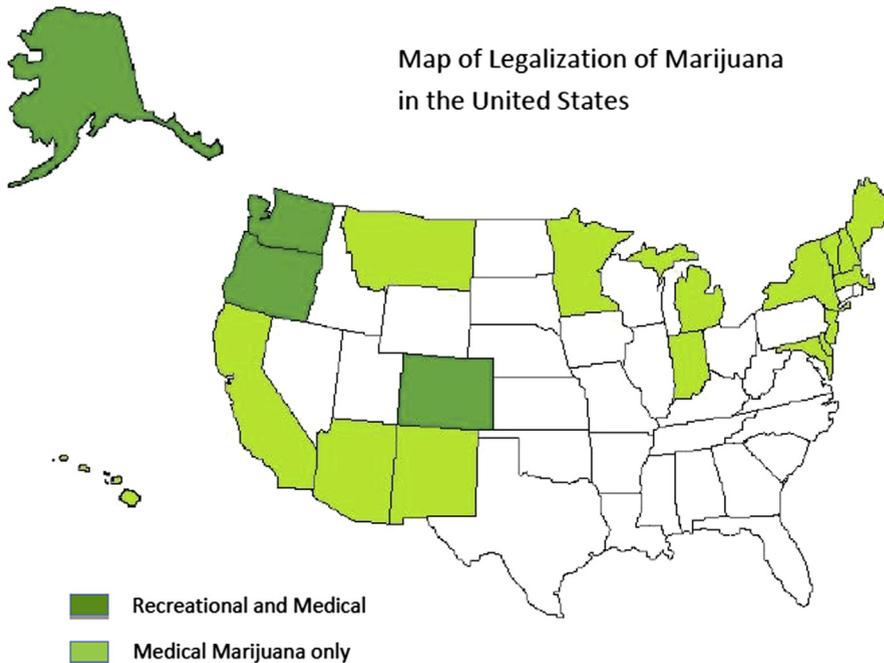
The Colorado experience

Medical marijuana was legalized in Colorado in the year 2000. However, it was not until 2009 when the US Attorney General issued a statement passing the jurisdiction of marijuana law enforcement to state governments that we saw a sharp increase in the number of medical marijuana users in the state.⁷ In 2012, recreational marijuana was legalized in the state of Colorado with the passing of Amendment 64. There is no stipulation in the law stating that pregnant women cannot purchase or possess marijuana.

Sales of recreational marijuana have been steadily increasing since the opening of the first recreational dispensaries on

FIGURE

An increasing number of states in the United States have legalized both medicinal and recreational marijuana use



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Jan. 1, of 2014. The state of Colorado does not publish overall sales amounts but does publish tax revenue on a monthly basis. In January 2014, the revenue was 3.5 million dollars. The monthly tax revenue is now up to 7.6 million dollars for the month of October 2014, showing a steady increase in sales and consumption.⁸ In addition, there has been an increase in the use of alternative forms of consumption such as vaping (heating the cannabis to release THC and cannabinoids without making it smoke), lotions, and edibles.^{7,9}

Following the legalization of marijuana, we have noted several unanticipated adverse consequences of the increase in marijuana availability including an increase in pediatric overdoses and emergency visits for marijuana toxicity.⁷

Attitudes and beliefs

When women have been followed up longitudinally during pregnancy, a decrease in marijuana use has been noted across trimesters of pregnancy. In a 1 year

prospective cohort study, marijuana use in pregnancy declined from 32% in the first trimester to 16% in the third trimester.⁵

Similarly, a longitudinal prospective study on drug use in pregnancy (n = 86), the Development and Infancy Study, found that the percentage of women who used marijuana throughout the pregnancy declined. However, approximately 60% of women who used marijuana in the year prior to pregnancy continued to use more than 10 joints per week, indicating that many women continue use throughout pregnancy.¹⁰ It should be noted that the women in the Development and Infancy Study¹⁰ smoked an average of 21 joints per week in the month prior to pregnancy and may not be representative of less frequent users of marijuana.

Two thirds of adults surveyed in a UK study noted that cannabis was either “not very harmful” or “not at all harmful.”¹¹ This is in contrast to other recreational drugs such as heroin or cocaine in which less than 5% of adults surveyed

perceived them to be either “not very harmful” or “not at all harmful.”¹¹ The perceived safety likely contributes to the high prevalence of its use in pregnancy.

Screening and testing for marijuana use

The American College of Obstetricians and Gynecologists and the American Academy of Pediatrics support screening all women for drug use at the time of entry to prenatal care.¹² Verbal screening for self-reported use was noted to be acceptable to patients in one study.¹³ Women who report use should then be encouraged to stop and referred to local substance use disorder programs if needed.

Unfortunately, maternal and fetal testing for marijuana exposure is fraught with error.¹⁴ Maternal urine testing is the most accurate method of testing. However, the duration of a positive urine toxicology result from the last use depends on many factors including chronicity of use (Table 1). Despite its limitations, urine is easy to obtain, has a high concentration of metabolites, and is therefore the preferred method of screening.⁶

Testing of maternal hair samples is inaccurate and may remain positive despite no recent use. Neonatal hair and meconium can also be tested (Table 1); however, because of the cost of testing, delay in results, and a high false-positive rate in laboratory testing of meconium by different techniques,¹⁵ neonatal testing is rarely used in clinical practice.

There are no good methods to quantify the amount of marijuana ingested using biological sampling in a clinical setting. The amount of THC in various forms of marijuana varies by the extraction process from the plant, *Cannabis sativa*, which also results in challenges in quantifying self-reported use. In addition, the various forms of consumption result in different rates of absorption and peak blood concentrations.⁷ In a report from the University of Mississippi’s Potency Monitoring Project, the average concentration of THC in seized samples in the United States in 2008 was 13.0%, which was an increase from 3.2% in 1983.¹⁶

Nausea and vomiting in pregnancy

As with any drug or medication in pregnancy, possible benefits must be weighed against possible adverse effects. There are few data on the possible benefits of marijuana use in pregnancy. Interest in the use of marijuana as an antiemetic has been propagated by its efficacy in oncology patients.^{17,18}

There are 2 studies investigating the relationship between marijuana use and nausea and vomiting of pregnancy.^{19,20} Roberson et al²⁰ used the pregnancy risk assessment monitoring system data (n = 4375) and found that women who used marijuana in pregnancy were more likely to report severe nausea (3.7% vs 2.3%; prevalence ratio, 1.63; 95% confidence interval [CI], 1.08–2.44). The treatment of nausea with marijuana was not specifically addressed in the study.

Westfall et al¹⁹ reported on the prevalence of nausea among 79 women who used medicinal marijuana in pregnancy. Forty of these women (51%) used marijuana to treat nausea and vomiting of pregnancy, and 92% of them believed it was effective. There was no control group, no documentation of quantity used, or a demonstration of effect on symptoms of nausea other than subjective report by survey after the pregnancy.

In summary, the effect of marijuana use on nausea and vomiting of pregnancy is unknown.

Anesthetic considerations

Marijuana use can affect the safety and administration of anesthesia surrounding delivery. In high doses, marijuana can cause bradycardia and hypotension, but more commonly, low or moderate doses can cause tachycardia.²¹ If tachycardia is present or marijuana use is suspected, drugs that increase heart rate such as ketamine, pancuronium, and epinephrine should be avoided. Because marijuana is often inhaled, it can also cause upper-airway irritation and edema, making anesthetic administration more complicated.²¹

Adverse perinatal outcomes

Fetal growth

Many of the human studies of marijuana in pregnancy focus on fetal growth

TABLE 1
Testing for marijuana in biological samples

Biological sample	Duration of positive result	Test limitations
Maternal urine	2–3 days in occasional users ⁶³ Several weeks in chronic users ⁶⁴	Chronicity of use determines duration of positive result ⁶³
Maternal serum	2–3 days in occasional users ⁶ Several weeks in chronic users ⁶	Chronicity of use determines duration of positive result ⁶³ Invasive sample Shorter half-life than urine ⁶
Maternal hair	Several weeks ⁶⁵	Less accurate for marijuana than other drugs ⁶⁵ False positives from passive exposure ⁶⁵ Not clinically used due to cost and inaccuracy
Meconium	Positive result indicates second- and third-trimester exposure ^{26,66,67}	Small amount of detectable THC in the samples ⁶⁸ High false-positive rate (up to 43%) ¹⁵ Send out to reference laboratory Costly and impractical at many sites
Neonatal hair	Positive result indicates third-trimester exposure ⁶⁶	Costly and impractical at many sites Less sensitive than meconium ⁶⁶

THC, delta-9-tetrahydrocannabinol.

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(Table 2).^{4,22-35} Abnormalities in growth are biologically plausible, given the passage of cannabinoids across the placenta. There are some data suggesting that cannabis affects glucose and insulin regulation and therefore may affect the fetal growth trajectory.²⁵ However, data regarding fetal growth with marijuana exposure are mixed, with some studies demonstrating a decrease in birthweight and/or growth and others demonstrating no association (Table 2). In part, the controversy may be a result of differing methodology for the ascertainment of marijuana exposure, varying from a single question about self-reported use at study entry to detailed longitudinal frequency of use data and biological sampling (Table 2). In addition, many early studies did not account for concurrent exposure to tobacco.³⁶

A metaanalysis by English et al³⁶ (1997) focused on the association between marijuana exposure and birthweight. This metaanalysis included 10 studies in which the investigators adjusted for the effect of tobacco exposure. Whereas women who consumed

large quantities of marijuana (more than 4 times per week) had babies that weighed less than nonusers by 131 g on average, the pooled odds ratio for low birthweight with any marijuana use was 1.09 (95% CI, 0.94–1.27).³⁶

Most of the trials included in the metaanalysis utilized self-report of marijuana use as the predictor of low birthweight rather than biological sampling. Zuckerman et al³⁵ found an association between a positive urine toxicology screen for THC and lower birthweight (79 g decrease in birthweight, $P = .04$). However, there was no observed association when only self-report was considered. These authors argued that a lack of association between marijuana and lower birthweight in other studies may have been a result of incomplete ascertainment of exposure by relying on self-report.³⁵

There is only one study, a secondary analysis of the Generation R data (a large prospective trial to assess early environmental and genetic determinants of health in The Netherlands), in which serial ultrasounds were performed to

TABLE 2
Summary of marijuana and fetal growth restriction studies

Study and number in cohort	Marijuana-exposed women, n (%)	Setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs or regression coefficients with 95% CIs reported when available)	Limitations and comments
Prospective cohort studies ^a							
Day et al, 1991 ³² (n = 519)	324 (62)	Single institution	Self-report by prenatal interview in each trimester of pregnancy	Frequency: light (0–2.9 joints/wk), moderate (3–6.9/wk), and heavy (≥1 joints/d)	SES, obstetric hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with SGA Isolated higher birthweight in heavy third-trimester users compared with nonusers (3357 g vs 3215 g; <i>P</i> = .04)	Increase in birthweight in marijuana users compared with nonusers Women who use marijuana were intentionally oversampled
El Marroun et al, 2009 ²⁵ (n = 7452)	459 (6)	Population-based study in The Netherlands	Self-report at study enrollment	Frequency: daily, weekly, monthly Reported use: only before pregnancy, use in early pregnancy, or ongoing use	Standard demo, psych hx, EtOH, fetal sex, tobacco Excluded women with other drugs	Use before pregnancy did not affect growth Early pregnancy use decreased growth 11.18 g (–15.26 to –7.10)/wk Ongoing marijuana use decreased growth 14.44 g (–22.94 to –5.94)/wk	Only study with serial ultrasounds to assess fetal growth (detailed in fetal growth section of text) Marijuana use not well quantified
Fergusson et al, 2002 ²² (n = 12,129)	606 (5)	Population-based study in United Kingdom	Self-completed questionnaire at 18–20 wks gestation	Frequency: 1 time/day, 2–4 time/wk, 1 time/wk, <1 time/wk before pregnancy, first trimester and ongoing	Standard demo, other drugs, EtOH, tobacco	Ongoing use 1 or more/wk throughout pregnancy was not associated with lower birthweight –84.20 g (–174.70 to 6.40)	Self-report data collected at 18–20 wks' gestation, no later pregnancy data
Fried et al, 1984 ³³ (n = 583)	84 (14)	Referred to study by primary obstetrician/study ads	Self-report by prenatal interview in each trimester of pregnancy	Frequency: irregular users (≤1 joint/wk), moderate (2–5/wk), heavy (>5/wk)	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with LBW	Marijuana use not well quantified, averaged over the course of pregnancy
Gray et al, 2010 ²⁶ (n = 86)	38 (44)	Single institution	Self-report by prenatal interview in each trimester of pregnancy Biological samples	Frequency: number of joints/day by trimester Presence of THC in maternal saliva and meconium	Standard demo, OB hx (parity only), tobacco Excluded women with other drugs, or heavy EtOH	THC in meconium associated with lower birthweight (3429 g vs 2853 g; <i>P</i> < .001), persistent effect in multivariable logistic regression Self-report alone was not associated with lower birthweight	Study designed to assess tobacco exposure primarily Sampling strategy for high prevalence of use not reported

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(continued)

TABLE 2
Summary of marijuana and fetal growth restriction studies (continued)

Study and number in cohort	Marijuana-exposed women, n (%)	Setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs or regression coefficients with 95% CIs reported when available)	Limitations and comments
Hatch et al, 1986 ⁶⁹ (n = 3857)	366 (10)	Planned delivery at single institution	Self-report by structured interview early in pregnancy	Frequency: none, occasional (≤ 1 times/mo), regular (≥ 2 times/mo)	OB hx, standard demo, other drugs, EtOH, tobacco	Regular use in white women associated with LBW (OR, 2.6; 95% CI, 1.1–6.2) Regular use in white women associated with SGA (OR, 2.3; 95% CI, 1.3–4.1)	Self-report data collected early in pregnancy, no later pregnancy data Differing results by racial group Marijuana use not well quantified
Hingson et al, 1982 ³⁴ (n = 1690)	237 (14)	Single institution	Self-report by structured interview postpartum	Frequency: < 1 time/mo, < 1 /wk, 1–2 times/wk, ≥ 3 per week	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Neonates 95 g smaller than nonusers with use < 3 times/wk ($P < .01$) Neonates 139 g smaller than nonusers with use ≥ 3 times/wk ($P < .01$)	Possible recall bias, most exposure data collected postpartum, small subset with prenatal interview (n = 328) Marijuana use not well quantified
Hurd et al, 2005 ²⁷ (n = 139)	44 (32)	Women undergoing elective termination at a single center at 17–22 wks	Self-report by structured interview at time of termination Biological samples	Frequency: light (0–0.4 joints/d), moderate (0.41–0.88/d), and heavy (≥ 0.89 joints/d), THC in maternal urine or meconium	Standard demo, gestational age at termination, EtOH, tobacco Excluded women with cocaine/opiates	Increasing self-reported use not associated with decreasing weight Decreased birthweight in marijuana-exposed (either urine/meconium positive toxicology or self-report) fetuses by 14.53 g (–28.21 to –0.86)	Growth assessed in midgestation prior to presentation of most growth abnormalities Women in study were undergoing elective termination of pregnancy
Kliegman et al, 1994 ⁷⁰ (n = 425)	34 (8)	Single institution	Self-report by structured interview at time of delivery Biological samples	THC in maternal urine at time of delivery	SES, OB hx, standard demo, other drugs, EtOH, tobacco	No association with LBW (OR, 2.28; 95% CI, 0.27–19.5)	Study designed to assess cocaine exposure primarily Marijuana use not quantified
Linn et al, 1983 ⁷¹ (n = 12,424)	1246 (10)	Single institution	Self-report by structured interview postpartum	Frequency: occasional, weekly or daily use	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with LBW for any use of marijuana (OR, 1.07; 95% CI, 0.87–1.31)	Possible recall bias, exposure data collected postpartum Marijuana use not well quantified

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TABLE 2
Summary of marijuana and fetal growth restriction studies (continued)

Study and number in cohort	Marijuana-exposed women, n (%)	Setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs or regression coefficients with 95% CIs reported when available)	Limitations and comments
Tennes et al, 1985 ⁵ (n = 756)	257 (34)	Two affiliated institutions	Self-report by structured interview at 1 prenatal visit and postpartum	Frequency quantified by trimester: light (≤ 1 times/wk), moderate (>1 time/wk but <1 time/d), heavy (≥ 1 times/d)	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No effect on birthweight when considered by trimester or as a total amount consumed during pregnancy	Possible recall bias, only 2 sessions of self-report, which was then reported by trimester of use Marijuana use not well quantified
Zuckerman et al, 1989 ³⁵ (n = 1226)	331 (27)	Single institution	Self-report by structured interview at 1 prenatal visit and postpartum Biological sampling	Reported use: yes/no THC in maternal urine at time of prenatal or postpartum interview	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Positive urine toxicology screen for THC associated with 79 g decrease in birthweight ($P = .04$) No association when only self-report considered	Possible recall bias, only 2 sessions of self-report Marijuana use not well quantified
Secondary analysis of prospective cohort ^a							
Bada et al, 2005 ²⁸ (n = 8637)	812 (9)	Multicenter, 4 university-based centers	Self-report by structured interview prior to delivery	Reported use: yes/no	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with LBW (OR, 1.08; 95% CI, 0.85–1.36) or SGA (OR, 0.9; 95% CI, 0.73–1.11)	Not designed to assess marijuana specifically (Maternal Lifestyle Study ⁷²) Marijuana use not quantified
Gibson et al, 1983 ⁷³ (n = 7301)	392 (5)	Two affiliated institutions	Self-report by structured interview at 1 prenatal visit and postpartum	Frequency: ≤ 1 times/wk, >1 time/wk	Standard demo, OB hx (parity only), EtOH, tobacco	No association with LBW after excluding premature neonates	Marijuana use not well quantified
Janisse et al, 2014 ²⁹ (n = 3090)	748 (24)	Single institution	Self-report by structured interview at each prenatal visit	Proportion of prenatal visits with reported use: 1–33%, 34–66%, or 67–100%	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	55 g decrease in fetal growth with ongoing marijuana use (reported at 67–100% of visits) ($P < .004$)	Study designed to assess EtOH exposure Population limited to African Americans Marijuana use not well quantified

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TABLE 2
Summary of marijuana and fetal growth restriction studies (continued)

Study and number in cohort	Marijuana-exposed women, n (%)	Setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs or regression coefficients with 95% CIs reported when available)	Limitations and comments
Kline et al, 1987 ⁷⁴ (n = 2815)	275 (10)	Two overlapping prospective cohorts at 3 urban hospitals	Self-report by structured interview at 1 prenatal visit	Frequency: <1 time/mo, 2–3 times/mo, 2–3 times/wk, 4–6 times/wk, daily	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with FGR in early cohort Decreased growth with increased use (127 g less with 2–3 times/wk, 143 g less with 4–6 times/wk and 230 g less with daily) in late cohort	Study designed as a case-control study with SAB as primary outcome Differing results for 2 overlapping prospective cohorts Marijuana use not well quantified
Saurel-Cubizolles et al, 2014 ⁴ (n = 13,545)	156 (1)	Population-based study, all births in France during a single week	Self-report by structured interview 2–3 d postpartum	Frequency: <1 time/mo, 1–9 times/mo, ≥10 times/mo	SES, standard demo, EtOH, tobacco	No association with SGA for <1 time/mo use (OR, 1.29; 95% CI, 0.61–2.72) or for use ≥1 times/mo use compared with nonusers (OR, 1.30; 95% CI, 0.66–2.56) Also no association with SGA for non-tobacco users, marijuana only	Recall bias Low prevalence of use concerning for ascertainment bias for marijuana exposure Marijuana use not well quantified
Shiono et al, 1995 ⁴⁰ (n = 7470)	822 (11)	Multicenter, 7 university-based clinics	Self-report by structured interview at 1 prenatal visit Biological samples	Frequency: number of times/wk THC in maternal serum	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with LBW when marijuana use assessed by self-report or positive serum assay for THC (OR, 1.1; 95% CI, 0.9–1.5) Increased odds of LBW with positive serum assay in isolation but not with self-report	Study designed to assess association between vaginal infections and PTB Marijuana use not well quantified
Teitelman et al, 1990 ⁷⁵ (n = 1206)	95 (8)	Planned delivery at single institution	Self-report by structured interview early in pregnancy	Reported use: yes/no	OB hx, standard demo, other drugs, EtOH, tobacco	No association with LBW (OR, 1.57; 95% CI, 0.54–4.52)	Study designed to assess associations between maternal work activity and LBW Same cohort as Hatch et al ⁶⁹ study with different inclusion criteria (employed women) No quantification of marijuana use

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TABLE 2
Summary of marijuana and fetal growth restriction studies (continued)

Study and number in cohort	Marijuana-exposed women, n (%)	Setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs or regression coefficients with 95% CIs reported when available)	Limitations and comments
van Gelder et al, 2010 ³⁰ (n = 5871)	189 (3)	Population-based, US National Birth Defects Prevention Study	Self-report by structured interview 6 wks to 24 mo after delivery	Reported use: yes/no by trimester	SES, OB hx, medical hx, standard demo, other drugs, ETOH, tobacco	No association with LBW (OR, 0.7; 95% CI, 0.3–1.6) No difference in mean birthweight (–17 g; P = .65) No difference by trimester of use	Recall bias, interviews up to 2 years postpartum Not designed for marijuana exposure specifically (birth defects registry) Marijuana use not well quantified

CI, confidence interval; ETOH, alcohol use; FGR, fetal growth restriction (estimated fetal weight less than the 10th percentile); LBW, low birthweight (defined <2500 g unless otherwise noted); Medical hx, medical history; OB hx, obstetrical history; OR, odds ratio; PTB, preterm birth (<37 weeks); SAB, spontaneous abortion; SES, socioeconomic status; SGA, small for gestational age (birthweight less than the 10th percentile); Standard demo, some measure of standard demographics including maternal age, race, body mass index; THC, delta-9-tetrahydrocannabinol.

^a Studies that did not adjust for tobacco use and retrospective cohorts are not included in this summary table.
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assess fetal growth rather than using birthweight as the outcome.²⁵ Women were followed up prospectively with growth ultrasounds at less than 18 weeks, 18-25 weeks, and 25 weeks or longer. Marijuana exposure data were by self-report. Fetuses exposed to marijuana in early pregnancy (n = 214) grew 11.2 g/wk less than the nonusers, and this effect was more pronounced in women with continued use throughout pregnancy (Table 2). The long-term effects of this small growth decrement are unknown.

A small subset of the Generation R cohort had Doppler ultrasound scans performed between 28 and 34 weeks.³⁷ Women with cannabis use in early pregnancy (n = 14) and women with ongoing cannabis throughout pregnancy (n = 9) were compared with nonusers. There was no difference in the umbilical artery pulsatility index or fetal cerebral blood flow. Women who used cannabis throughout pregnancy had a higher uterine artery pulsatility index and resistance index than nonusers after adjusting for fetal weight, fetal sex, and maternal education. The authors appropriately cautioned against drawing widespread conclusions from these data, given the small sample size.

It should be noted that in contrast to all other studies finding a small growth decrement or no difference in birthweight, one prospective study noted an increase in birthweight among neonates exposed to heavy use of marijuana (more than 1 joint/day) in the third trimester (3357 g v 3215 g; P = .04).³² This finding has not been replicated in any other studies and was not demonstrated with use in other trimesters.

In summary, there may be a small decrease in growth with exposure to marijuana in pregnancy. However, the clinical significance of this decrease is questionable, with reported growth differences on the order of 100 g.

Stillbirth

Many of the prior studies of marijuana in pregnancy exclude women with a stillbirth, so data regarding stillbirth and marijuana use are scant. However, a recent case-control study by Varner et al³⁸

TABLE 3
Summary of marijuana and preterm birth studies

Study and number in cohort	Marijuana-exposed women, n, %	Study design and setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs with 95% CIs reported when available)	Limitations and comments
Prospective cohort ^a							
Day et al, 1991 ³² (n = 519)	324 (62)	Single institution	Self-report by prenatal interview in each trimester of pregnancy	Frequency: light (0–2.9 joints/wk), moderate (3–6.9/wk), and heavy (≥ 1 joints/d)	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No effect on length of gestation No association with PTB	Women who use marijuana were intentionally oversampled
Dekker et al, 2014 ³⁹ (n = 3184)	213 (7) with pre-pregnancy exposure	International multicenter	Self-report by structured interviews at 20 wks	Timing of use: before pregnancy, first trimester	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Prepregnancy use associated with spontaneous PTB with intact membranes (OR, 2.34; 95% CI, 1.22–4.52)	Study designed to develop screening tests for PTB and other adverse obstetrical outcomes Marijuana use not quantified
Fried et al, 1984 ³³ (n = 583)	84 (14)	Referred to study by primary obstetrician/study ads	Self-report by prenatal interview in each trimester of pregnancy	Frequency: irregular users (≤ 1 joints/wk), moderate (2–5 joints/wk), heavy (> 5 joints/wk)	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Heavy use of marijuana reduced the length of gestation by 0.8 wks ($P = .008$) Increasing use associated with decreasing length of gestation	Marijuana use not well quantified by trimester, averaged over the course of pregnancy
Hatch et al, 1986 ⁶⁹ (n = 3857)	366 (10)	Planned delivery at single institution	Self-report by structured interview early in pregnancy	Frequency: none, occasional (≤ 1 times/mo), regular (≥ 2 times/mo)	OB hx, standard demo, other drugs EtOH, tobacco	Use associated with increased rate of PTB (< 37 wks) in white women (OR, 1.9; 95% CI, 1.0–3.9) No association with PTB in women of other races	Marijuana use not well quantified, especially for more frequent users Self-report data collected early in pregnancy, no later pregnancy data
Kliegman et al, 1994 ⁷⁰ (n = 425)	34 (8)	Single institution	Self-report by structured interview at time of delivery Biological samples	THC in maternal urine at time of delivery	SES, OB hx, standard demo, other drugs, EtOH, tobacco	No association with PTB (OR, 1.89; 95% CI, 0.34–10.50)	Study designed to assess cocaine exposure primarily Marijuana use not quantified

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TABLE 3
Summary of marijuana and preterm birth studies (continued)

Study and number in cohort	Marijuana-exposed women, n, %	Study design and setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs with 95% CIs reported when available)	Limitations and comments
Linn et al, 1983 ⁷¹ (n = 12,424)	1246 (10)	Single institution	Self-report by structured interview postpartum	Frequency: occasional, weekly, or daily use	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with PTB (OR, 1.02; 95% CI, 0.82–1.27)	Possible recall bias, exposure data collected postpartum Marijuana use not well quantified
Tennes et al, 1985 ⁵ (n = 756)	257 (34)	Two affiliated institutions	Self-report by structured interview at 1 prenatal visit and postpartum	Frequency quantified by trimester: light (≤ 1 times/wk), moderate (> 1 time/wk but < 1 times/d), heavy (≥ 1 times/d)	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No odds ratio reported for PTB (0% PTB rate in ≥ 3 times/wk users and 7% in nonusers) Total marijuana use in pregnancy positively correlated with increased gestational age at birth (r = 0.10), average of 2 d longer gestation with daily use	Possible recall bias, only 2 sessions of self-report No PTB (0%) in the nonusers as comparison group Finding of longer length of gestation not replicated in other human studies
Secondary analysis of a prospective cohort ^a							
Bada et al, 2005 ²⁸ (n = 8637)	812 (9)	Multicenter, 4 university-based centers	Self-report by structured interview prior to delivery	Reported use: yes/no	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with PTB (OR, 1.21; 95% CI, 0.9–1.61)	Not designed to assess marijuana specifically (Maternal Lifestyle Study ⁷²) Marijuana use not quantified
Gibson et al, 1983 ⁷³ (n = 7301)	392 (5)	Two affiliated institutions	Self-report by structured interview at 1 prenatal visit and postpartum	Frequency: ≤ 1 times/wk, > 1 time/wk	Standard demo, OB hx (parity only), EtOH, tobacco	High proportion of PTB among > 1 time/wk users (25% vs 6% in nonusers; $P < .001$)	Marijuana use not well quantified
Janisse et al, 2014 ²⁹ (n = 3090)	748 (24)	Single institution	Self-report by structured interview at each prenatal visit	Proportion of prenatal visits with reported use: 1–33%, 34–66%, or 67–100%	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Not associated with PTB	Study designed to assess EtOH exposure primarily Population limited to African Americans Marijuana use not well quantified

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(continued)

TABLE 3
Summary of marijuana and preterm birth studies (continued)

Study and number in cohort	Marijuana-exposed women, n, %	Study design and setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs with 95% CIs reported when available)	Limitations and comments
Saurel-Cubizolles et al, 2014 ⁴ (n = 13,545)	156 (1)	Population-based study, all births in France during a single week	Self-report by structured interview 2-3 d postpartum	Frequency: <1 time/mo, 1-9 times/mo, ≥10 times/month	SES, standard demo, EtOH, tobacco	Any marijuana use associated with spontaneous PTB (OR, 2.15; 95% CI, 1.10–4.18) No association with PTB when only women with marijuana use and no concurrent tobacco use were analyzed (OR, 1.22; 95% CI, 0.29–5.06)	Recall bias Low prevalence of use concerning for ascertainment bias for marijuana exposure Marijuana use not well-quantified
Shiono et al, 1995 ⁴⁰ (n = 7470)	822 (11)	Multicenter, 7 university-based clinics	Self-report by structured interview at 1 prenatal visit, biological samples	Frequency: number of times per week THC in maternal serum	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with PTB when marijuana use assessed by self-report or positive serum assay for THC (OR, 1.1; 95% CI, 0.8–1.3) Increased odds of PTB with positive serum assay in isolation but not with self-report	Study designed to assess association between vaginal infections and PTB Marijuana use not well quantified
van Gelder et al, 2010 ³⁰ (n = 5871)	189 (3)	Population-based, US National Birth Defects Prevention Study	Self-report by structured interview 6 wks to 24 mo after delivery	Reported use: yes/no by trimester	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	No association with PTB (OR, 1.0; 95% CI, 0.6–1.9) No difference by trimester of use	Recall bias, interview up to 2 y postpartum Not designed for marijuana exposure specifically (birth defects registry) Marijuana use not well quantified

CI, confidence interval; EtOH, alcohol use; Medical hx, medical history; OB hx, obstetrical history; OR, odds ratio; PTB, preterm birth (<37 wks); SAB, spontaneous abortion; SES, socioeconomic status; Standard demo, some measure of standard demographics including maternal age, race, body mass index; THC, delta-9-tetrahydrocannabinol.

^a Studies that did not adjust for tobacco use and retrospective cohorts are not included in this summary table.

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in the Stillbirth Collaborative Research Network demonstrated an increased risk of stillbirth among women who used marijuana in pregnancy as demonstrated by THC in the umbilical cord homogenate (odds ratio [OR], 2.34; 95% CI, 1.13–4.81).

These data are valuable, given their cross-sectional nature, diverse population, and objectivity. However, there are limitations including lack of quantification and timing of marijuana use. In addition, the authors noted concern for possible residual confounding by tobacco use, which attenuated the association between THC in the cord homogenate and stillbirth by approximately 10%, thereby further stressing the importance of accounting for concurrent tobacco use in marijuana research.

Preterm birth

Data on the association between marijuana use and preterm birth are mixed, with some studies demonstrating an increased risk of preterm birth and others demonstrating no association (Table 3). This is likely a result of differing methodological approaches including poor quantification of marijuana exposure and a lack of documentation of the indication for preterm birth in many studies (Table 3). Only 2 studies specify an outcome of spontaneous preterm birth^{4,39} rather than a generic outcome of any preterm birth (<37 weeks).

There are 2, large retrospective, population-based Australian studies supporting an increased risk of preterm birth with marijuana use. The first was a cohort (n = 24,874) who self-reported marijuana use at their intakes for prenatal care. After adjusting for alcohol, tobacco, and other illicit drugs, marijuana use was associated with preterm birth (OR, 1.5; 95% CI, 1.1–1.9).²³ A second study using *International Classification of Diseases*, 10th revision, codes for substance use similarly noted an increased incidence of preterm birth among marijuana users (18.8% vs 5.8%; $P < .001$).²⁴

In contrast, in the Avon Longitudinal Study of Pregnancy and Childhood, which is a population-based cohort from the United Kingdom to study

environmental exposures that affect the health and development of children (n = 12,129), the preterm birth rate in women who used marijuana weekly beyond the first trimester was exactly the same as nonusers at 4.6% ($P = .976$).²²

One prospective multicenter study by Shiono et al⁴⁰ highlights one of the difficulties in marijuana research. Only 31% of the women with a positive serum screen for THC (n = 585) also self-reported use in a structured interview. Conversely, only 43% of women who self-reported use had a positive serum assay for THC. These investigators grouped women who reported use and/or had a positive drug assay for THC and demonstrated no association between preterm birth and marijuana use (OR, 1.1; 95% CI, 0.8–1.3).⁴⁰ There was, however, an association with preterm birth in women when only the women with a positive serum assay (possibly more chronic users) were considered marijuana exposed (OR, 1.3, 95% CI, 1.0–1.7).

Multiple other prospective cohort studies and secondary analyses fail to provide a definitive answer regarding preterm birth and marijuana use (Table 3). The majority of studies demonstrate no increased risk of preterm birth. However, the 2 studies mentioned in previous text that use spontaneous preterm birth as the outcome demonstrate an association with marijuana use (Table 3).^{4,39} Further research with detailed documentation of obstetrical history (specifically history of preterm birth and risk factors for preterm birth), quantification of marijuana use, and indication for delivery is needed.

Congenital anomalies

There are 2 studies in which data were collected prospectively to assess for an association of marijuana exposure with congenital anomalies (Table 4). Neither of these demonstrated an association between marijuana use and major congenital anomalies. There are also several large retrospective cohort studies examining whether there is an association between marijuana and birth defects, with mixed results (Table 4).^{41–43} Unfortunately, the majority of these

studies are based on birth defects registries with incomplete ascertainment of confounding factors and potential for recall bias with the exposure data collected long after delivery.

Current evidence does not support an association between marijuana exposure and any specific congenital birth defect (Table 4).

Neurodevelopment

There have been multiple animal studies and retrospective human studies looking at the effect of maternal marijuana use during pregnancy on neurodevelopment, behavior, and intelligence.

Animal studies have shown alterations in neurotransmitter and neuroendocrine systems in the offspring of rodents exposed to cannabinoids. This effect is particularly pronounced within dopaminergic pathways.⁴⁴ In addition, there have been some animal studies that show a marked increase in hyperactivity and exploratory behaviors in female rats.⁴⁵ Other rat studies have shown persistent deleterious effects on learning and memory functions in exposed offspring.⁴⁶ Whereas rodent animal studies cannot be extrapolated directly to humans, they can help elucidate some of the mechanisms by which marijuana affects the developing brain.

In human research, there is one published series of postmortem fetal brains (n = 44) from 17–22 week elective terminations exposed to marijuana.²⁷ Dopamine receptors were reduced in the amygdala of marijuana exposed-compared with nonexposed fetuses. This effect was most prominent in male fetuses and was directly correlated with the amount of marijuana used during the pregnancy.⁴⁷

Human research on drug exposure in utero and its subsequent effects is challenging because of confounding psychosocial issues and ongoing exposures that are impossible to fully adjust for in multivariable modeling.⁴⁸ The following data must therefore be interpreted with caution.

One study assessed 26 infants born to adolescent mothers who were exposed to marijuana in utero compared with

TABLE 4
Summary of marijuana and congenital anomalies studies

Study and number in cohort	Marijuana-exposed women, n (%)	Study design and setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs with 95% CIs reported when available)	Limitations and comments
Any major congenital malformation							
Linn et al, 1983 ⁷¹ (n = 12,424)	1246 (10)	Prospective cohort Single institution	Self-report by structured interview postpartum	Frequency: occasional, weekly, or daily use	SES, OB hx, medical hx, standard demo, other drugs, EtOH, tobacco	Rate of major malformation: 2.6% nonusers, 3.2% occasional, 3.9% weekly, 3.6% daily No association with major congenital anomalies (OR, 1.36; 95% CI, 0.97–1.91)	Possible recall bias, exposure data collected postpartum No data on trimester of exposure Marijuana use not well quantified
Gibson et al, 1983 ⁷³ (n = 7301)	392 (5)	Secondary analysis of a prospective cohort Two affiliated institutions	Self-report by structured interview at 1 prenatal visit and postpartum	Frequency: ≤1 times/wk, >1 time/wk	Standard demo, OB hx (parity only), EtOH, tobacco	Rate of major malformation: 4.2% in cohort, rates by nonusers and users of marijuana not provided No association with congenital anomalies, no OR reported	No data on trimester of exposure Marijuana use not well quantified
Gastroschisis							
Forrester et al, 2007 ⁸⁴¹ (n = 316,508)	829 (0.3)	Retrospective cohort from a Hawaiian birth defects registry	Self-report in medical record or positive urine toxicology screen at delivery	Reported use: yes/no THC in maternal urine at time of delivery (ordered clinically)	None	n = 109 total cases of gastroschisis, n = 3 cases of gastroschisis in marijuana-exposed Rate ratio of marijuana users compared with women with other live births, 23.11; 95% CI, 4.69–69.34	Low prevalence of marijuana use (0.3%) indicates incomplete ascertainment of exposure No adjustment for possible confounders No data on trimester of exposure Marijuana use not well quantified
van Gelder et al, 2009 ⁸⁴² (n = 10,241 cases with anomalies and 4967 controls)	610 (4)	Retrospective cohort from a multistate birth defects registry	Self-report by structured interview 6 wks to 24 mo after delivery	Reported use: yes/no from 1 month before pregnancy to end of pregnancy	Standard demo including maternal age at delivery, EtOH, tobacco, folic acid use, maternal diabetes	n = 485 total cases of gastroschisis, n = 189 cases of gastroschisis in marijuana-exposed No association with gastroschisis (OR, 1.3; 95% CI, 0.9–1.8)	Possible recall bias, women interviewed 6 wks to 24 mo after delivery No data on trimester of exposure Marijuana use not well quantified

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(continued)

TABLE 4
Summary of marijuana and congenital anomalies studies (continued)

Study and number in cohort	Marijuana-exposed women, n (%)	Study design and setting	Data source	Marijuana measure	Other variables considered in analysis	Findings (adjusted ORs with 95% CIs reported when available)	Limitations and comments
Ventricular septal defect							
Williams et al, 2004 ⁴³ (n = 122 cases with a VSD and 3029 controls)	253 (8)	Retrospective cohort from Atlanta Birth Defects Case-Control Study	Self-report by structured telephone interview postpartum	Reported use: <2 d/wk, ≥3 d/wk from 3 months before pregnancy to end of first trimester	Cases matched to controls by birth year, race, birth period, and hospital of birth Adjusted for maternal age, multivitamin use, maternal diabetes	n = 122 total cases of VSD, n = 20 cases of VSD in marijuana exposed Marijuana use associated with VSD (adjusted OR, 1.90; 95% CI, 1.29–1.81)	Possible recall bias, women interviewed after delivery Incomplete ascertainment of confounding factors Marijuana use not well quantified
van Gelder et al, 2009 ⁴² (n = 10,241 cases with anomalies and 4967 controls)	610 (4)	Retrospective cohort from a multistate birth defects registry	Self-report by structured interview 6 wks to 24 mo after delivery	Reported use: yes/no from 1 mo before pregnancy to end of pregnancy	Standard demo including maternal age at delivery, EtOH, tobacco, folic acid use, maternal diabetes	n = 927 total cases of perimembranous VSD, n = 34 cases of perimembranous VSD in marijuana-exposed No association with VSD (OR, 0.9; 95% CI, 0.6–1.4)	Possible recall bias, women interviewed 6 wks to 24 mo after delivery No data on trimester of exposure Marijuana use not well quantified
Anencephaly							
van Gelder et al, 2009 ⁴² (n = 10,241 cases with anomalies and 4967 controls)	610 (4)	Retrospective cohort from a multistate birth defects registry	Self-report by structured interview 6 wks to 24 mo after delivery	Reported use: yes/no from 1 month before pregnancy to end of pregnancy	Standard demo including maternal age at delivery, EtOH, tobacco, folic acid use, maternal diabetes	n = 244 total cases of anencephaly, n = 12 cases of anencephaly in marijuana exposed Marijuana use associated only with anencephaly in a subanalysis restricted to first month after conception exposure (OR, 2.5; 95% CI, 1.3–4.9)	Possible recall bias, women interviewed 6 wks to 24 mo after delivery No data on trimester of exposure Marijuana use not well quantified

CI, confidence interval; EtOH, alcohol use; Medical hx, medical history; OB hx, obstetrical history; OR, odds ratio; SES, socioeconomic status; Standard demo, some measure of standard demographics including maternal age, race, body mass index; THC, delta-9-tetrahydrocannabinol; VSD, ventricular septal defect.

^a Rates of other birth defects that were higher in women with isolated (no other drug use) marijuana use in study by Forrester et al⁴¹ (2007) were encephalocele, hydrocephaly, microcephaly, anotia/microtia, tetralogy of Fallot, atrial septal defect, pulmonary valve atresia/stenosis, hypoplastic left heart syndrome, cleft lip and palate, pyloric stenosis, anal/rectal/large intestinal atresia/stenosis, obstructive genitourinary defect, polydactyly, syndactyly, and reduction deformity of upper limbs. These findings are not further described, given the limitations in the methodology of this study with no correction for possible confounders; ^b No association was found between marijuana and several other birth defects in the study by van Gelder et al⁴² (2009) including spina bifida, anotia/microtia, d-transposition of the great arteries, tetralogy of Fallot, hypoplastic left heart syndrome, coarctation of the aorta, pulmonary valve stenosis, atrial septal defect, cleft lip and palate, esophageal atresia, anorectal atresia, hypospadias, transverse limb deficiency, craniosynostosis, and diaphragmatic hernia.

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nonexposed infants of demographically matched mothers.⁴⁹ Exposure was confirmed by maternal hair samples and neonatal meconium testing. Those exposed to marijuana had significantly different arousal, regulation, and excitability on the Neonatal Intensive Care Unit Network Neurobehavioral Scale.

There are 2 large cohorts with both short- and long-term follow-up of children exposed to marijuana in utero. The Ottawa Prenatal Prospective Study looked at the effects of prenatal marijuana and tobacco use on 180 offspring of primarily middle class, white, low-risk patients in Ottawa, Canada, at various developmental ages.⁵⁰ Younger than age 4 years, there were no differences in behavior problems, intellect, visual perception, language, or sustained attention and memory tasks between children born to mothers who used marijuana and those who did not. However, after the age of 4 years, there were differences in behavioral problems and poorer performance on visual perception tasks as well as language comprehension and sustained attention and memory difficulties in exposed children.⁵⁰ By the age of 9-12 years, there was no difference between exposed and unexposed children in global intelligence quotient scores or performance on visual tasks and impulse control.⁵¹

Although the Ottawa Prenatal Prospective Study provides much-needed long-term follow-up of exposed children, it has limitations. It does not adequately correct for environmental factors, does not clearly report differences based on the quantity of marijuana used, and is a relatively homogenous population.

The other large cohort with long-term follow-up is the Maternal Health Practices and Child Development Project (MHPCD) from Pittsburgh, PA, which consists of mostly high-risk, low-income minority women and their children.⁵² Whereas there were no differences in intelligence testing at 3 years of age,⁵² maternal use of 1 or more joints per day during the first trimester was associated with decreased verbal reasoning by the age of 6 years.⁵³

TABLE 5

Recommendations for clinicians regarding marijuana use in pregnancy**Recommendations**

- Screen all women verbally for marijuana use at intake to obstetrical care
- Consider rescreen later in pregnancy
- Consider urine toxicology screening in high-risk patients
- Recommend avoiding marijuana in pregnancy
- Marijuana crosses the placenta
- Counsel women regarding uncertainty of effects on perinatal outcomes
- Possible increased risk of stillbirth
- Possible increased risk of preterm birth (mixed data)
- Counsel women regarding uncertainty of effects on offspring
- Possible adverse effects on neurodevelopment
- Possible increased risk of fetal growth restriction (mixed data)
- No established association with specific congenital anomalies
- Refer women who use marijuana and desire cessation to appropriate resources
- Local substance-use programs
- Do not otherwise modify clinical care
- Growth ultrasounds not indicated outside study protocols
- Screening for preterm birth with cervical length not indicated
- Antenatal surveillance not indicated
- Recommend avoiding marijuana while lactating
- Marijuana is passed to the neonate in breast milk
- Possible adverse effects on early neurodevelopment
- Provide counseling, but do not withdraw lactation support

Recommendations in the Table above reflect the opinions of the authors after a thorough review of the existing literature on marijuana in pregnancy and lactation.

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The MHPCD cohort was examined again at age 10 years (n = 636). Those children exposed during the first and third trimesters demonstrated decreased attention and more hyperactivity and impulsivity.⁵⁴ Academic performance in reading and spelling and by teacher report was worse in those exposed to at least 1 joint per day during pregnancy.⁵⁵ In the last assessment of the MHPCD cohort at the age of 14 years (n = 524), maternal use was associated with lower scores in reading, math, and spelling, most notably in those exposed to heavy use in the first trimester.⁵⁶ In addition, there was an earlier age of onset of substance use and greater duration of use than their matched counterparts, even after adjustment for home environment and parental substance use.⁵⁷

Although the human research in neonatal and childhood development following marijuana exposure is flawed by factors including the concurrent use of other substances, variability in exposure dosing and frequency, other genetic or environmental factors, and a

reliance on self-reported data, there is a concerning pattern of altered neurodevelopment with early, heavy maternal use of marijuana.

Breast-feeding

Cannabinoids consumed by lactating mothers reach the newborn during breast-feeding.⁵⁸ The amount that reaches the infant is estimated at 0.8% of the mother's exposure.⁵⁹ There is some evidence that marijuana use inhibits milk production by inhibiting prolactin secretion.⁶⁰

Astley and Little⁶¹ attempted to determine the effects of marijuana use on infant development at 1 year. Infants exposed to marijuana during lactation scored poorly on the Psychomotor Developmental Index compared with those not exposed. However, this result could not be separated from the effect of marijuana use during pregnancy.⁶¹ Eighty-four percent of users during pregnancy continued use during lactation.⁶¹

The American Academy of Pediatrics' policy statement on "Breastfeeding and

the Use of Human Milk” states that breast-feeding is contraindicated in women using illicit drugs.⁶² The statement does not address whether hospitals should withdraw lactation support in the form of facilitation of breast-feeding by nursing and lactation consultants. In our opinion, given the paucity of data regarding ongoing exposure to marijuana through breast milk, and multiple known benefits of breast-feeding, lactation support should not be withdrawn. However, women should clearly be educated regarding the potential adverse effects of ongoing marijuana exposure through breast milk and encouraged to stop using marijuana while lactating.

Future research

Despite a large volume of literature on the topic of marijuana in pregnancy, there is still a need for high-quality, contemporary, prospective data to better understand the effects of marijuana use in pregnancy and lactation.

We have identified the following research gaps as areas of focus for future studies: (1) determining whether there is an association between marijuana use and congenital anomalies, spontaneous preterm birth, pregnancy loss and still-birth, or poor fetal growth by serial ultrasound assessments; (2) confirming the long-term neurobehavioral consequences of marijuana exposure with longitudinal follow-up; (3) establishing whether there are adverse effects of breast-feeding in the setting of ongoing marijuana use; (4) characterizing the changes in the prevalence of use during pregnancy in states with legalized marijuana for medical and recreational use; (5) understanding women’s attitudes and beliefs regarding marijuana in pregnancy in the setting of increasing legalization; and (6) understanding the impact of different modes of consumption on outcomes with increased use of edible forms of marijuana that contain high concentrations of THC.

While performing any study on marijuana, it will be important to collect participant’s socioeconomic status, medical history, obstetrical history, use of other drugs, and alcohol and tobacco use in a detailed, methodical manner. In

addition, studies will need to quantify the timing and the amount of marijuana ingested and corroborate self-report with biological specimens.

Ultimately high-quality data will enable obstetricians to appropriately counsel women regarding marijuana use in pregnancy. If adverse effects are confirmed, intervention and education programs can be developed to minimize morbidity to mothers and their babies.

Summary

Summary recommendations for the practicing clinician are listed in **Table 5**. These recommendations are made after a thorough review of the existing literature but are based on studies of varying methodological quality with mixed results and reflect the opinions of the authors after completing this extensive review. Until further data are available, we should continue to discourage women from using recreational drugs, including marijuana, during pregnancy and lactation, given the uncertain short- and long-term outcomes. ■

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