

#### Legalization of cannabis: Implications for maternal and infant health in BC and emerging best practice for response

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Sabrina Luke, PhD Epidemiologist, Perinatal Services BC



#### **Legalization of Cannabis in Canada**

- Cannabis currently a Schedule II drug under the Controlled Drug and Substances Act.
- Cannabis Act (Bill C-45) introduced in 2017.
- Subject to parliamentary approval, cannabis could be legalized in Canada by July 2018.
- Recommendations for use in pregnancy?



#### **Trends of cannabis use in Canada**

- Canada has one of the highest rates of cannabis use among youth in the world.
- Cannabis use among Canadian women increased from 7.4% in 2013 to 9.7% in 2015.
- Cannabis is the most common illicit drug used by pregnant women.



## **Potential implications**

- Nearly half of pregnancies in Canada are unplanned.
- Cannabinoids cross the placenta with greater transfer in early pregnancy.
- Potential for fetal exposure in the first trimester before women know they are pregnant.





#### **PDR Data Analysis**

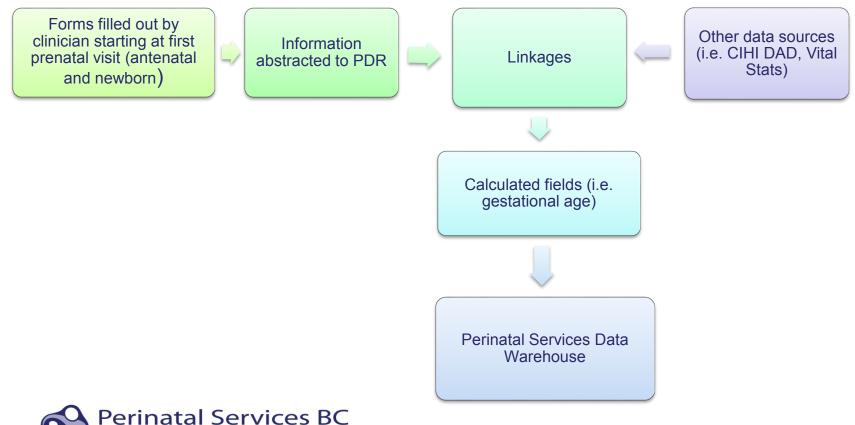
## **Objectives**

- To examine the trends of cannabis use among pregnant women in BC and related maternal characteristics.
- To determine the association between cannabis use in pregnancy and perinatal outcomes among women in BC including:
  - Stillbirth (antepartum and intrapartum)
  - Spontaneous preterm birth
  - Small-for-gestational age



## **Perinatal Data Registry (PDR)**

An agency of the Provincial Health Services Authority



#### **Antenatal Form**

8. Lifestyle & Social Discussed Concerns	Referred
Diet/Food Safety	
Folic acid	
Physical Activity/rest/work	
OTC drugs/vitamins	
Alcohol never quit (DD/MM/YYYY) Drinks/wk: before pregnancy current Binge drinking No Yes	
TWEAK score (see reverse)	
Substance use       No       Yes         Heroin       Cocaine       Marijuana         Methadone       Solvents       Other         Prescription       Unknown	
Smoking never quit (DD/MM/YYYY) Cig/day: before pregnancy current	
Exposure 2nd hand smoke INO Yes	



#### **Methods**

- Time period: April 1<sup>st</sup>, 2008 to March 31<sup>st</sup>, 2016.
- Excluded records:
  - missing data on outcomes and gestational age
  - gestational ages less than 20 weeks or greater than 44 weeks.
- Singleton births



#### **Methods**

- Trends of cannabis use among pregnant women in BC.
- Chi-square analysis of maternal characteristics and cannabis use.
- Logistic Regression:
  - Association between cannabis use and small-forgestational age (<10th) and spontaneous preterm birth (<37 weeks).</li>
- Cox Proportional Hazards Model:
  - Cannabis use and stillbirth
  - Ran separate models for overall stillbirth, antepartum and intrapartum
  - Excluded late termination



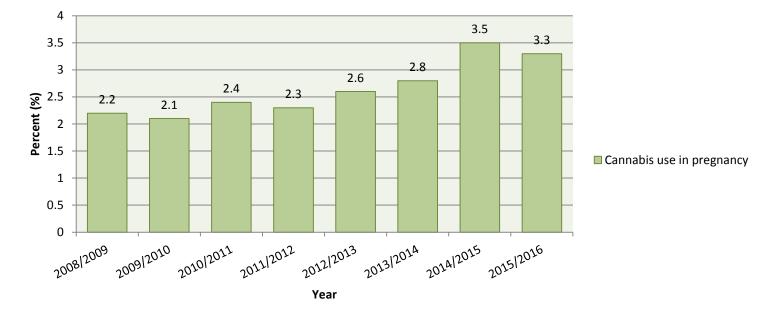
#### **Methods**

- Logistic regressions and Cox proportional hazards models adjusted for:
  - maternal age
  - maternal ethnicity (Nam Pehchan)
  - SES (QAIPPE)
  - pre-pregnancy BMI
  - tobacco use in pregnancy
  - alcohol use in pregnancy
  - illicit substances (cocaine, heroin/opioids, solvents, designer drugs, methadone, hallucinogens, stimulants, prescription medication, nonprescription medication, other unknown drug)
- Cox proportional hazards models also adjusted for any chromosomal or congenital anomalies.



#### **Results: Trends of cannabis use in BC**

• Cannabis use among pregnant women in BC has increased over the past decade.





#### **Results: Maternal characteristics and cannabis use**

Maternal Characteristics	Cannabis Use N=5801	No Cannabis Use N=237339	p-value
Maternal age			<0.0001
10-24	49.8	14.6	
25-29	25.7	27.9	
30-34	16.8	34.2	
35-39	6.6	19.1	
40+	1.2	4.4	
Neighborhood SES (QAIPPE)			<0.0001
1 - Lowest	34.1	21.0	
2	22.3	21.3	
3	18.0	20.9	
4	15.7	20.5	
5 – Highest	10.0	16.3	
Tobacco use	69.6	16.6	<0.0001
Any alcohol use	12.1	0.9	<0.0001
Other substance use	15.8	0.9	<0.0001
History of anxiety	17.9	6.5	<0.0001
History of depression	28.7	9.6	<0.0001
History mental illness	41.1	15.7	<0.0001

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#### **Results: Perinatal outcomes and cannabis use**

Perinatal outcomes	Cannabis Use N=5801	No Cannabis Use N=237339	p-value
Small-for-gestational age (<10 <sup>th</sup> percentile)	10.1	6.5	<0.0001
Large-for-gestational age	10.0	12.7	<0.0001
Spontaneous preterm birth	7.5	4.3	<0.0001
Stillbirth	0.5	0.3	0.0001
Antepartum Stillbirth	0.4	0.2	<0.0001
Intrapartum Stillbirth	0.14	0.02	<0.0001



## **Results: Risk for SGA, preterm birth and stillbirth**

#### Association between cannabis use and SGA, any spontaneous preterm birth and stillbirth

Birth Outcome	Cannabis use during pregnancy	
	Unadjusted OR [CI]	Adjusted OR [CI]*
SGA (<10th percentile)	1.63 [1.52-1.75]	1.48 [1.37-1.59]
Spontaneous Preterm Birth (<37 weeks)	1.84 [1.70-2.00]	1.31 [1.20-1.43]
	Unadjusted HR [CI]	Adjusted HR [CI]*
Stillbirth (All)	1.66 [1.19-2.32]	1.38 [0.95-1.99]
Antepartum Stillbirth <sup>a</sup>	1.47 [0.99-2.18]	1.34 [0.88-2.06]
Intrapartum Stillbirth <sup>b</sup>	4.84 [2.33-10.1]	2.84 [1.18-6.82]

\*Adjusted for maternal age, pre-pregnancy BMI, tobacco use, alcohol use, other substance use, QAIPPE and ethnicity.

<sup>a</sup> Stillbirth that occurred before the onset of labour.

<sup>b</sup> Stillbirth that occurred after the onset of labour.



#### **Mechanisms of action**

- Fetal oxygen and nutrient deprivation from altered blood supply through changes in placental vasculature.
- Changes in glucose and insulin regulation which restricts fetal growth (through insulin-like growth factors 1 and 2)?
- Similar to the effects of tobacco



# **Strength and Limitations**

#### Strengths

- Large dataset that includes >99% of births in BC
- PDR data includes information on tobacco and alcohol use in pregnancy, other illicit substances and prescription medications.

Limitations

- Self-report of cannabis use, tobacco, alcohol and other substances.
- Co-occurrence of cannabis use and tobacco use.
- Alcohol use underreported and difficult to adjust for.
- Ethnicity
- Neighborhood-based index of SES



#### **Conclusions**

- Cannabis use in pregnancy and history of mental illness.
- Association between cannabis use and SGA, and spontaneous preterm birth.
- Stillbirth needs further research
- Timing of cannabis exposure, type and frequency.
- Recommendations for cannabis use in pregnancy?



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