

#### Associations between PM2.5 components, brake and tire wear markers, ultrafine particles, and childhood cancers in Canada

#### Eric Lavigne, PhD





 Outdoor fine particulate matter (PM2.5) and UFPs are a heterogeneous mix



 Most epidemiological studies only contrast PM2.5 and UFP concentrations



#### PM2.5, UFPs & pediatric cancer development



Maternal exposure to ambient air pollution and risk of early childhood cancers: A population-based study in Ontario, Canada

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### PM2.5, UFPs & pediatric cancer development

RR for associations between PM2.5 and incidence of selected childhood cancers

|              | PM <sub>2.5</sub> (IQR=2.9 μg/m3) |                    |                      |                      |  |
|--------------|-----------------------------------|--------------------|----------------------|----------------------|--|
| Cancer site  | Obs. cases                        | Crude <sup>a</sup> | Model 1 <sup>b</sup> | Model 2 <sup>c</sup> |  |
|              |                                   | HR (95% CI)        | HR (95% CI)          | HR (95% CI)          |  |
| ALL          | 365                               | 1.04 (0.91-1.18)   | 1.05 (0.91-1.22)     | 1.03 (0.88-1.20)     |  |
| Astrocytoma  | 88                                | 1.14 (0.88-1.47)   | 1.38 (1.01-1.88)     | 1.35 (1.00-1.85)     |  |
| Wilms tumour | 88                                | 1.10 (0.84-1.44)   | 1.17 (0.86-1.59)     | 1.20 (0.88-1.64)     |  |

a Unadjusted model.

b Model adjusted for maternal age at delivery, infant sex, parity, year of birth, census tract median family income, census tract proportion of population who are visible minority and census tract proportion of the adult female population aged 25-64 years old who completed postsecondary education.

c Additional adjustment for maternal cigarette smoking during pregnancy using multiple imputation



#### PM2.5, UFPs & pediatric cancer development



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#### Ambient ultrafine particle concentrations and incidence of childhood cancers

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#### Table 4

Hazard ratios<sup>a</sup> (HR) and 95% confidence intervals (95% CI) for the associations between UFPs (per 10,000/cm<sup>3</sup>) over specific periods and childhood cancer risk  $\leq$  6 years of age with additional adjustment for PM<sub>2.5</sub> and NO<sub>2</sub>.<sup>b</sup>

| Cancer site              | 1st trimester    | 2nd trimester    | 3rd trimester    | Entire pregnancy | Childhood exposure |
|--------------------------|------------------|------------------|------------------|------------------|--------------------|
| $UFPs + NO_2$            |                  |                  |                  |                  |                    |
| All cancers              | 1.11 (1.02-1.21) | 1.00 (0.88-1.07) | 0.96 (0.89-1.06) | 1.00 (0.91-1.12) | 0.97 (0.92-1.03)   |
| ALL                      | 1.23 (0.87-1.74) | 0.90 (0.58-1.27) | 1.07 (0.78-1.44) | 1.06 (0.77-1.42) | 1.04 (0.75-1.40)   |
| Astrocytoma              | 0.49 (0.18-1.34) | 0.80 (0.55-1.45) | 1.00 (0.64-1.55) | 0.90 (0.51-1.48) | 0.71 (0.46-1.10)   |
| Neuroblastoma            | 0.91 (0.58-1.43) | 0.98 (0.70-1.31) | 0.88 (0.64-1.12) | 0.81 (0.59-1.11) | 1.11 (0.80-1.54)   |
| UFPs + PM <sub>2.5</sub> |                  |                  |                  |                  |                    |
| All cancers              | 1.10 (1.01-1.19) | 0.97 (0.86-1.07) | 0.91 (0.82-1.01) | 0.99 (0.88-1.11) | 0.96 (0.88-1.09)   |
| ALL                      | 1.14 (0.95-1.36) | 0.87 (0.60-1.27) | 1.07 (0.79-1.46) | 1.03 (0.75-1.44) | 0.95 (0.72-1.28)   |
| Astrocytoma              | 0.65 (0.37-1.13) | 0.85 (0.55-1.47) | 1.00 (0.61-1.50) | 0.82 (0.51-1.50) | 0.78 (0.50-1.18)   |
| Neuroblastoma            | 1.05 (0.77-1.49) | 1.01 (0.73-1.35) | 0.90 (0.69-1.18) | 0.98 (0.71-1.32) | 1.12 (0.79-1.54)   |
| $UFPs + PM_{2.5} + NO_2$ |                  |                  |                  |                  |                    |
| All cancers              | 1.13 (1.03-1.22) | 1.01 (0.89-1.08) | 0.97 (0.90-1.07) | 1.01 (0.92-1.13) | 0.97 (0.92-1.04)   |
| ALL                      | 1.22 (0.85-1.72) | 0.90 (0.58-1.28) | 1.07 (0.78-1.45) | 1.06 (0.76-1.43) | 1.04 (0.74-1.41)   |
| Astrocytoma              | 0.55 (0.20 1.07) | 0.85 (0.53-1.47) | 1.00 (0.60-1.58) | 0.91 (0.52-1.49) | 0.75 (0.48-1.12)   |
| Neuroblastoma            | 0.95 (0.63-1.45) | 0.98 (0.70-1.31) | 0.89 (0.65-1.13) | 0.88 (0.65-1.17) | 1.12 (0.81-1.55)   |
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Objective

#### EVALUATE THE ASSOCIATIONS BETWEEN PRENATAL & EARLY LIFE EXPOSURES TO PM2.5 COMPONENTS, BRAKE & TIRE WEAR MARKERS, ULTRAFINE PARTICLES AND CHILDHOOD CANCER RISK



#### Methods

- Study Design: Population-based retrospective cohort study
  - » Canadian Vital Statistics Birth Database linked to Canadian Cancer Registry from 2000 to 2021





### Exposure data

<u>Bi-weekly</u> PM2.5 total mass and component (SO4, NH4, NO3, SS, BC, dust, OM) estimates at 1x1km derived from satellite-based estimates with geographically weighted regression (2000 to 2021)



Long term measure of Barium [Ba] as a marker of brake dust and Zinc [Zn] as a tire and road wear marker (available for 40 sites across Canada) (data measured from 2016 to 2018)



#### Exposure data

Model predictions of within-city spatial variation in median <u>daily</u> outdoor UFP levels UFP number concentration for two largest cities in Canada (Toronto and Montreal)





- » Cox proportional hazards models assessing risk until 14 years of age
- » Distributed Lag Non-Linear Models (DLNM) extension accounting for <u>gestational weekly exposures</u> for PM components and UFPS
- » Long term measures for Ba and Zc
- » Hazard ratios (HR) presented per IQR increase
- Covariates: maternal age, parity, infant sex, birth weight, arealevel socioeconomic variables, geographic location (rural/urban), season of birth, year of birth
  - Maternal smoking only for the province of Ontario



### Results

Study population (2000 – 2021)

| Variable      | Total cohort -<br>all births | All pediatric<br>cancer cases | Acute<br>Lymphoid<br>Leukemia | Acute Myeloid<br>Leukemia | Astrocytoma  |
|---------------|------------------------------|-------------------------------|-------------------------------|---------------------------|--------------|
| Ν             | 8,383,345                    | 22,230                        | 5,240                         | 960                       | 1,780        |
| Maternal age  |                              |                               |                               |                           |              |
| at delivery,  | 29.52                        | 29.49                         | 29.56                         | 30.39                     | 29.11        |
|               | (5.40)                       | (5.52)                        | (5.54)                        | (5,70)                    | (5.49)       |
|               | (3.49)                       | (5.53)                        | (5.54)                        | (5.70)                    | (3.40)       |
| Male          | 51                           | 54                            | 56                            | 52                        | 51           |
| Female        | 49                           | 46                            | 44                            | 48                        | 49           |
| PM 2.5 during |                              |                               |                               |                           |              |
| pregnancy,    | /                            |                               |                               |                           |              |
| mean (SD)     | 7.57 ( 2.22)                 | 7.87 (2.14)                   | 7.79 ( 2.10)                  | 7.61 (2.22)               | 7.74 ( 2.14) |



## Results - HRs for gestational exposures to PM components on the risk of ALL



### Results - HRs for gestational exposures on the risk of leukemia



## Results - $SO_4$ concentration-response curve on the risk of leukemia



# Results – UFPs, Ba & Zc on the risk of selected cancer types



Hazard ratio for UFPs, Ba & Zc on risk of selected car



#### Discussion

- SO<sub>4</sub> & UFPs were associated with ALL
  - » Early gestation (first few weeks) appeared to be a sensitive period
- Barium was associated with risk of astrocytoma





#### Conclusion

- Future studies should investigate components with finer temporal resolution models
- UFPs & particle size and childhood cancers should be further studied
- Ongoing research investigating childhood cancer survival, other cancer incidence subtypes & childhood exposures



### **THANK YOU**

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Health Canada

