

Associations between Halogenated Flame Retardants Exposure and Childhood Gut Microbiome Multi-omics Profiles in a Canadian Cohort

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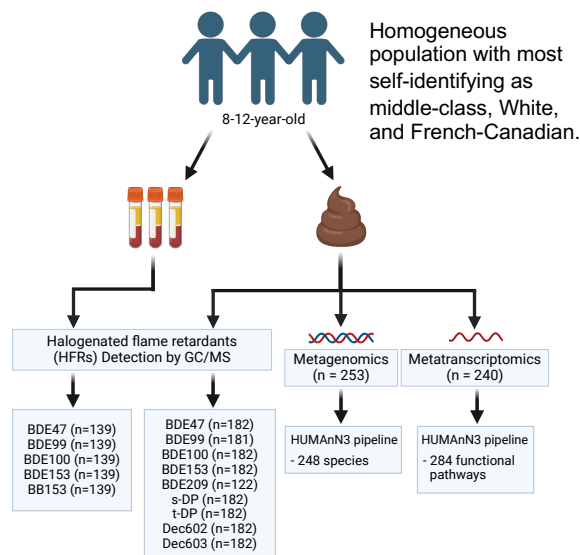
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Background & Objective

- Imbalance in the childhood gut microbiota has been linked to long-term adverse health outcomes, including metabolic disorders, infections, and autoimmune diseases. Early-life exposure to halogenated flame retardants (HFRs) may alter the gut microbiota, but this association remains unexplored.
- This study examined the associations of HFRs in plasma and stool with gut microbiome and transcriptome profiles in children aged 8-12 years from a birth cohort in Sherbrooke, Quebec, Canada.

Methods



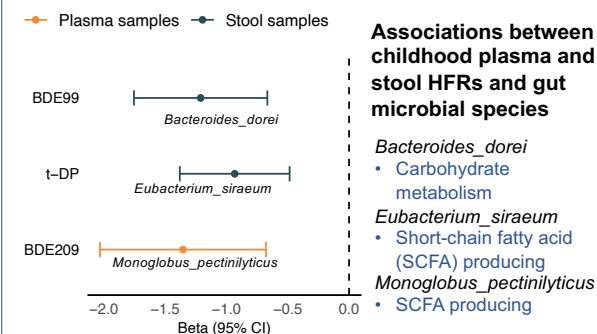
- Maaslin3 package for screening associations between HFRs and microbiome features;
- Models were adjusted for child's sex, family income, antibiotics intake in the past year, dietary patterns at the 8- to 12-year-old follow-up, and sequencing depth.

Study population

Characteristics	Distributions	HFRs	Detection (%)	Mean $\pm$ SD (ng/g)
Sex (%)		Plasma		
Male	106 (44.5)	BDE47	83.7	10.04 $\pm$ 19.77
Female	132 (63.5)	BDE99	53.5	3.53 $\pm$ 12.17
Family income (1,000\$ (mean (SD)))	104.2 (71.4)	BDE100	80.2	4.64 $\pm$ 8.33
Parents' education level (%)		BDE153	95.9	1.22 $\pm$ 6.93
College and University	151 (63.4)	BB153	44.7	6.02 $\pm$ 7.23
Vocational Training or Secondary school	85 (35.7)	Stool		
Other	2 (0.9)	BDE47	78.6	3.57 $\pm$ 6.60
Food Frequency Questionnaire Class (%)		BDE99	53.4	0.23 $\pm$ 0.03
Class 1	37 (15.5)	BDE100	71.3	0.09 $\pm$ 0.15
Class 2	86 (36.1)	BDE153	83.9	0.06 $\pm$ 0.09
Class 3	115 (48.4)	BDE209	95.4	154 $\pm$ 711
Metagenomic sequencing depth ^a (mean (SD))	34.7 (27.0)	s-DP	77.6	0.04 $\pm$ 0.14
Metatranscriptomic sequencing depth ^a (mean (SD))	33.26 (31.4)	t-DP	91.7	0.35 $\pm$ 2.84
		Dec602	35.9	0.006 $\pm$ 0.001
		Dec603	37.5	0.003 $\pm$ 0.005

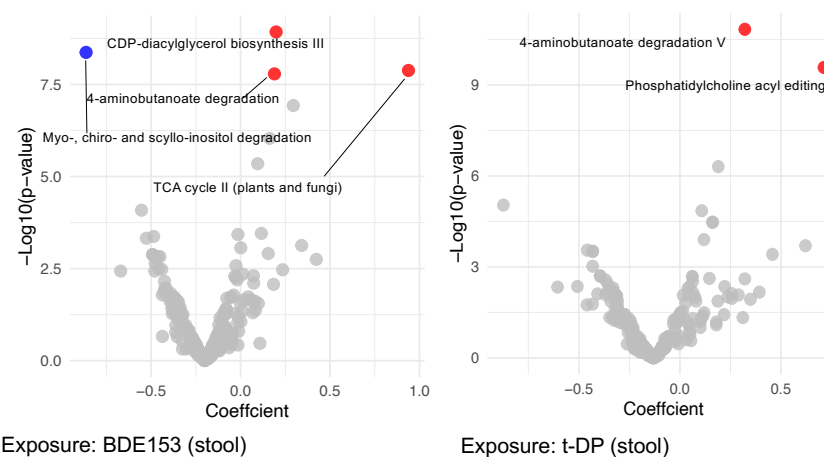
^aThe unit for sequencing depth is million reads

HFRs ~ microbiome species



HFRs ~ microbiome functional pathways

Associations between childhood plasma and stool HFRs and gut microbial transcriptomic functional pathways: six pathways are associated with stool HFRs, while no significant associations were observed between plasma HFRs and functional pathways.



- 4-aminobutanoate degradation;
 - Associated with γ -aminobutyric acid (GABA) degradation
- CDP-diacylglycerol biosynthesis III
 - Phospholipid synthesis, cell membrane integrity, gut barrier.
- Myo-, chiro- and scyllo-inositol degradation
 - Enhances lipid absorption and obesity
- TCA cycle II (plants and fungi)
 - Regulating energy metabolism
- Phosphatidylcholine acyl editing
 - Phosphatidylcholine (PC) metabolism, associated with gut barrier function.

Conclusions

In children from the GESTE cohort, both plasma and stool HFRs exposure were associated with alterations of gut microbiome, with stronger associations observed for stool HFRs.

Funding: National Institute of Environmental Health Sciences: R01ES027845; P30ES009089; Canadian Institutes of Health Research: MOP-84551; Natural Sciences and Engineering Research Council of Canada: CRC-950-230570; The University of Texas System Rising STARS award