

Exposome Analysis with Bayesian Group Index Regression

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Bayesian Group Index Regression

- Bayesian group index model:

$$y_i \sim \text{Bernoulli}(p_i)$$
$$\text{logit}(p_i) = \beta_0 + \sum_{k=1}^K \beta_k \left(\sum_{j=1}^{C_k} w_{kj} q_{kij} \right) + \mathbf{z}_i^T \phi,$$

- Priors:

$$\beta_k \sim \text{Normal}(0, \tau_k)$$

$$\tau_k = 1/\sigma_k^2$$

$$\sigma_k \sim \text{Uniform}(0, 100)$$

$$w_{k1}, \dots, w_{kC_k} \sim \text{Dirichlet}(\alpha_{k1}, \dots, \alpha_{kC_k}), \text{ for } k=1, \dots, K$$

- Dirichlet prior assures that the variable weights $w_{kj} \in (0, 1)$ and $\sum_{j=1}^{C_k} w_{kj} = 1$.

Exposome Data Analysis

- Modeled child asthma risk with predictors from the air pollution, indoor air, metal, organochlorine, pesticide, PBDE, PFAS, phenol, phthalate, traffic, natural space, meteorological, lifestyle, and built environment families using positive and negative association groups when mixed.

Parameter	Index	Predictors	Parameter	Index	Predictors
β_1	Air Pollution	NO2 (yr), PM10 (yr), PM25 (yr), PM25 Abs (yr)	β_{11}	Phthalates (-)	MBZP, MECPP, MEHHP, MEHP, MEOHP, OXOMINP
β_2	Indoor Air (-)	Absorbance, NO2	β_{12}	Phthalates (+)	MEP, MIBP, MNBP, OHMINP
β_3	Indoor Air (+)	PM, Benzene	β_{13}	Traffic	Traffic Load, Traffic Density
β_4	Metals (-)	As, Co, Cs, Cu, Pb	β_{14}	Natural Spaces	NDVI (h), NDVI (s)
β_5	Metals (+)	Cd, Hg, Mn, Mo	β_{15}	Meteorological	Humidity (month), UV dose (month)
β_6	Organochlorines	DDE, DDT, HCB, PCB138, PCB153, PCB170, PCB180	β_{16}	Lifestyle (-)	Sleep Time, KIDMED
β_7	Pesticides	DEP, DETP, DMP	β_{17}	Lifestyle (+)	Sedentary Time, MVPA
β_8	PBDEs	PBDE153, PBDE 47	β_{18}	Built Environment (-)	Access Points (h), Building Density (h), Population Density (h), Building Density (s), Population Density (s)
β_9	PFASs	PFHXS, PFOA, PFOS, PFUNDA	β_{19}	Built Environment (+)	Access Points (s), Connectivity (h), Connectivity (s), Facility Density (h), Facility Density (s), SEI (h), SEI (s)
β_{10}	Phenols	BPA, BUPA, ETPA, MEPA, OXBE, PRPA			

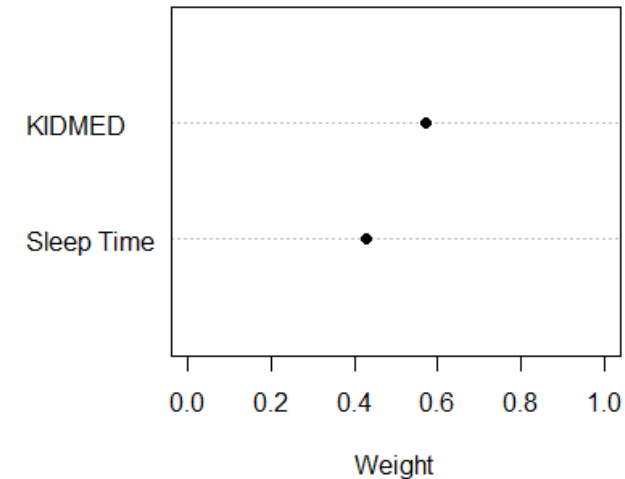
Exposome Data Analysis

Main Model - OR estimates for family groups and covariates

Variable	Odds Ratio	2.5% CI	97.5% CI
Family Groups			
Air Pollution	0.69	0.41	1.12
Indoor Air (-)	1.08	0.81	1.43
Indoor Air (+)	1.23	0.97	1.57
Metals (-)	0.74	0.50	1.02
Metals (+)	1.28	0.93	1.72
Organochlorines	1.14	0.86	1.55
Pesticides	0.77	0.60	0.98
PBDEs	0.72	0.56	0.92
PFASs	1.07	0.82	1.44
Phenols	0.95	0.71	1.28
Phthalates (-)	0.84	0.61	1.12
Phthalates (+)	1.36	0.99	1.94
Traffic	0.94	0.74	1.17
Natural Spaces	0.74	0.53	1.03
Meteorological	1.00	0.71	1.42
Lifestyle (-)	0.77	0.59	0.99
Lifestyle (+)	1.06	0.84	1.35
Built Environment (-)	0.69	0.47	1.16
Built Environment (+)	1.23	0.84	1.93

Variable	Odds Ratio	2.5% CI	97.5% CI
Cohort			
Cohort 1	5.89	1.99	18.31
Cohort 2	4.31	1.22	15.77
Cohort 4	1.92	0.57	7.46
Cohort 5	2.39	0.68	9.21
Cohort 6	1.16	0.33	4.36
Other Covariates			
Child Sex (Male)	1.63	1.06	2.49
Maternal Age	0.86	0.71	1.05

Weights for Lifestyle (-)



Exposome Data Analysis

- The significant cohort covariates indicated effect modification.
- Cohorts 3 and 6 had few asthma cases.
- Family indices were once again split when containing mixed associations with the outcome, so while similar to the main model the stratified model specifications are not identical.

Main Model:

- N=1301
- 19 indices
- Air Pollution predictors together in one index

Cohort 1 Model:

- N=202
- 20 indices
- Air Pollution (-) contains PM25 and PM25 Abs
- Air Pollution (+) contains NO2 and PM10

Exposome Data Analysis

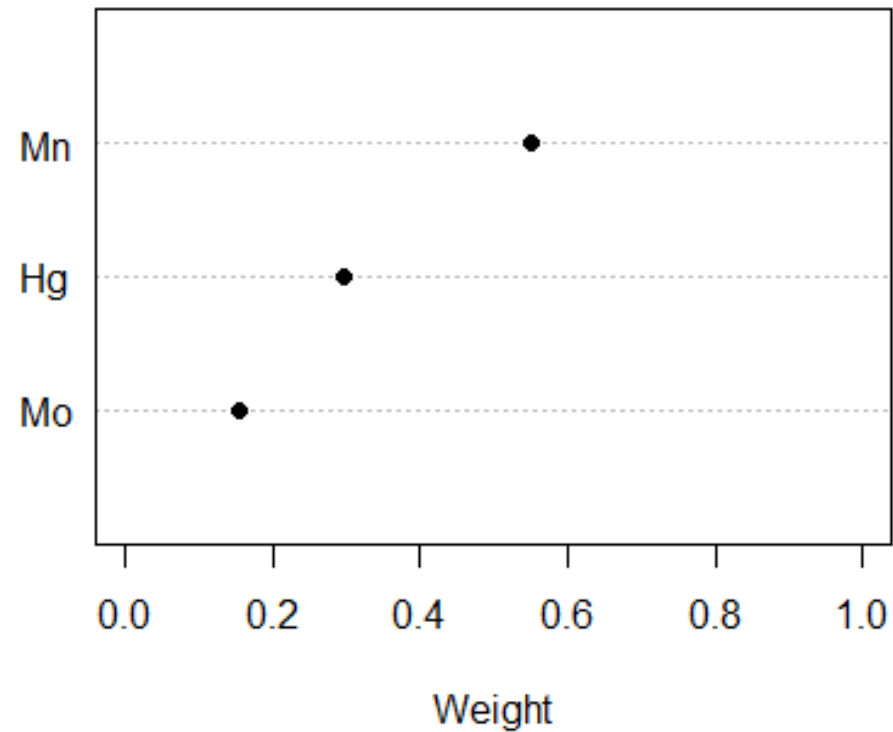
Significant Cohort Findings

Model	Significant Parameters	OR (95% CI)	Important Predictors (Weight)
Cohort 1			
Negative Effects:	Metals (-)	0.17 (0.04, 0.60)	Copper (0.21), Arsenic (0.21)
	Phthalates (-)	0.37 (0.12, 0.94)	OXOMINP (0.28)
Positive Effects:	Phthalates (+)	2.99 (1.27, 8.18)	MEP (0.51), OHMINP (0.49)
	Metals (+)	2.39 (1.02, 6.05)	Manganese (0.55)
	Phenols (+)	2.23 (1.01, 5.26)	OXBE (0.75)
	Child Sex (Male)	4.11 (1.22, 15.20)	
Cohort 2			
Positive Effects:	Organochlorines (+)	2.50 (1.04, 6.85)	PCB 138 (0.42)
Cohort 4			
Negative Effects:	Built Environment (-)	0.10 (0.01, 0.59)	Home and School Connectivity Density (0.27 and 0.26)
	Metals (-)	0.11 (0.01, 0.79)	Cadmium (0.23)
	Phenols (-)	0.14 (0.01, 0.82)	BPA (0.26)
	Natural Spaces	0.21 (0.04, 0.65)	Home NDVI (0.64)
Positive Effects:	PBDEs	0.27 (0.06, 0.80)	PBDE 153 (0.58)
	Pesticides (+)	2.80 (1.01, 10.96)	DMTP (0.60)
Cohort 5			
Negative Effects:	Built Environment (-)	0.17 (0.05, 0.43)	Home Access Points (0.67)
Positive Effects:	Traffic	2.03 (1.01, 4.12)	Traffic Density (0.79)

Exposome Data Analysis

Cohort 1: OR = 2.39 (1.02, 6.05)

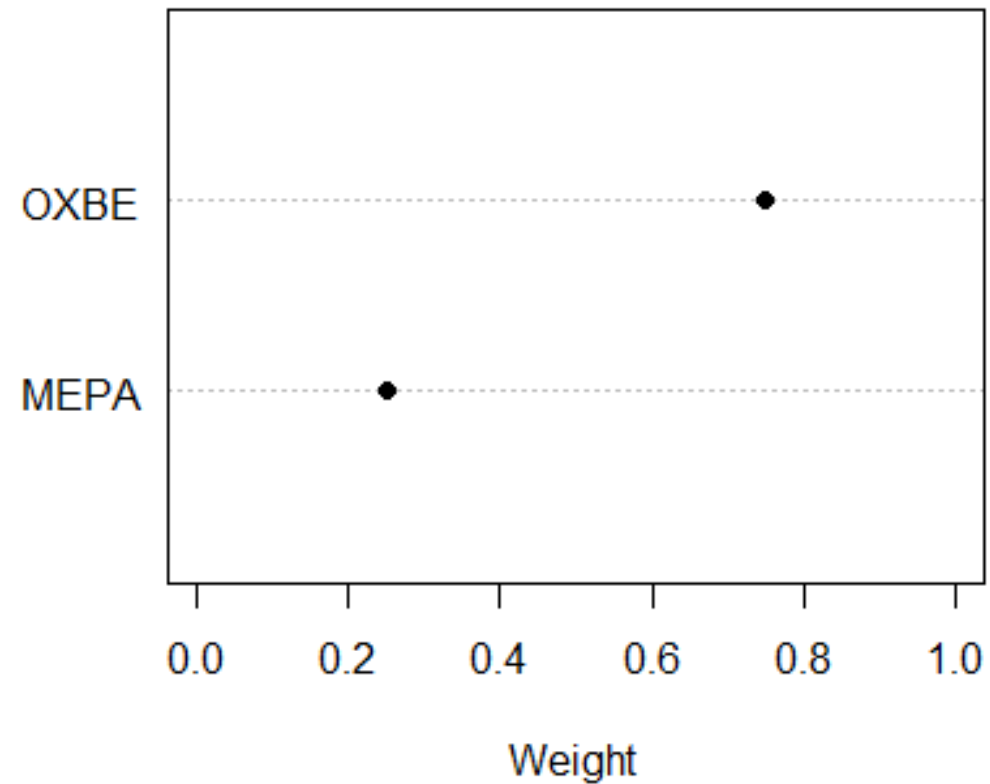
Weights for Metals (+)



Exposome Data Analysis

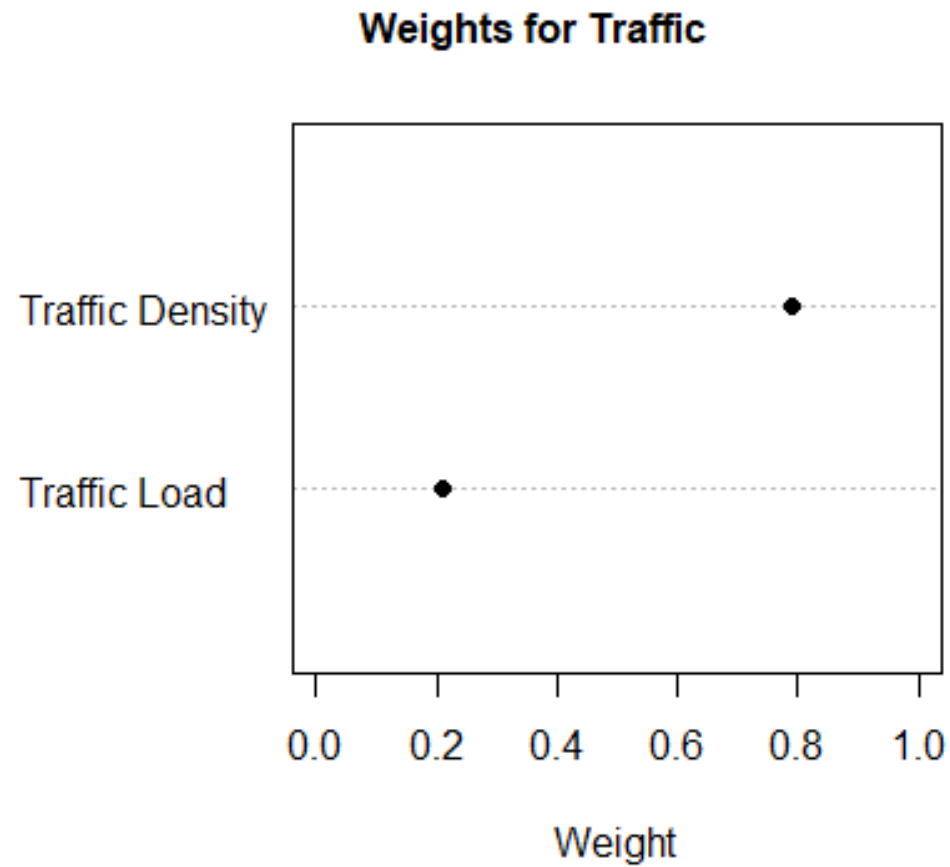
Cohort 1: OR = 2.23 (1.01, 5.26)

Weights for Phenols (+)



Exposome Data Analysis

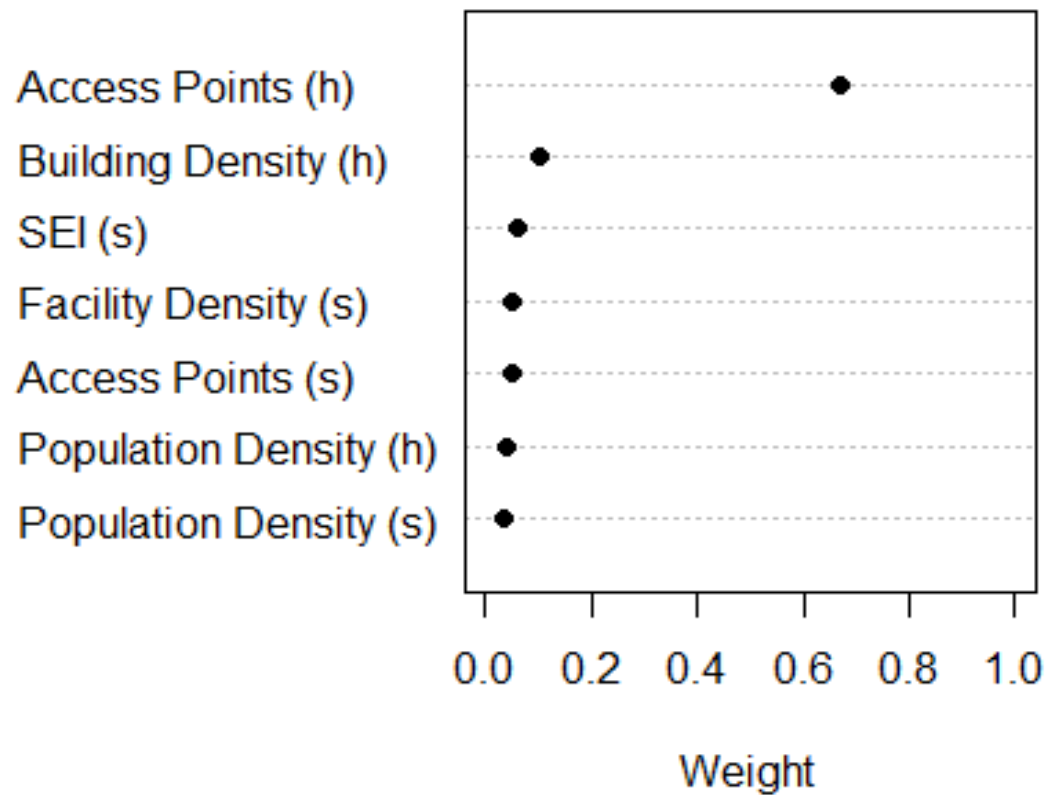
Cohort 5: OR = 2.03 (1.01, 4.12)



Exposome Data Analysis

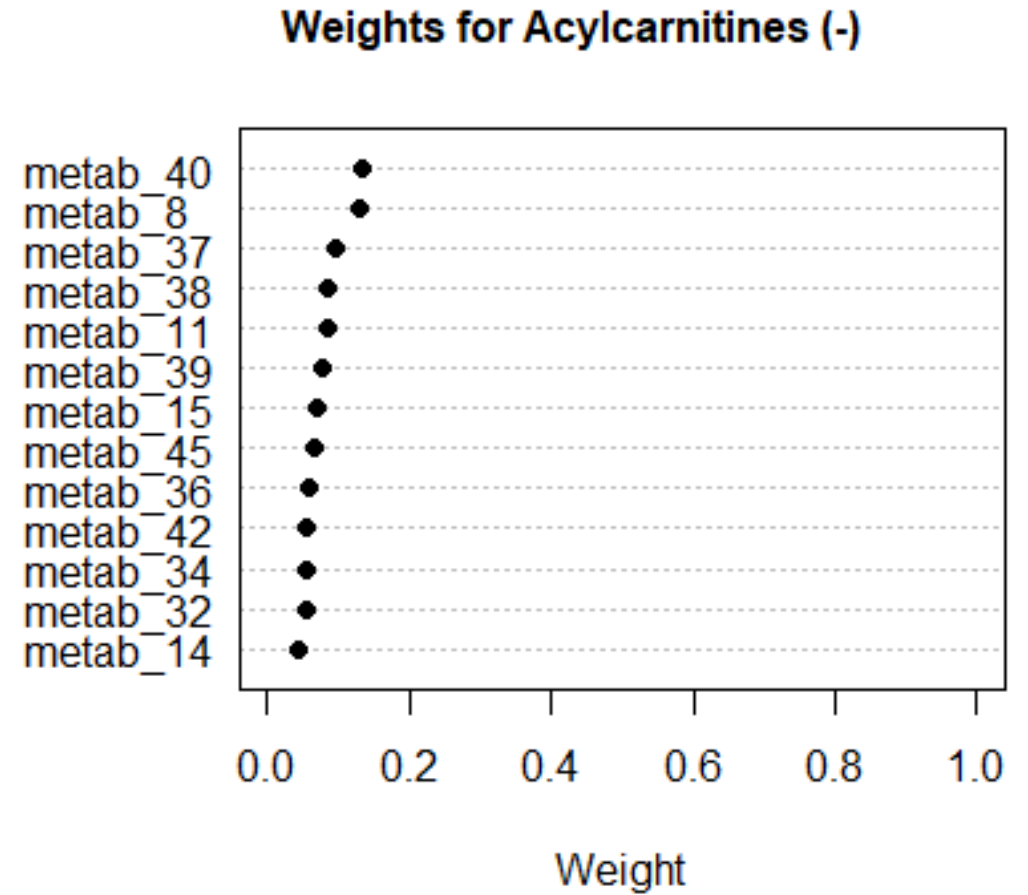
Cohort 5: OR = 0.17 (0.05, 0.43)

Weights for Built Environment (-)



Incorporating -omics Data

- Took the Main Model from Challenge 1 and incorporated the serum metabolomics data.
- In addition to original 19 family indices this model included Biogenicamines (-), Biogenicamines (+), Aminoacids (-), Aminoacids (+), Acylcarnitines (-), Acylcarnitines (-), Glycerophospholipids (-), Glycerophospholipids (+), Sphingolipids (-),and Sphingolipids (+) indices.
- Acylcarnitines (-) was significant with OR = 1.52 [1.02, 2.34].



Related References

- Friedman, Michael S, Powell, et al. "Impact of Changes in Transportation and Commuting Behaviors During the 1996 Summer Olympic Games in Atlanta on Air Quality and Childhood Asthma." *JAMA : The Journal of the American Medical Association* 285.7 (2001): 897-905. Web.
- Haimerl, Pascal, Bernhardt, Ulrike, et al. "Inflammatory Macrophage Memory in Nonsteroidal Anti-inflammatory Drug–exacerbated Respiratory Disease." *Journal of Allergy and Clinical Immunology* 147.2 (2021): 587-99. Web.
- Patel, Molini M, Quinn, James W, et al. "Traffic Density and Stationary Sources of Air Pollution Associated with Wheeze, Asthma, and Immunoglobulin E from Birth to Age 5 Years among New York City Children." *Environmental Research* 111.8 (2011): 1222-229. Web.
- Vaidya, Swapnil V, and Kulkarni, Hemant. "Association of Urinary Bisphenol A Concentration with Allergic Asthma: Results from the National Health and Nutrition Examination Survey 2005-2006." *The Journal of Asthma* 49.8 (2012): 800-06. Web.
- Wheeler DC, Carli M. 2020. BayesGWQS: Bayesian Grouped Weighted Quantile Sum Regression. R package. URL: <https://cran.r-project.org/web/packages/BayesGWQS/index.html>.
- Wheeler DC, Rustom S, Carli M, Whitehead T, Ward MH, Metayer C. Assessment of grouped weighted quantile sum regression for modeling chemical mixtures and cancer risk. *International Journal of Environmental Research and Public Health* 2021.
- Wheeler DC, Rustom S, Carli M, et al. Bayesian Group Index Regression for Modeling Chemical Mixtures and Cancer Risk. *International Journal of Environmental Research and Public Health*. 2021 Mar;18(7). DOI: 10.3390/ijerph18073486.
- Wittczak, Tomasz, Dudek, Wojciech, et al. "Occupational Asthma Due to Manganese Exposure: A Case Report." *International Journal of Occupational Medicine and Environmental Health* 21.1 (2008): 81-83. Web

Thank you!

Any questions?

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