Traffic-Related Air Pollution and Biomarkers of Effect in Children in Fresno

John R. Balmes, MD
University of California, San Francisco and
Berkeley





Disclosures



- I have no financial conflict of interest
- I am the Physician
 Member of the
 California Air Resources
 Board

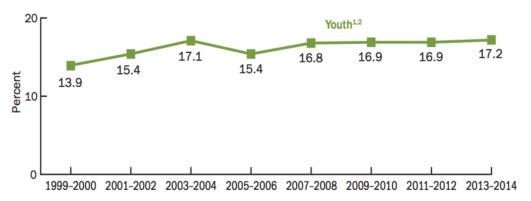


Epidemic of Obesity in U.S.

- 38% of Americans are overweight (BMI 25-29.9) and 35% are obese (BMI≥30)
- 17% of children (ages 2-19) are obese
 - Latino: 21.9%; African-American: 19.5%
 - ages 2-5: 8.9%
 - ages 6-11: 17.5%
 - ages 12-19: 20.5%



Trends in obesity prevalence among youth aged 2–19 years: United States, 1999–2000 through 2013–2014





Obesity Epidemic

- Thought to be due primarily to
 - increased caloric intake from high consumption of sugar-containing drinks and high-caloric-density fast food
 - decreased energy expenditure from a sedentary lifestyle
- Animal evidence suggests that chemicals in the environment may be "obesogens"
 - Organochlorines
 - Bisphenol A







Other Environmental Exposures and Obesity

- Dietary composition
- Gut microbiome
- Built environment through its role in exercise
 - and food consumption
- ? Exposure to ambient air pollutants
 - -? in utero/early childhood





Air Pollution and Obesity

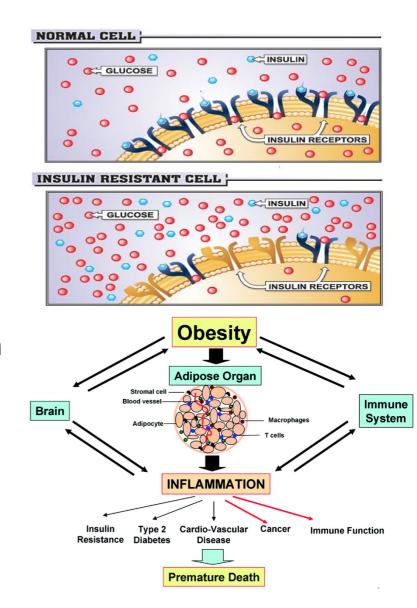
Several studies of children in LA and NYC have shown associations with traffic-related air pollution (TRAP)





Linkage of Diabetes and Obesity

- Type 2 diabetes is a disorder of glucose metabolism
 - the body's cells fail to take up glucose from the blood due to insulin resistance
- 80% of those who develop type 2 diabetes are obese
 - obesity is associated with insulin resistance and β-cell failure
- Both diabetes and obesity are associated with increased systemic inflammation



Air Pollution and Diabetes

- Several studies have shown associations between diabetes in adults and exposure to traffic-related air pollution (TRAP)
- Recent studies from USC showed TRAP exposure in minority children was associated with higher fasting glucose and insulin resistance





Potential Mechanism

- Air pollution can induce oxidative stress and systemic inflammation
- Exposure to fine particles induced fat tissue inflammation and insulin resistance in a mouse model of diet-induced obesity
- Hypothesis:
 - Exposure to air pollution in utero and in early childhood increases risks of abnormal glucose metabolism and obesity in later childhood











Fresno Air Pollution



·	Jnhealthy	Air
Metro Area	Ozone Rank	Particle Rank
Bakersfield	2	1
Fresno- Madera	3	2 (tie)
Visalia- Porterville- Hanford	4	2 (tie)
Modesto- Merced	6	4





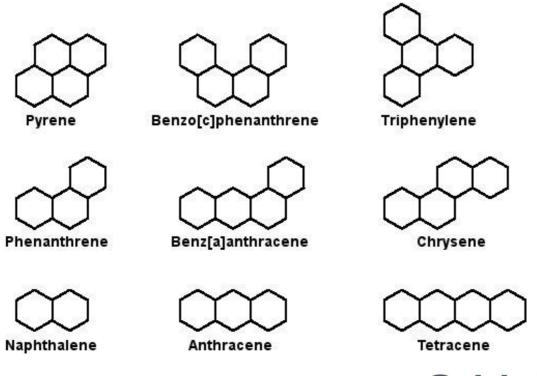
Does Air Pollution Contribute to Obesity and Glucose Dysregulation in Children?





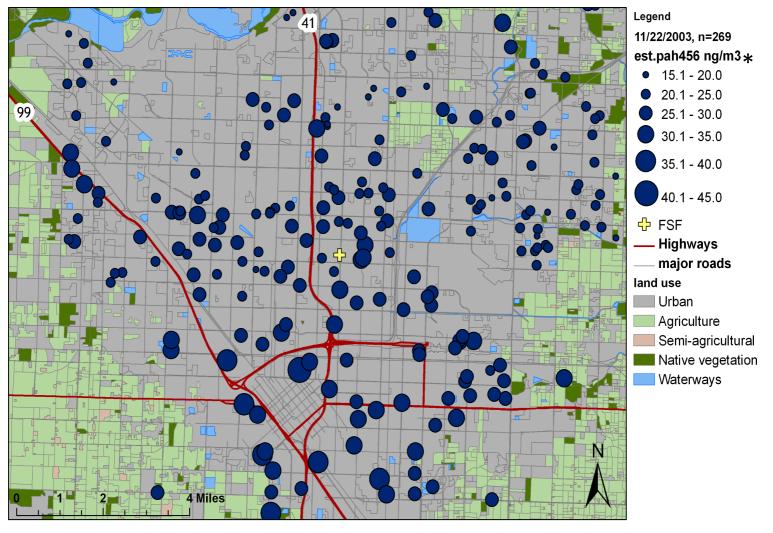
Polycyclic Aromatic Hydrocarbons (PAHs)

 PAHs are formed by incomplete combustion of carbon-containing materials (wood, coal, diesel, gas; also cooked food and tobacco smoke)





PAH cumulative exposure in Fresno





Research Question

- Are exposures to ambient air pollutants, especially PAHs, associated with:
 - increased BMI
 - biomarkers of oxidative stress, systemic inflammation, and abnormal fat and glucose metabolism
 - increased blood pressure



CHAPS Study Design

- Birth cohort, including cord blood; annual visit ages 1 and 2; n=200)
- Child cohort (age 7, follow-up visit at age 9; n=300)
- For all participants
 - Anthropometry and biomarker measurements
 - Estimated air pollution exposure (elemental carbon [EC], nitrogen dioxide $[NO_2]$, nitrogen oxides $[NO_x]$, PAH456*, fine particulate matter $[PM_{2.5}]$, and carbon monoxide [CO])



Description of Child Cohort (n=299)

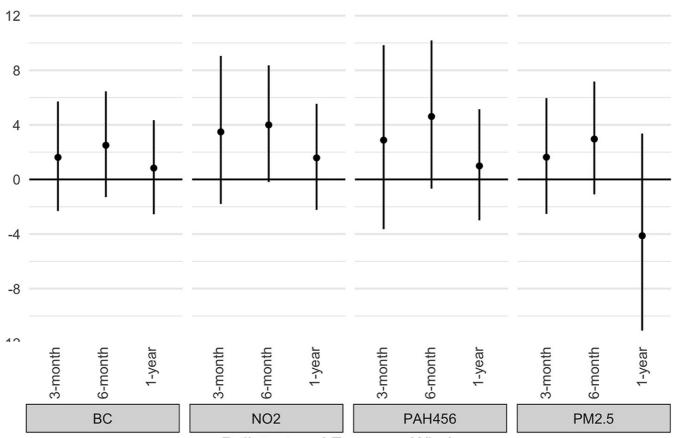
Characteristic	Mean (SD)	%
Age (months)	95.6 (7.0)	
Weight (kg)	31.5 (8.8)	
Male		53.2
Race/Ethnicity		
Hispanic/Latinx		79.6
Black		11.4
Non-Hispanic White		6.0
Asian/Pacific Islander		3.0
Primarily Spanish-speaking		17.4
Renter*		78.0
Annual household income <\$15K*		28.0
Obese**		24.8
Overweight**		16.1

^{*}Responses were refused, not applicable or unknown for n=3 for each of the home renting and household income questions **Using age-and sex-specific percentiles of the 2000 CDC growth charts, obese was defined as BMI \geq 95th percentile and overweight was defined as BMI 85th to <95th percentiles.



BMI-percentile

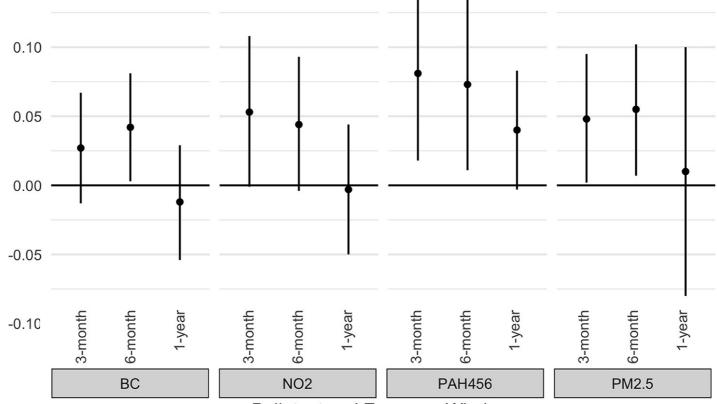
(B) Estimated % change in BMI relative to the 95th percentile





Hemoglobin A1c

(A) Estimated change in HbA1C (%)



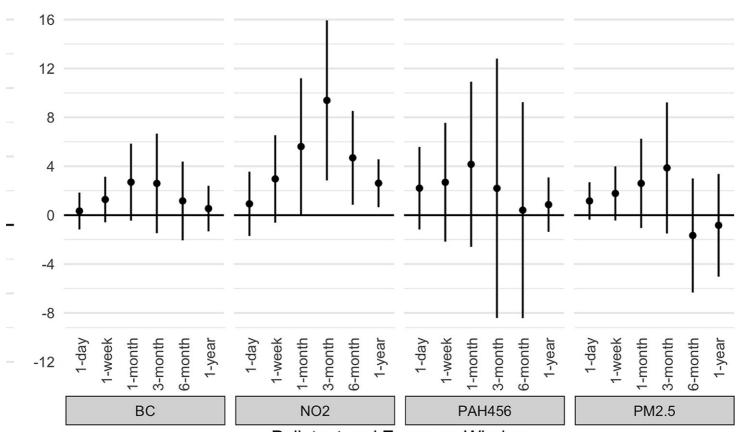
Pollutant and Exposure Window



0.15

Systolic Blood Pressure

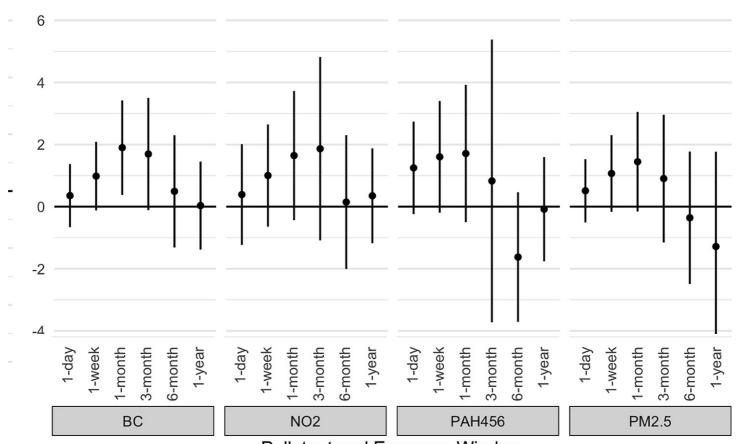
(D) Estimated change in Systolic Blood Pressure (mmHg)





Diastolic Blood Pressure

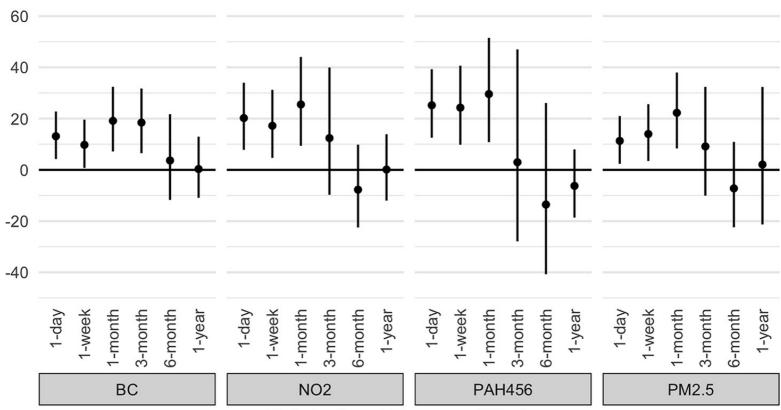
(E) Estimated change in Diastolic Blood Pressure (mmHg)





Urinary 8-isoprostane

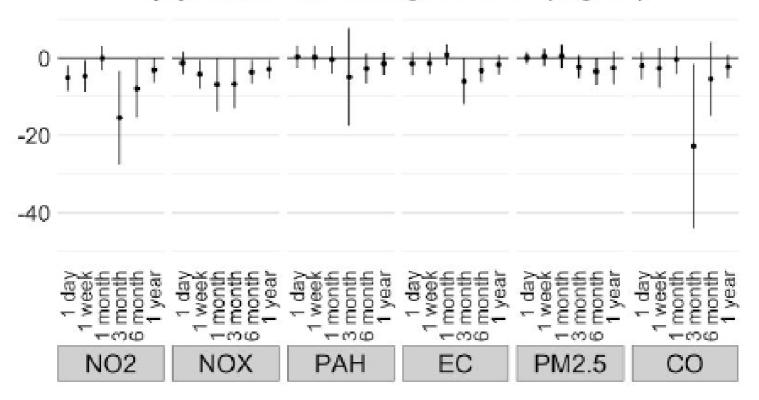
(F) Estimated % change in creatinine-adjusted 8-Isoprostane (ng/mg)





High-Density Lipoprotein (Age 9, n=182)

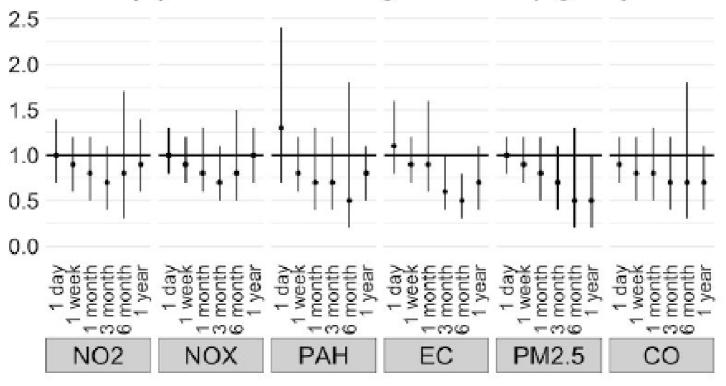
(A) Estimated change in HDL (mg/dL)





Urinary Club Cell Protein-16 (Age 9, n=122)

(D) Estimated change in CC16 (ng/mL)





Summary

- The prevalence of both obesity and pre-diabetes is high among Latinx youth in the SJV
- Air pollution may increase the risk of both conditions by inducing oxidative stress, airway inflammation, and ? systemic inflammation
- Childhood exposure to TRAP is associated with outcomes consistent with increased risk of metabolic syndrome (BMI, HbA1c, HDL, and BP)





Acknowledgements

Jennifer Mann

Helene Margolis

Liza Lutzker

Ellen Eisen

Andreas Neophytou

Sadie Costello

Tim Tyner

Mary Prunicki

Kari Nadeau

Betsey Noth

Kathie Hammond

Fred Lurmann

Gwen Tindula

Nina Holland

Stephanie Holm

Amy Zhang













Thank you





