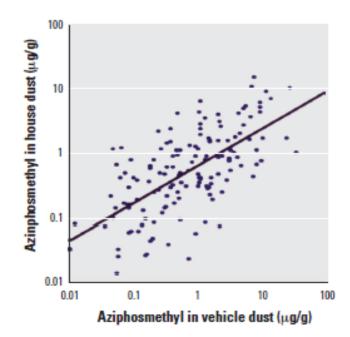
# Dietary Exposure to Agricultural Pesticides (are you what you eat?)

Cynthia Curl, PhD Assistant Professor, Department of Community and Environmental Health Boise State University

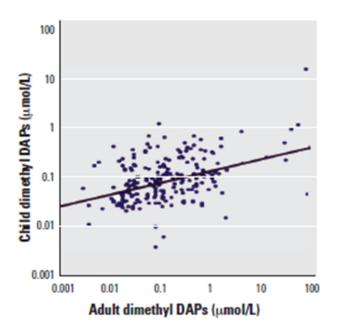
## **Evaluation of Take-Home Organophosphorus Pesticide Exposure among Agricultural Workers and Their Children**

Cynthia L. Curl,<sup>1</sup> Richard A. Fenske,<sup>1</sup> John C. Kissel,<sup>1</sup> Jeffry H. Shirai,<sup>1</sup> Thomas F. Moate,<sup>2</sup> William Griffith,<sup>1</sup> Gloria Coronado,<sup>3</sup> and Beti Thompson<sup>3,4</sup>

<sup>1</sup>Department of Environmental Health, School of Public Health and Community Medicine, University of Washington, Seattle, Washington, USA; <sup>2</sup>SNBL USA Ltd., Everett, Washington, USA; <sup>3</sup>Cancer Prevention Research Program, The Fred Hutchinson Cancer Research Center, Seattle, Washington, USA; <sup>4</sup>Department of Health Services, School of Public Health and Community Medicine, University of Washington, Seattle, Washington, USA



**Figure 1.** Plot of azinphosmethyl concentrations (n = 145) in vehicle dust  $(\mu g/g)$  versus azinphosmethyl concentrations in house dust  $(\mu g/g)$  by household  $(r^2 = 0.41; p < 0.0001)$ . This plot includes samples with residue levels below the LOQ.



**Figure 2.** Plot of urinary dimethyl DAP concentrations (n = 206) in adult urine (µmol/L) versus child urine (µmol/L) by household ( $r^2 = 0.18$ ; p < 0.0001). Creatinine-adjusted results are similar ( $r^2 = 0.15$ ; p < 0.0001). This plot includes samples with metabolite levels below the LOQ.

## Biologic Monitoring to Characterize Organophosphorus Pesticide Exposure among Children and Workers: An Analysis of Recent Studies in Washington State

Richard A. Fenske,<sup>1</sup> Chensheng Lu,<sup>2</sup> Cynthia L. Curl,<sup>3</sup> Jeffry H. Shirai,<sup>1</sup> and John C. Kissel<sup>1</sup>

<sup>1</sup>Department of Environmental and Occupational Health Sciences, School of Public Health and Community Medicine, University of Washington, Seattle, Washington, USA; <sup>2</sup>Department of Environmental and Occupational Health, Rollins School of Public Health,

## **Expected**

Adult farmworkers Kids of pesticide applicators Kids of farmworkers = Kids in farming communities Kids in Seattle

\*During "non-spray" seasons



### Prohibits use of:

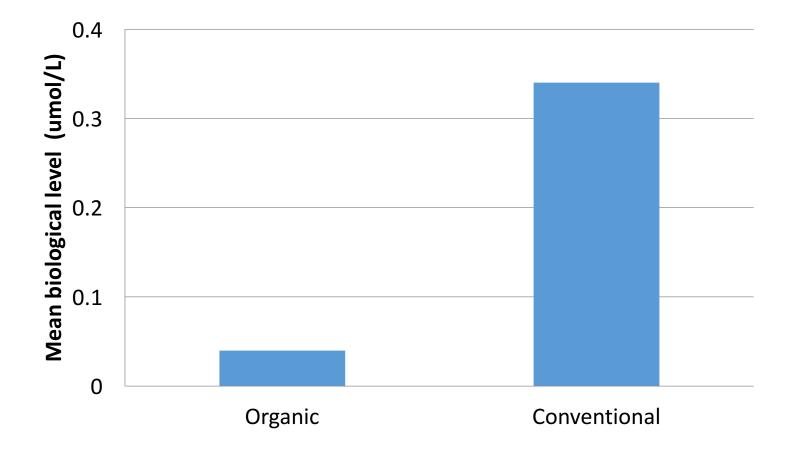
- Most synthetic pesticides, including organophosphates
- Synthetic fertilizers
- Genetic modification
- Irradiation
- Human biosolids ("sewage sludge")
- Antibiotics/synthetic hormones

## Children's Health Articles

#### Organophosphorus Pesticide Exposure of Urban and Suburban Preschool Children with Organic and Conventional Diets

#### Cynthia L. Curl, Richard A. Fenske, Kai Elgethun

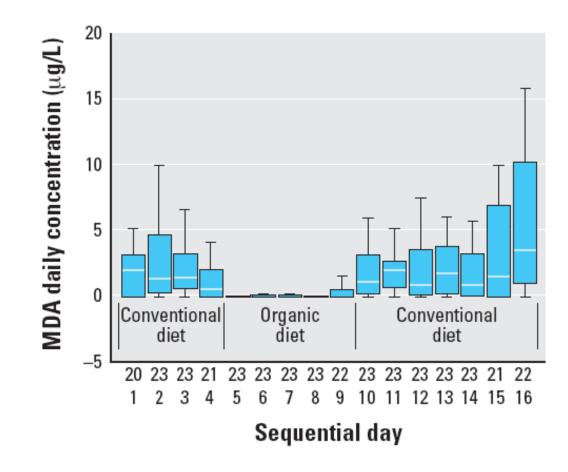
Department of Environmental Health, School of Public Health and Community Medicine, University of Washington, Seattle, Washington USA



## Organic Diets Significantly Lower Children's Dietary Exposure to Organophosphorus Pesticides

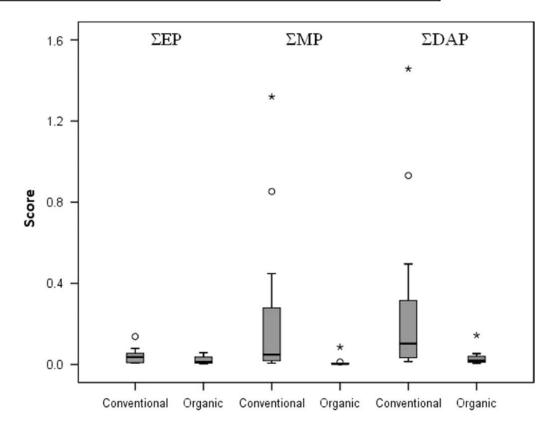
#### Chensheng Lu,<sup>1</sup> Kathryn Toepel,<sup>2</sup> Rene Irish,<sup>2</sup> Richard A. Fenske,<sup>2</sup> Dana B. Barr,<sup>3</sup> and Roberto Bravo<sup>3</sup>

<sup>1</sup>Department of Environmental and Occupational Health, Rollins School of Public Health, Emory University, Atlanta, Georgia, USA; <sup>2</sup>Department of Environmental and Occupational Health Sciences, University of Washington, Seattle, Washington, USA; <sup>3</sup>National Center for Environmental Health, Centers for Disease Control and Prevention, Atlanta, Georgia, USA





Liza Oates <sup>a,\*</sup>, Marc Cohen <sup>a</sup>, Lesley Braun <sup>b,1</sup>, Adrian Schembri <sup>c</sup>, Rilka Taskova <sup>d</sup>



**Fig. 3.**  $\Sigma DAP$ ,  $\Sigma MP$  and  $\Sigma EP$  (creatinine corrected). Mild outliers are marked with a circle (O) and extreme outliers are marked with an asterisk (\*) on the boxplot.



The New York Times

ENVIRONMENT

Study of Organic Crops Finds Fewer Pesticides and More Antioxidants

Time Magazine

NUTRITION

## Is It Worth Buying Organic? Maybe Not

New research questions whether organic produce and meats are really more nutritious or healthier than conventional varieties (MORE: Does Organic Food Turn You into a Jerk?)

# Maternal/neonatal exposure

- Mother-child cohort studies suggest that low-level pesticide exposure *in utero* can have neurological and cognitive effects
- Unknown if pesticide exposure due to conventional diet is substantial enough to cause these detriments
- Unknown if organic diet could result in measurable health benefit

#### Research Children's Health

### Seven-Year Neurodevelopmental Scores and Prenatal Exposure to Chlorpyrifos, a Common Agricultural Pesticide

#### Virginia Rauh,<sup>1</sup> Srikesh Arunajadai,<sup>2</sup> Megan Horton,<sup>3,4</sup> Frederica Perera,<sup>4</sup> Lori Hoepner,<sup>4</sup> Dana B. Barr,<sup>5</sup> and Robin Whyatt<sup>4</sup>

<sup>1</sup>Heilbrunn Center for Population and Family Health, Mailman School of Public Health, <sup>2</sup>Department of Biostatistics, Mailman School of Public Health, <sup>3</sup>Sergievsky Center, and <sup>4</sup>Columbia Center for Children's Environmental Health, Mailman School of Public Health, Columbia University, New York, New York, USA; <sup>4</sup>Enrory University, Atlanta, Georgia, USA

BACKGROUND: In a longitudinal birth cohort study of inner-eity mothers and children (Columbia Center for Children's Environmental Health), we have previously reported that prenatal exposure to chlorpyrifos (CPF) was associated with neurodevelopmental problems at 3 years of age.

**OBJECTIVE:** The goal of the study was to estimate the relationship between prenatal CPF exposure and neurodevelopment among cohort children at 7 years of age.

METHODS: In a sample of 265 children, participants in a prospective study of air pollution, we measured prenatal CPF exposure using umbilical cord blood plasma (picograms/gram plasma) and 7-year neurodevelopment using the Wechsler Intelligence Scale for Children, sth edition (WISC-IV). Linear regression models were used to estimate associations, with covariate selection based on two alternate approaches.

RESULTS: On average, for each standard deviation increase in CPF exposure (4.61 pg/g), Full-Scale intelligence quotient (1Q) declined by 1.4% and Working Memory declined by 2.8%. Final covariates included maternal educational level, maternal 1Q, and quality of the home environment. We found no significant interactions between CPF and any covariates, including the other chemical exposures measured during the prenatal period (environmental tobacco smoke and polycyclic aromatic hydrocarbons).

CONCLUSIONS: We report evidence of deficits in Working Memory Index and Full-Scale IQ as a function of prenatal CPF exposure at 7 years of age. These findings are important in light of continued widespread use of CPF in agricultural settings and possible longer-term educational implications of early cognitive deficits.

KEY WORDS: chlorpyrifos, neurodevelopment, pesticides. *Environ Health Perspect* 119:1196–1201 (2011). doi:10.1289/ehp.1003160 [Online 21 April 2011]

Each year, thousands of new chemicals are released in the United States, with very little documentation about potential long-term human health risks (Landrigan et al. 2002). First registered in 1965 for agricultural and pest control purposes, chlorpyrifos (CPF; 0,0-diethyl-0-3,5,6-trichloro-2-pyridyl phosphorothioate) is a broad-spectrum, chlorinated organophosphate (OP) insecticide. Before regulatory action by the U.S. Environmental Protection Agency (EPA) to phase out residential use beginning in 2000, CPF applications were particularly heavy in urban areas, where the exposed populations included pregnant women (Berkowitz et al. 2003; Whyatt et al. 2002, 2003). In a sample of pregnant women in New York City (Perera et al. 2002) detectable levels of CPF were found in 99.7% of personal air samples, 100% of indoor air samples, and 64-70% of blood samples collected from umbilical cord plasma at delivery (Whyatt et al. 2002)

Early concerns about the possible neurotoxicity of OP insecticides for humans derived from rodent studies showing that prenatal and early postnatal exposures to CPF were associated with neurodevelopmental deficits, and these effects have been seen at exposure levels well below the threshold for systemic toxicity caused by cholinestrase inhibition

in the brain (e.g., Slotkin and Seidler 2005). Evidence has accumulated over the past decade showing that noncholinergic mechanisms may play a role in the neurotoxic effects of CPF exposure in rodents, involving disruption of neural cell development, neurotransmitter systems (Aldridge et al. 2005; Slotkin 2004), and synaptic formation in different brain regions (Qiao et al. 2003). Such developmental disruptions have been associated with later functional impairments in learning, short-term working memory, and long-term reference memory (Levin et al. 2002).

In humans, OPs have been detected in amnionic fluid (Bradman et al. 2003) and are known to cross the placenta (Richardson 1995; Whyatt et al. 2005), posing a threat to the unborn child during a period of rapid brain development. Using urinary metabolites as the biomarker of exposure, several different birth cohort studies have reported that prenatal maternal nonspecific OP exposure was associated with abnormal neonatal reflexes (Engel et al. 2007; Young et al. 2005), mental deficits and pervasive development disorder at 2 years (Eskenazi et al. 2007), and attention problem behaviors and a composite attention-deficit/hyperactivity disorder indicator at 5 years of age (Marks

et al. 2010).

Using a different biomarker of exposure (the parent compound of CPF in umbilical cord plasma), we have previously reported (in the same cohort as the present study) significant associations between prenatal exposure to CPF (> 6.17 pg/g) and reduced birth weight and birth length (Whyatt et al. 2004), increased risk of small size for gestational age (Rauh V, Whyatt R, Perera F, unpublished data), increased risk of mental and motor delay (< 80 points) and 3.5- to 6-point adjusted mean decrements on the 3-year Bayley Scales of Infant Development (Rauh et al. 2006), and evidence of increased problems related to attention, attention deficit hyperactivity disorder, and pervasive developmental disorder as measured by the Child Behavior Checklist at 2-3 years (Rauh et al. 2006). Taken together, these prospective cohort studies show a consistent pattern of early cognitive and behavioral deficits related to prenatal OP exposure, across both agricultural and urban populations, using different biomarkers of prenatal exposure.

We undertook the present study to identify the developmental consequences of prenatal exposure to CPF in a sample of New York City children at 7 years of age. Given the mechanisms proposed in the rodent literature, and early findings from prospective human studies involving nonspecific OP exposures, we hypothesized that prenatal exposure to CPF would be associated with

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1289/chp.1003160 via http://dx.doi.org/). We are grateful to the families of northern Manhattan who have so generously contributed their time and effort to the study.

This study was supported by the National Institute of Environmental Health Sciences (grants 5P01ES09600, P50ES015905, and 5R01ES08977), the U.S. Environmental Protection Agency (grants R82/027, 8260901, and RR00615), the Educational Foundation of America, the John and Wendy Neu Family Foundation, the New York Community Trust, and the Trustees of the Blanchette Hooker Rockefeller Fund.

The authors declare they have no actual or potential competing financial interests.

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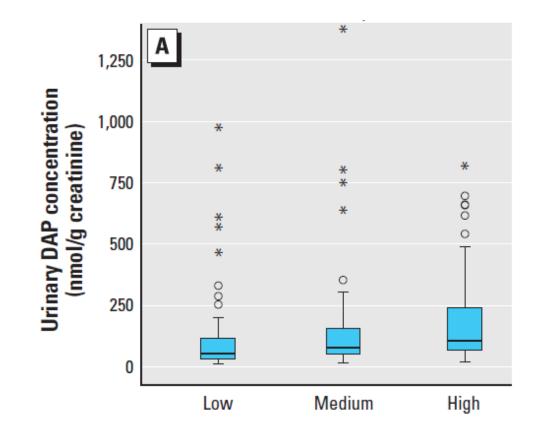
Remaining uncertainties regarding longterm exposure in vulnerable populations

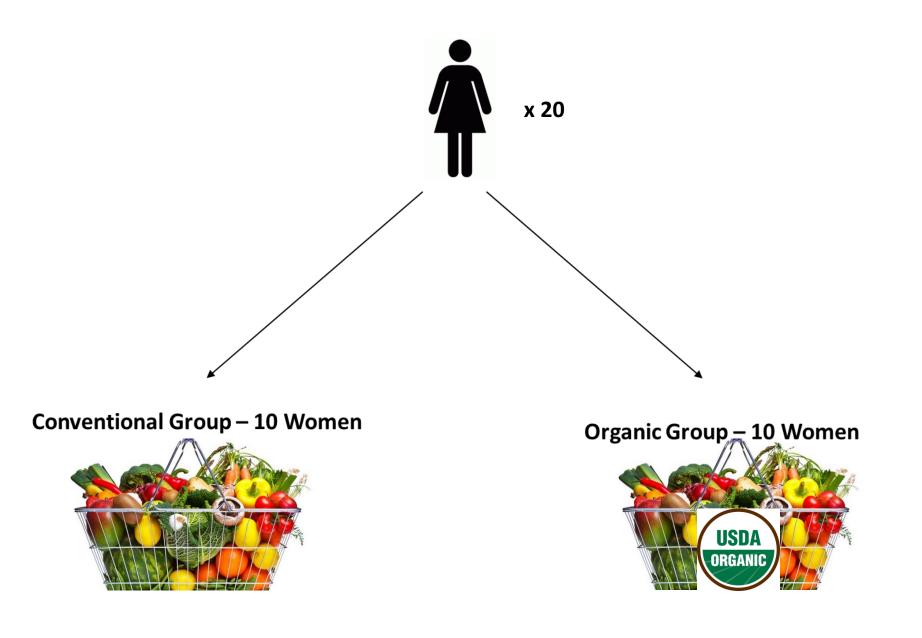
- Chronic vs acute exposure measurement
  - Short half-lives
  - Short interventions
- Realistic dietary interventions
  - Most "organic consumers" don't eat 100% organic
- Vulnerable population
  - No organic intervention studies to date in pregnant women
- Confounding
  - Observational studies are challenged by factors associated with the choice to consume organic

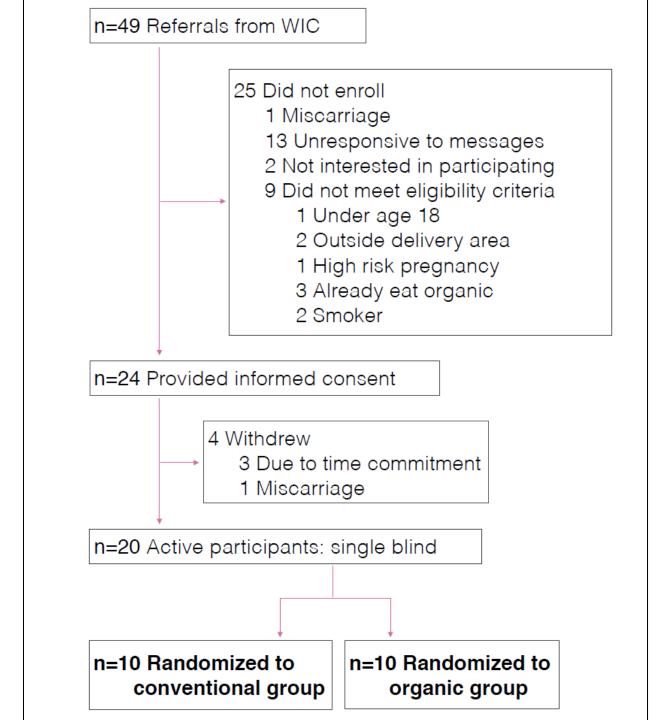
## Estimating Pesticide Exposure from Dietary Intake and Organic Food Choices: The Multi-Ethnic Study of Atherosclerosis (MESA)

Cynthia L. Curl,<sup>1</sup> Shirley A.A. Beresford,<sup>2</sup> Richard A. Fenske,<sup>1</sup> Annette L. Fitzpatrick,<sup>2</sup> Chensheng Lu,<sup>3</sup> Jennifer A. Nettleton,<sup>4</sup> and Joel D. Kaufman<sup>1,2,5</sup>

<sup>1</sup>Department of Environmental and Occupational Health Sciences, and <sup>2</sup>Department of Epidemiology, University of Washington, Seattle, Washington, USA; <sup>3</sup>Department of Environmental Health, Harvard University, Boston, Massachusetts, USA; <sup>4</sup>Department of Epidemiology, Human Genetics, and Environmental Sciences, University of Texas Health Science Center, Houston, Texas, USA; <sup>5</sup>Department of Medicine, University of Washington, Seattle, Washington, USA









BOUT ADMISSIONS ACADEMICS RES

/ PROTECTED: PRODUCE ORDER FORM

#### PROTECTED: PRODUCE ORDER FORM

#### Produce Delivery Order

STEP 2 OF 2	
Total	
\$0.00	
Fruits	
Apples - Gala (	3 pound bag, approximately x apples)
Price: \$4.16	Quantity:
Apples - Red D	elicious (3 pound bag, approximately x apples)
Price: \$3.50	Quantity:
Banana (6 cour	t) (Irr
Price: \$4.34	Quantity:
Blueberries (1	container)
Price: \$3.33	Quantity:
Cantaloupe (1	cantaloupe)
Price: \$2.25	Quantity:
Grapes - Green	(2 pound bag)
Price: \$5.00	Quantity:

- \$20-worth of fresh fruit and vegetables delivered weekly to participants homes
- Deliveries continue throughout second and third trimester (up to \$480 total)
- Partnership with local produce distributors and delivery service
- Convenient and easy for participants

# Developed a study-specific food diary app

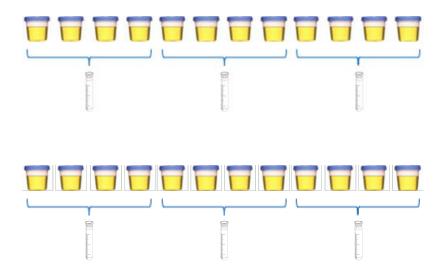
- Participants captured:
  - amount and type of all produce consumed (including photos)
  - whether it was provided by the study or not
  - and if not, whether it was conventional or organic
- Used to calculate number of conventional and organic servings of each type of produce consumed during the study



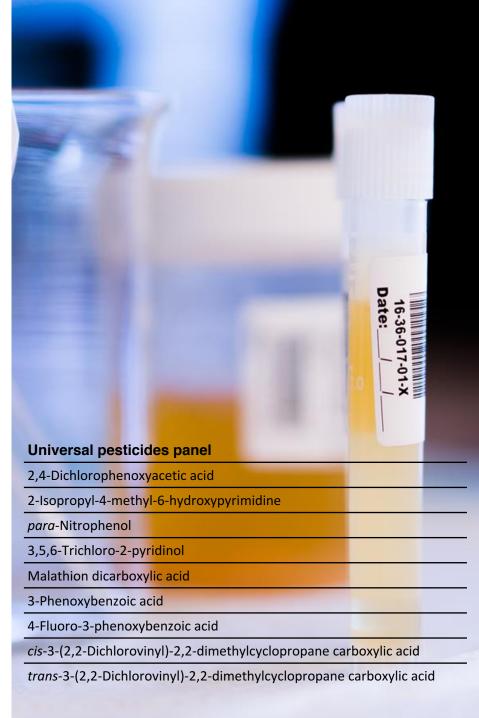
We collected 461 samples from 20 women over a 1-year period, representing an average of 23 weeks (~6 months) per participant.

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25				•						•	•		•			•	•				
20 mples						•														•	
Number of Samples 51																					
2 10																					
5																					
0	<ul><li>1</li></ul>	2	3	4	5	6	7	8	9 Par	<ul> <li>10</li> <li>rticip</li> </ul>	11	• 12 : ID	<b>•</b> 13	<b>•</b> 14	<b>•</b> 15	<b>•</b> 16	<ul> <li>17</li> </ul>	<b>9</b> 18	<b>9</b>	20	21

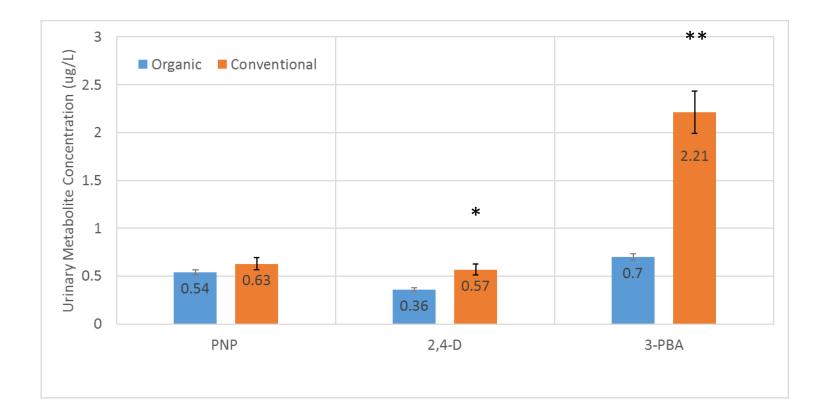




24 weekly samples composited in 1 mL aliquots to create six "monthly" aggregate samples representing the second and third trimesters



Provision of organic fruits and vegetables *significantly* decreased exposure to 2,4-D (an herbicide) and 3-PBA (a metabolite of pyrethroid insecticides)



Curl et al. (Work in progress)

## Conclusions

- Consumption of an organic diet significantly reduces exposure to synthetic agricultural pesticides, including herbicides and several classes of insecticides
  - This is true even when the diet is only partially organic
- For populations without occupational or residential sources, diet is the dominant route of exposure to several classes of synthetic agricultural pesticides
- This work demonstrates the feasibility of a randomized organic diet intervention study among pregnant women, which may be the best way to evaluate the health effects of dietary exposure to agricultural pesticides in conventionally grown food

Unpublished work presented here was funded by the Institute for Translational Health Sciences at the University of Washington.