#### PEHSU NATIONAL Pediatric Environmental Health Specialty Units CLASSROOM

www.pehsu.net/nationalclassroom.html



### Riesgos Ambientales: Pesticidas Environmental Risks: Pesticides

Beatriz Tapia, MD, MPH, EdD

\*Adapted from Environmental Impacts on Reproductive Health: Pesticide Exposure



# **Objectives**

- Discuss the impact of pesticide exposure during pregnancy.
- Identify critical windows of susceptibility for pesticide exposures.
- Describe pesticide exposures among pregnant women in South Texas.
- Identify resources to facilitate patient counseling of risk and risk reduction of hazardous exposures.

# **Pesticides Defined**

# Chemicals used against unwanted organisms:

Insects Rodents Plants Fungi



## **Pesticide Use Is Widespread in United States**



### More than 1.2 billion pounds used annually

Kiely T, et al. 2004 US EPA, 2008



# A well-known harmful pesticide



# **Types of Pesticides**

# Examples:

Type of Pesticide	Target Organisms	Chemical Class
Insecticide	Aphids, fleas, beetles	Organophosphates, pyrethroids
Rodenticide	Mice, rats	Coumarins
Herbicide	Invasive grasses	Acetyl-coenzyme A carboxylase inhibitors

Délye C, et al. Plant Physiol, 2005 EPA, 2008

# **Sources of Pesticide Exposure**



Residues on food



Tap water

EPA, 2008 NPIC, 2008



Community applications



Household use



Occupational exposure



Personal use

# Household Use Is Common

**75%** of US households use at least 1 pesticide product indoors.

May be used to: Eliminate insects and rodents Care for lawn and garden Prevent fleas and ticks



# **Pesticides Contain Active and Inert Ingredients**



more...

Cox C. Environ Health Perspect, 2006 PANNA, 2008

# Inert Ingredients: Examples of Effects

# Some inert ingredients have been found to:

- Decrease heart rate and blood pressure
- Reduce mitochondrial activity
- Be toxic to human placenta cell cultures

# Data come primarily from animal studies

Cox C. Environ Health Perspect, 2006 Richard S, et al. Environ Health Perspect, 2005

# Routes of Exposure





more...

Klaassen C. In: Casarett & Doull's Toxicology: The Basic Science of Poisons. 7<sup>th</sup> ed. 2007

# Acute vs. Chronic Pesticide Exposure

# Acute exposure

- Increase likelihood to identify the source
- Initial insult can be responsible for various health effects

# Chronic exposure

• Not obvious to patient thus less likely to be reported to physicians as a health concern

### Environmental Exposures and Adverse Pregnancy Outcomes

Birth Outcor	me Pollutant	Author and Year Published	Study Type	Exposure Characterization <sup>a</sup>	Sample Size
Fetal growth	Particulates	Wang et al <sup>28</sup> Bobak <sup>29</sup> Dejmek et al <sup>30</sup> Ha et al <sup>31</sup> Wilhelm and Ritz <sup>32</sup> Parker et al <sup>33</sup>	Retrospective cohort Retrospective cohort Retrospective cohort Time series Retrospective cohort Retrospective cohort	Ambient monitoring Ambient monitoring Ambient monitoring Ambient monitoring Ambient monitoring Ambient monitoring	$\begin{array}{l} n = 74\ 671\\ n = 73\ 148\\ n = 4883\\ n = 276\ 763\\ n = 498\ 235\\ n = 18\ 247 \end{array}$
	Organophosphate	Perera et al <sup>41</sup>	Prospective cohort	Serum levels	n = 263
	insecticides	Levario-Carrillo et al <sup>42</sup>	Case-control	Residential history	n = 371
	Triazine herbicides	Munger et al <sup>43</sup>	Cross sectional	Drinking water monitoring	n = 9551
		Dabrowski et al <sup>44</sup>	Case-control	Residential and occupational history	n = 494
		Villanueva et al <sup>45</sup>	Retrospective cohort	Municipal drinking water monitoring	n = 9721
Preterm birth		Bobak <sup>29</sup> Wilhelm and Ritz <sup>32</sup> Sagiv et al <sup>48</sup> Huynh et al <sup>49</sup>	Retrospective cohort Retrospective cohort Time series Matched case–	Ambient monitoring Ambient monitoring Ambient monitoring Ambient monitoring	$\begin{array}{l} n = 73 \ 148 \\ n = 498 \ 235 \\ n = 187 \ 997 \\ n = 42 \ 692 \end{array}$
	SO <sub>2</sub>	Bobak <sup>29</sup> Liu et al <sup>38</sup> Liu et al <sup>38</sup>	Retrospective cohort Retrospective cohort Retrospective cohort	Ambient monitoring Ambient monitoring Ambient monitoring	$n = 108 \ 173$ $n = 229 \ 085$ $n = 229 \ 085$
ngenital	Chlorophenoxy	Garry et al <sup>50</sup>	Cross sectional	Occupational history	n = 210
abnormalities	herbicides	Garry et al <sup>51</sup>	Cross sectional	Occupational history	n = 1532
		Basso 1999 <sup>52</sup>	Cohort	Occupational history	n = 8671
		Schreinemachers <sup>53</sup>	Ecologic	Pesticide use and	n = 43 6
	Other pesticides	Rull 2006 <sup>54</sup>	Case-control	Pesticide use and	n = 1671
				proximity	
	byproducts	Toledano et al <sup>62</sup>	Cohort	water utility monitoring data Monitored tap water extrapolated to populations	n = 920 571

#### Stillerman, et al. Reprod Sci, 2008

#### Table 1. (continued)

Birth Outcome	Pollutant	Author and Year Published	Study Type	<b>Exposure</b> Characterization <sup>a</sup>	Sample Size
	DDT	Korrick et al <sup>63</sup>	Case–control	Serum levels	n = 30
		Longnecker et al <sup>64</sup>	Prospective cohort	Serum levels	n = 1717
		Law et al <sup>65</sup>	Prospective cohort	Serum levels	n = 390
		Cocco et al <sup>66</sup>	Retrospective cohort	Occupational history	n = 105
		Venners et al <sup>67</sup>	Prospective cohort	Serum levels	n = 388
	Bisphenol-A	Sugiura-Ogasawara et al <sup>68</sup>	Case-control	Serum levels	n = 45

<sup>a</sup> Ambient monitoring for the common air pollutants is assumed to be a reasonable surrogate for individual level exposures.

<sup>b</sup> The study evaluated families with children with birth defects, the n is for the number of children in the study.

#### Stillerman, et al. Reprod Sci, 2008

## **NHANES: Chemicals in Pregnant Women**

#### Table 1. Chemical classes measured in biological tissue of pregnant women, NHANES 2003–2004.

	No. of chemical analytes measured			
Chemical class	Blood	Serum	Urine	Total
Cotinine		1		1
Environmental phenols			4	4
Metals	4			4
Organochlorine pesticides		13		13
Organophosphate insecticides			6	6
Perchlorate			1	1
Phthalates			13	13
PBDEs and other brominated flame retardants		11		11
PCBs and dioxin-like chemicals		55		55
PAHs			10	10
PFCs		12		12
VOCs	33			33

See Supplemental Material, Table 1 (doi:10.1289/ehp.1002727), for individual chemical analytes included in each chemical class.

Woodruff TJ, et al. Environ Health Perspect, 2011

# NHANES: Chemicals in Pregnant Women



**Figure 3.** Number of chemicals detected by chemical class in U.S. pregnant women, NHANES subsample B [metals, cotinine, organochlorine (OC) pesticides, phthalates, brominated flame retardants (PBDEs), and PAHs], 2003–2004 (n = 54). Each vertical bar represents one study participant. Other subsamples showed similar results.

Woodruff TJ, et al. Environ Health Perspect, 2011

# **Reproduction-Related Effects**

"...every child conceived today in the Northern hemisphere is exposed to pesticides from conception throughout gestation and lactation regardless of where it is born."

> Colborn T. Environ Health Perspect. 2006

## Pesticide Exposure During Pregnancy among Hispanic Women in South Texas

# **Research Question:**

How do pesticide exposures of pregnant women living in the Lower Rio Grande Valley compare with those of pregnant women living in New York City (NYC)?

(in partnership with U.S. Hispanic Nutrition Research and Education Center)





Prenatal insecticide exposure, birth weight and length among an urban minority cohort. Whyatt, et al. Environ Health Perspect. 2004 Jul; 112(10):1125-32.

- Inner city of NYC has highest pesticide application rates in NY State, principally used for roach control in low-income housing.
- Measured pesticides in dust, air, maternal blood, etc.

Results: Babies born to mothers exposed to organophosphate pesticides had decreased birth length and head circumference. Currently assessing cognitive/behavioral measures up to age seven. N=314

Whyatt, et al. Environ Health Perspect, 2004

# **Study Population:**

25 pregnant Hispanic Women, 18-35 y/o, 30-34 weeks gestation, recruited from maternity clinics in Hidalgo County

### **Inclusion criteria:**

- Stable residency within drivable distance
- Non smoker/no illicit drugs/moderate drinker
- No major health problems (DM, AHT, HIV, NS)
- Homemakers who spend majority day in home

# **Pesticide Exposure in South Texas**

# **Methods:**

- Questionnaire covering demographics, home characteristics, residential history, and lifestyle
- Installation of PUF sampler in home
- Two weeks later follow-up questionnaire
- All samples (air and dust) sent for analysis



# Pesticides analyzed in home air and dust (~45 total)

#### Organophosphates

- Azinophos-methyl
- Chlorpyrifos
- Diazinon
- Ethyl Parathion
- Malathion
- Methyl parathion
- Propetamophos

#### Carbamates

- Bendiocarb
- Carbaryl
- Carbofuran
- Fenoxycarb (also IGR)
- Propoxur

#### Synergists

- MGK 264
- Piperonyl butoxide

#### Fungicides

Captan

#### Synthetic Pyrethroids

- Bioallethrin
- Bifenthrin
- Cis-permethrin
- Cyfluthrin
- Cypermethrin
- Deltamethrin/tralomethrin
- Fenvalerate
- Lamda-cyhalothrin
- Prallethrin
- Sumithrin
- Tetramethrin
- Trans-permethrin

# **Pesticides Analyzed (continued)**

### Insect Growth Retardant (IGR)

- Fenoxycarb (also carbamate)
- Hydroprene
- Methoprene

### Organochlorines

- 4,4'-DDD
- 4,4,'-DDE
- 4,4'-DDT
- Alpha-chlordane
- Dieldrin
- Gamma-chlordane
- Heptachlor
- Lindane

### Herbicides

- Atrazine
- Metolachlor
- Pendimethalin
- Simazinc
- Trifluralin

### Others

- Fipronil
- Orthophenylphenol
- Sulfluramid

# Pesticide Exposure in South Texas

# **Results:**

68% of these households reported pesticide use vs. 85% of previously studied NYC households

35% used two or more pest control methods

14 pesticides detected including several organophosphates: ortho-phenylphenol in 92% of home air samples, followed by chlorpyrifos in 80%, propoxur in 76%, diazinon in 72%, and trifluralin in 60%



# Pest control methods used and target pests for Texas women who reported pest control measures used in their homes during

#### oregnancy. N=17



steer

# **Conclusions Result cont.**

Household pesticide exposures during pregnancy in South Texas were similar to those in NYC

In both Hidalgo County and NYC, the principal reason for organophosphate pesticide use were **roaches** 

Offspring of mothers exposed in NYC showed significant neurodevelopmental problems

Neurodevelopmental studies among South Texas children exposed to pesticides are lacking

## **Studying Pesticide Exposure Has Inherent Challenges**

# •Limitations impeding research:

- Human trials precluded by ethical considerations
- Difficulties in assessing impacts
- Difficulties in measuring outcomes



### Many Complex Factors Interact to Affect the Impact of Exposures



Adapted from Hubbs-Tait, et al. Psychological Science in the Public Interest, 2005

# Environmental Exposures and Critical Windows of Susceptibility



Woodruff TJ, et al. Fertil Steril, 2008

### **Reproduction-Related Effects: Women**



Bretveld RW, et al. Reprod Biol Endocrinol, 2006

# Reproduction-Related Effects: Women

# Changes in Menstrual Cycle (1.5 ↑ odds )

- Longer cycles
- Missing periods
- Bleeding mid cycle

### **Consequently affecting Fertility!**

Pesticides: Lindane, atrazine and mancozeb

Farr SL, et al. Am J Epidemiol, 2004

### **Reproduction-Related Effects: Men**

# Testicular damage:

- Azoospermia, oligospermia
- Damage to germinal epithelium
- Genetic alterations in sperm
- Reduced fertility
- Altered hormone function



Figà-Talamanca I, et al. Occup Med, 2001 Whorton MD, et al. Lancet, 1977

# **Delayed Effects of Exposure**



Critical windows of susceptibility:

- Preconception
- Prenatal
  - Postnatal (lactation)
  - Childhood and adult cancer
  - Delayed development
  - Childhood asthma and allergies
  - Infections
  - Postnatal growth effects

Wigle DT, et al. J Toxicol Environ Health B Crit Rev, 2008

# The Barker Hypothesis

"Exposures to adverse insults during critical...windows of development can permanently reprogram normal physiologic responses, and thus give rise to...disorders later in life."

> Woodruff TJ, et al. *Fertil Steril.* 2008

Barker DJ. Trends Endocrinol Metab, 2002; Woodruff TK, Walker CL. Fertil Steril, 2008; Woodruff TJ, et al. Fertil Steril, 2008



# **Delayed Development**

Strong evidence implicating pesticides and developmental disorders, including PDD-NOS

Organochlorine exposure in utero

- > abnormal reflexes in neonates
- ↓Psychomotor ↓mental at 12 months
- ↓General cognitive, memory, verbal and executive functions at 4 years

Rosas LG, et al. Currt Opin Pediatr, 2008

# Fetal Origin of Adult Disease

# Parkinson Disease

- ↑ evidence that the environment plays a major role
- Animal studies of faulty neurochemistry in offspring are linked to low level exposure of pesticides.
- Alterations to the CNS may induce a "silent" state of dopamine dysfunction leaving individuals vulnerable

# "So What Do I Do?"

- Science regarding environmental exposures and reproductive health is:
  - Primarily based on animal studies
  - Warrants guidance to limit/avoid exposure
- Approach patients on case-by-case basis
- Exposure is unavoidable, but specific changes can make a difference

# Using your flip chart



#### **Insect Control and Pesticide Use**

- · Pesticides can be highly toxic.
- Seal or caulk crevices to keep insects and rodents out.
- Use boric acid or traps to control pressure







#### **Insect Control and Pesticide Uses**

#### Pesticides

- Pesticides are chemical products that include herbicides, rodenticides, and insecticides. They can be extremely toxic.
   Harm they produce:
- Chronic exposure to pesticides may cause learning and developmental problems in children as well as autism. Adults may develop cancer.
   Prevention:
- Seal or caulk crevices where insects can enter your home.
- Avoid leaving food or dishes to be washed lying out in your kitchen.
- Take out garbage daily.
- Use insect traps and baits instead of sprays wherever possible.
- Use boric acid powder to help control cockroaches in your kitchen. Place it behind the refrigerator, stove, etc., keeping it away from places children or pets can reach.
- Minimize "harborage," or clutter, where insects tend to hide and breed.
- Roaches often obtain water from condensation on pipes, so insulate cold water lines with foam wrap and seal wrap with tape
- Keep flour and sugar in resealable plastic bags.
- If you have many cockroaches, even if they are coming from neighboring residences, consider hiring a professional exterminator who uses integrated pest management (IPM). Baits and traps are preferable to insecticides that add chemicals to the air. Explain your child's condition to the pest control service before allowing anyone to treat you house.

# Using your flip chart

#### **Illegal Pesticides**

#### What are they?

- Illegal pesticides may be more toxic than legal ones and they may even have unknown chemicals as part of their formula.
- Legal pesticides are the ones registered and approved by the LLS Environmental Protection Age
- For example, the "Chinese Cł registered with EPA.
- Manufacturer states that the " any harm to humans or pets.

#### Harm they produce:

 Children may confuse "Chines have been reports of children such as vomit, abdominal pair "Chinese Chalk."

#### Prevention

 Do not purchase illegal pestic your house.

#### **Illegal Pesticides**

#### **Airplane Powder**

- Airplane powder is a very powerful pesticide.
- Chronic exposure may cause irreversible damage to the nervous, cardiovascular, and reproductive organs. It has been reported that it may also cause Parkinson's and Alzheimer's.

#### How to know if a pesticide is legal?

- Check for the EPA registration number on the product's label.
- Purchase products only from authorized vendors and never from a Street vendor.
- In the U. S. call the National Pesticide Information Center at 1-800-858-7378 if you have questions or need more information.
- In Mexico call your local Red Cross or local Health Community Center.



# Focus on Windows of Susceptibility

- For male and female adolescents
- For male and female patients who experience unintended pregnancy
- For women and men during pregnancy planning
- For pregnant women
- For male and female patients with newborns and children

# The Environmental Health History

# Section Incorporate into reproductive health history

Identify and reduce
 or eliminate
 potentially harmful
 exposures

### When?

#### Vulnerable Stages:

- Early childhood
- Puberty
- Adolescence
- Preconception planning (men & women)
- Pregnancy

#### Guide patients in making decisions

Identifying and reducing exposures to potentially harmful toxicants *now*, when couples plan (or not) a pregnancy, increase the likelihood of a successful outcome.



# Association of Reproductive Health Professionals (ARHP) Resources

- •Learn more at the ARHP Website:
  - Click on Environmental and Reproductive Health topic area
  - www.arhp.org/topics/enviro-repro-health
    - Fact Sheet: Environmental and Reproductive Health Resources for Health Care Providers
    - Patient handout: Health Matters: The Connection Between Your Health and the Environment

# Pesticide Resources

- Fact sheets from CDC's National Agricultural Safety Database (http://nasdonline.org/)
- National Pesticide Information Center (http://npic.orst.edu/)
- EPA's Pesticides Information Web site (includes information for children) (www.epa.gov/pesticides)
- Pesticide Action Network (PAN) database (www.pesticideinfo.org)

# **Questions?**

# Thank you!