# The Primary Prevention of Asthma

#### Risk Factors Associated with Asthma Onset

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### Asthma: a complex disease(s)

- Common signs and symptoms: airway narrowing, wheeze, cough, shortness of breath
- Different kinds of asthma; pathophysiology varies
- Implications for primary prevention differ

#### Asthma: Lifecourse perspective



### Key Risk Factors in Asthma Onset

- Pets
- Obesity, Nutrition
- Immigration and Infection
- Prenatal exposures to chemical contaminants
- Psychosocial environment/stress
- Allergens indoors
- Asthmagens in the workplace
- Air pollution
- Tobacco

### Pets in the home

- "Pet exposure" is complex:
  - Allergen exposure
  - Inhaled and ingested microbial exposure
  - Exposure to support or stress (bites)
  - Exercise

### Pets

- Early-life "pet exposure" is <u>probably</u> associated with asthma onset
- But, the evidence for <u>the direction of the</u> <u>association (beneficial, risk factor, no</u> <u>association) varies</u>, and it varies more for cat than for dog ownership.

#### Pets

- Variability in response to pet allergens or microbes likely depends on
  - Dose from pet in the home
  - Timing of exposure
  - Inheritance
  - Background dose of pet allergen in the community
  - The microbial organisms that the pet carries
  - Unmeasured or measured cofactors influencing susceptibility to allergy, airway inflammation, airway irritability and asthma

# Immigration and infection

- Enough studies have been done to show that asthma prevalence is lower in people living in many low-income countries and that it increases after immigration to developed countries.
- The effect sizes are large.
- BUT, the reasons for this are not yet clear.
- Therefore research needs to focus on why.

# Immigration and Infection: hypotheses

- Language, culture, and literacy
- "Hygiene" hypothesis
- Chemicals, including air pollution
- Vitamin D
- Selection bias in immigration

### **Obesity and Nutrition**

### Obesity

- Maternal obesity during pregnancy
- Early childhood obesity
- Adult obesity

#### Nutrition

- Antioxidants (eg, fruits & vegetables)
- omega-3 fatty acids (eg, fish)
- Vitamin D (sunlight + diet/supplements)

### Obesity, nutrition

- **Obesity** across life span:
  - During pregnancy: probably associated
  - Early childhood: probably associated
  - Adult: known association
- Nutrition during pregnancy & early childhood:
  - Antioxidants (fruit & vegetables): <u>possibly</u> associated
  - omega-3 fatty acids (fish): possibly associated
  - Vitamin D (sunlight + intake): <u>not classifiable</u>

# Chemicals

- Metals: <u>Possibly</u> associated with asthma
- Organochlorine Compounds: <u>Not classifiable</u>; inadequate evidence
- Bisphenol A: Possibly associated with asthma
- Perfluorinated Compounds: <u>Not classifiable</u>; inadequate evidence
- Phthalates: <u>Not classifiable</u>; inadequate evidence
- [Magnetic field: <u>Not classifiable</u>; inadequate evidence]

### Psychosocial stress and asthma

- Biological plausibility
- Temporal sequence prospective prenatal & early childhood cohorts a particular strength
- Exposure-response relationship
- Robust to adjustment for a number of important confounders
- Robust to sensitivity analyses
- <u>Known</u> to be associated with early asthma phenotypes

### **Psychosocial stress**

- Socially toxic environments are NOT simply a marker of a more toxic physical environments
- Social contexts and consequent stress may be as detrimental to children's health as chemical toxicants
- "Social pollutants"
- Psychological stress disrupts biological systems overlapping with those altered by physical pollutants/toxicants

### **Psychosocial stress**

- Independent effects
- Interactive (joint) effects
- Individual- and place-based psychosocial stress may impact host resistance such that physical toxicants (e.g., indoor allergens, traffic-related air pollution) may have adverse effects at relatively lower doses
- Epidemiological studies and interventions need to address physical toxicants and social stress jointly to impact public health most effectively

### Indoor allergens

- Exposure to cockroach and mouse allergens: <u>probably</u> associated with asthma onset
- Sensitization to dust mites, cat and dog dander <u>known</u> to be associated with asthma onset
  - Timing of exposure to animal dander is important
  - Most wheezers with no atopy lose their symptoms by school age
- Sensitization to molds <u>known</u> to be associated with asthma onset
  - As with animal dander, differs between atopic and non-atopic wheezers
  - May be variable response to various mold species
- Interactions with stress, chemicals, and outdoor air pollution may be important

### **Occupational** asthma

- Occupational Asthma
  - Sensitizer induced
  - Irritant induced
    - Reactive Airways Dysfunction Syndrome (RADS)
    - Low dose irritant induction of asthma
- Work-exacerbated asthma
  - Irritant exposures
  - Workplace exposures to sensitizers

#### Ten most frequently reported agent categories; state-based surveillance systems



# **Outdoor** air pollution

- Traffic related air pollution:
  - <u>Probably</u> associated with asthma onset
- Ozone and PM 2.5
  - Possibly associated with asthma onset
- Special considerations:
  - Non-specific exposure metrics, (e.g. TRAP and proximity to traffic)
  - Possible synergistic effects with stress and allergens

### Tobacco smoke

- Meets all Bradford Hill criteria for causality
  - Consistency and coherence
  - Strength of association
  - Plausible mechanism(s)
    - Oxidative stress from free radicals, epithelial damage, inflammation
  - Temporality
  - Dose-response relationship
- Thus, tobacco smoke exposure is <u>known</u> to be associated with risk of asthma onset