Chronic Disease: Introductory Concepts

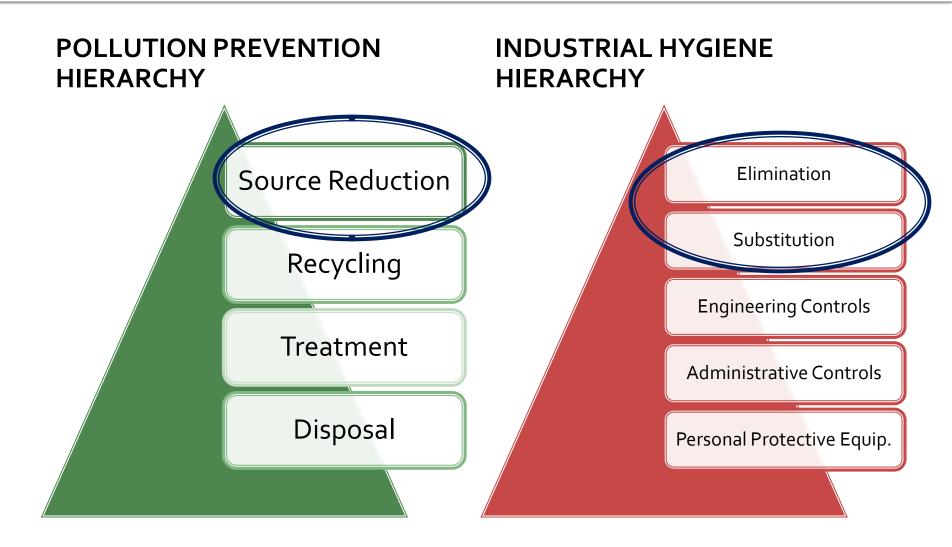
Outline

- Primer on Important Chronic Disease Concepts
 - Primary Prevention: Familiar Territory to TUR Planners
 - Mechanisms of Disease Causation: 2 Useful Metaphors
 - Critical Windows of Development/Timing of Exposure
 - Low Dose Effects
 - Disease Latency

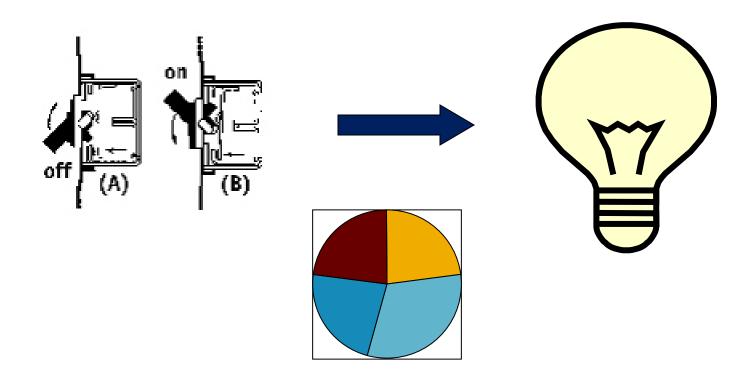
Primary, Secondary & Tertiary Prevention

- Primary Prevention: Interventions that keep diseases from ever developing in the first place
- Secondary Prevention: Interventions to diagnose and treat early stages of a disease before it causes significant morbidity
- Tertiary Prevention: Interventions to reduce the negative impact of disease by restoring function and reducing disease-related complications

Toxics Use Reduction: A Crucial Primary Prevention Strategy



Chronic Disease: A Multi-Factorial, Multi-Stage Process



Disease Causation: Critical Windows of Development/Timing of Exposure







- Examples from cancer studies
 - Woburn MA: Childhood leukemia 8.3 times higher among children whose mothers drank contaminated water while pregnant.
 - Girls exposed to DDT before puberty were 5 times more likely to develop breast cancer when they reach middle age
 - 3. Cancer risk among workers exposed to radiation is greater among older workers than younger worker when exposed to the same dose.

(1) References: <u>Sci Total Environ.</u> 2002 Dec 2;300(1-3):23-35; (2) <u>Environ Health Perspect.</u> 2007 Oct;115(10):1406-14. <u>Am J Ind Med.</u> 1993 Mar;23(3):371-89; <u>Int J Epidemiol.</u> 1999; (3) Jun;28(3):428-36.

Low Dose Effects

- "Low dose": doses below those tested in traditional toxicological assessments
- Primary mechanism: endocrine disruption by mimicking the action of hormones.
 - Small changes in hormone concentration can have biologically important consequences.

Chemical	Endpoint affected at low doses
BPA	Prostate and mammary gland, brain development and behavior, reproduction, immune system, metabolism
Dioxin (TCDD)	Spermatogenesis, immune function and oxidative stress, tooth and bone development, female reproduction, mammary gland, behavior
Perchlorate	Thyroid stimulating hormone levels

Reference: Endocr Rev. 2012 Jun;33(3):378-455

Disease Latency

- Disease latency: "The time interval between disease occurrence and its subsequent detection.*"
 - Important for studying disease-exposure connection
 - Cancer: long latency
 - In adults: Exposure to causal agents likely occurred 10-30 years (depends on the type of cancer) prior to diagnosis.
 Children: latency 0-10 years
 - Sensitizing asthma: shorter latency
 - 1st exposure does not cause disease. Requires sensitizing period.

^{*}Reference: Rothman. *Epidemiology: An Introduction*. New York, NY: Oxford Univ Press.