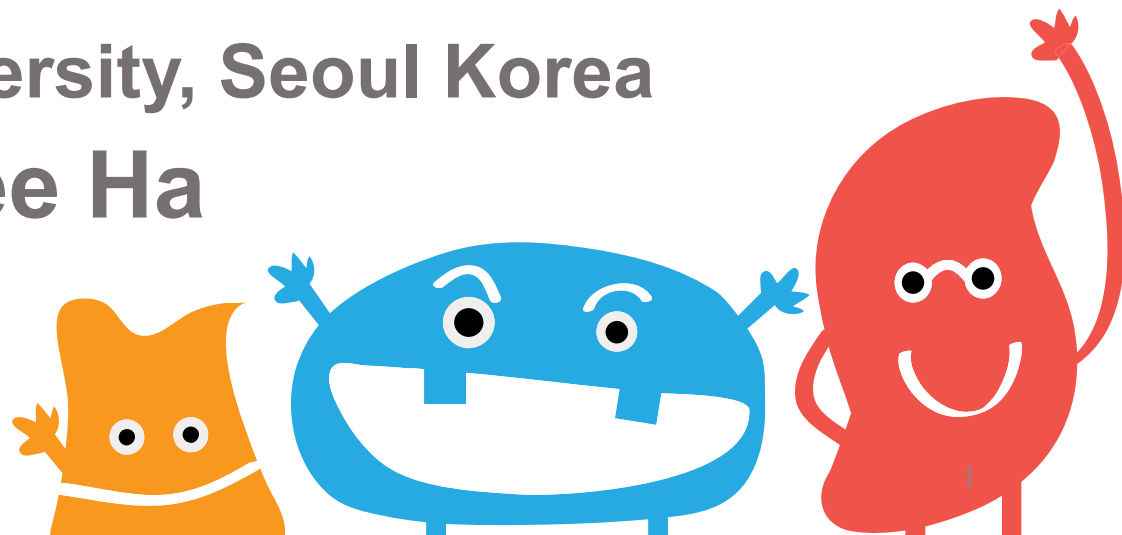
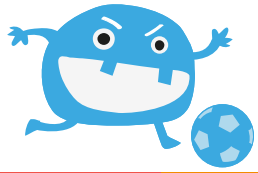


# Air pollution and pregnancy outcome and children's health

Ewha Womans University, Seoul Korea  
Eunhee Ha

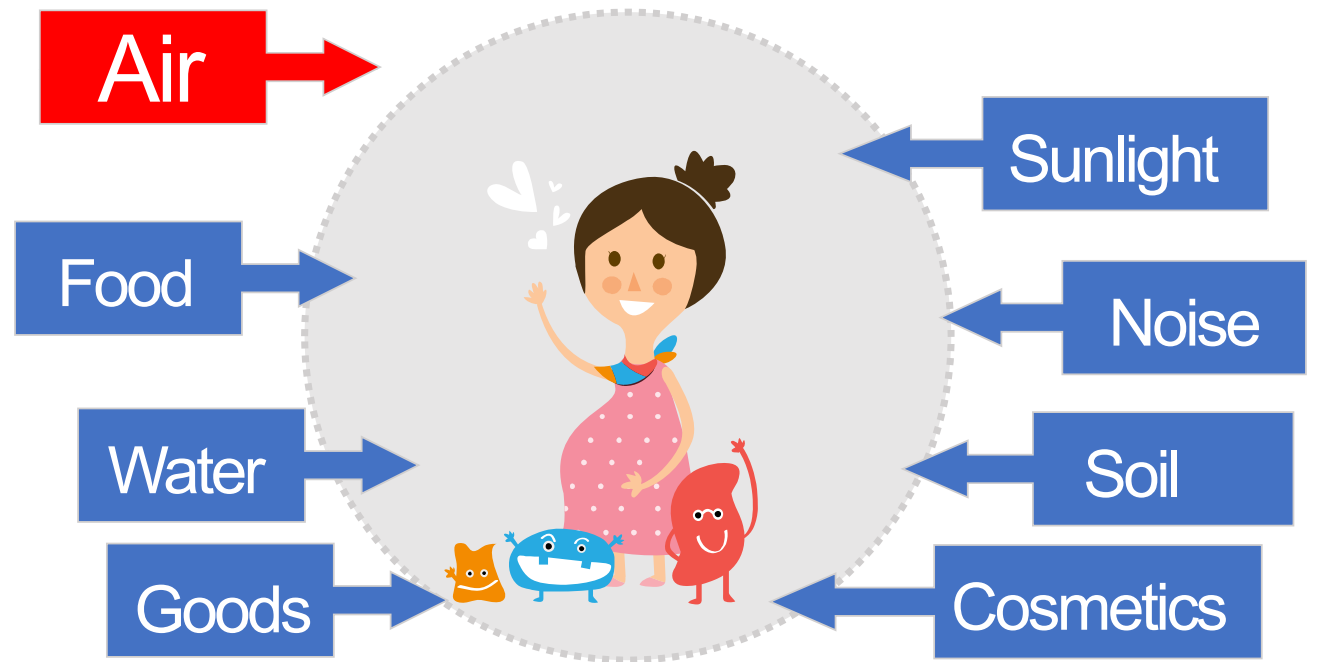




# Index

- I. Environmental pollutants
- II. Children and Environment
- III. Air pollution and health
- IV. Air pollution and Pregnancy outcome  
and Children's Health from MOCEH
- V. Conclusion

# I. Environmental pollutants

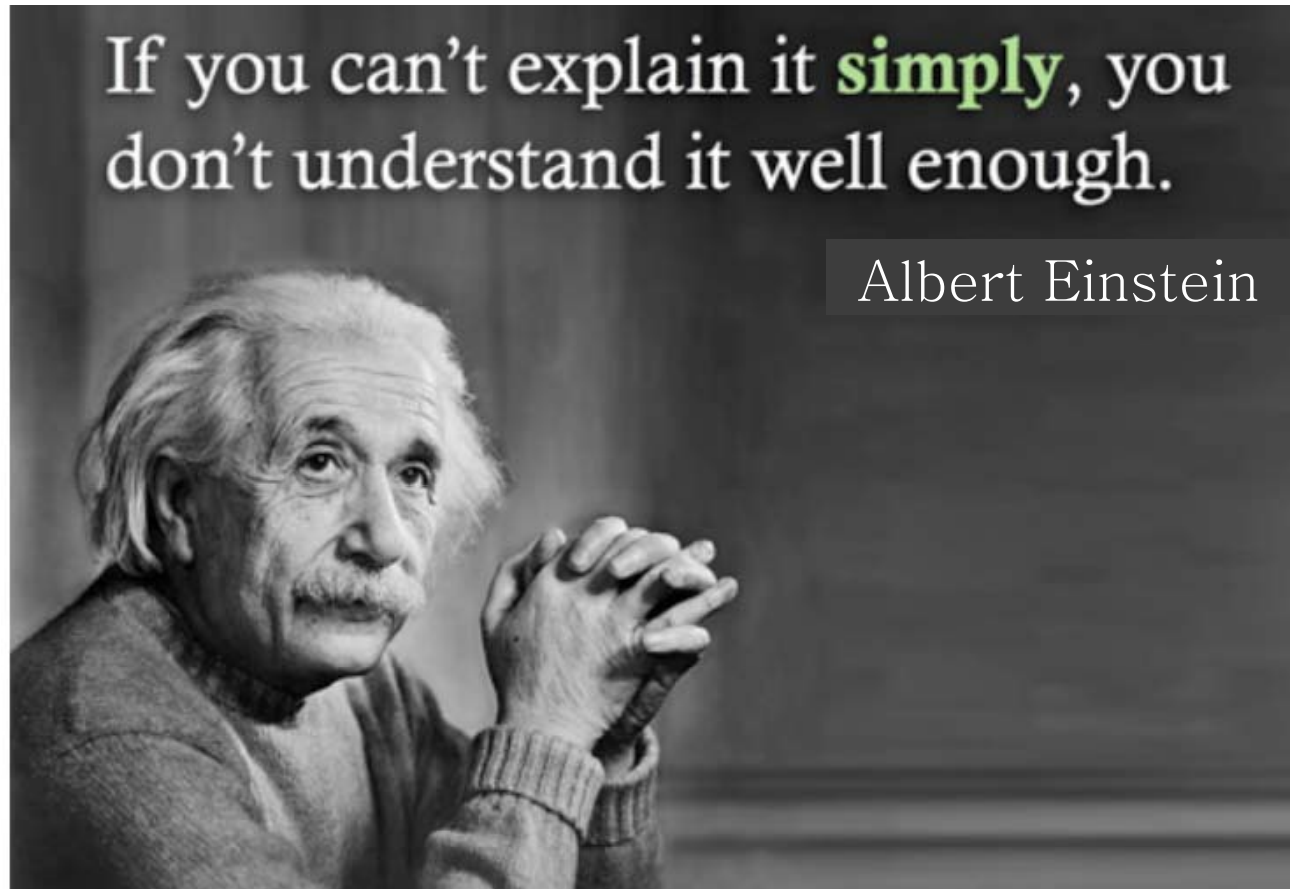




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# 1. what is the environment?

**"The environment is everything that is not me"** Albert Einstein



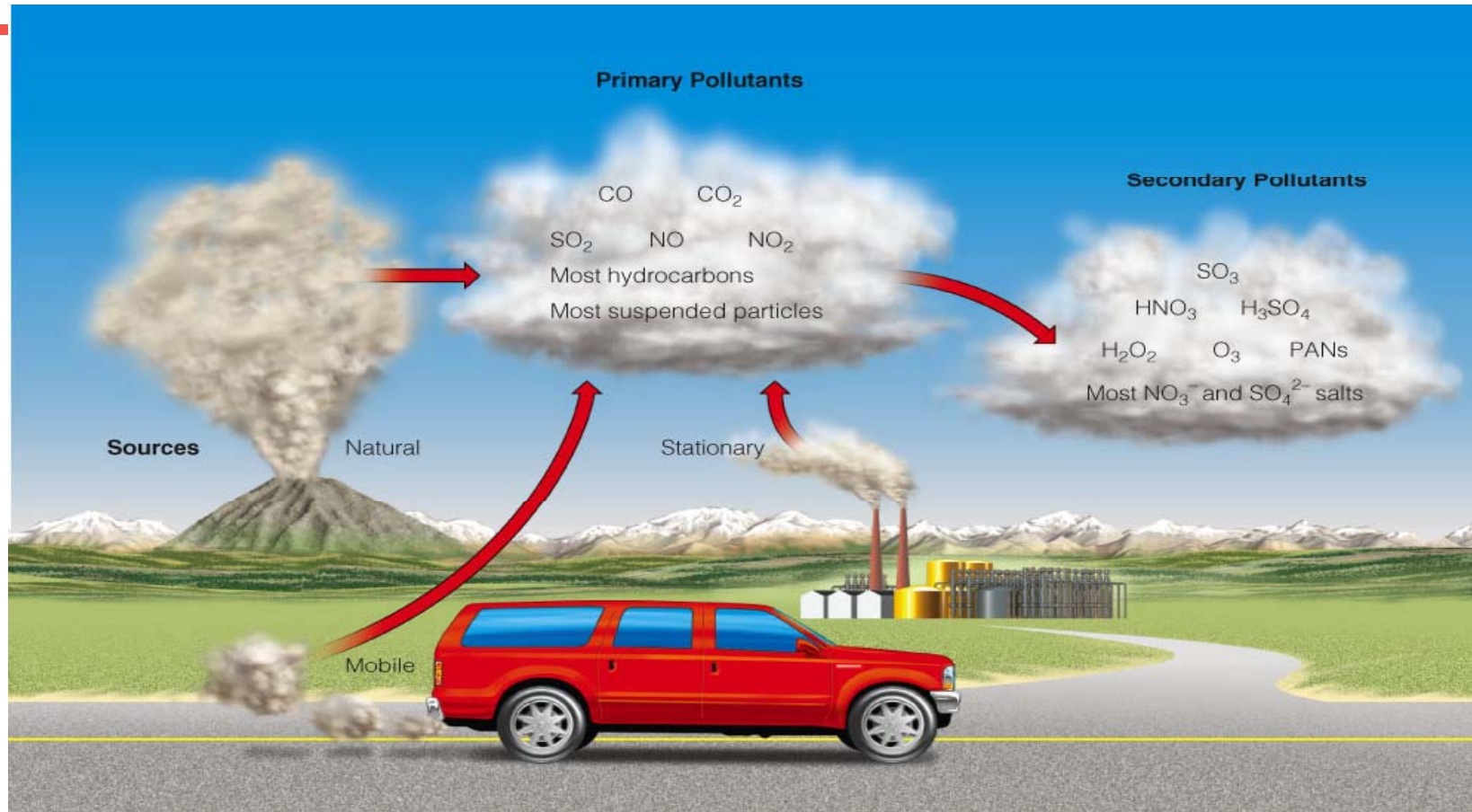
## 2. Environmental pollutant is..

- Focuses on exposure to **contaminants found in the environment**(specifically during critical periods of development)
- All the physical, chemical, biological and social factors that may affect the origin, growth, development and survival of an organism in a given setting.
- *Some examples include:*
  - **Air contaminants**
  - Specific synthetic chemicals
  - Some metals



**Still an emerging issue!**

### 3. Ambient(outdoor) air pollutant is...



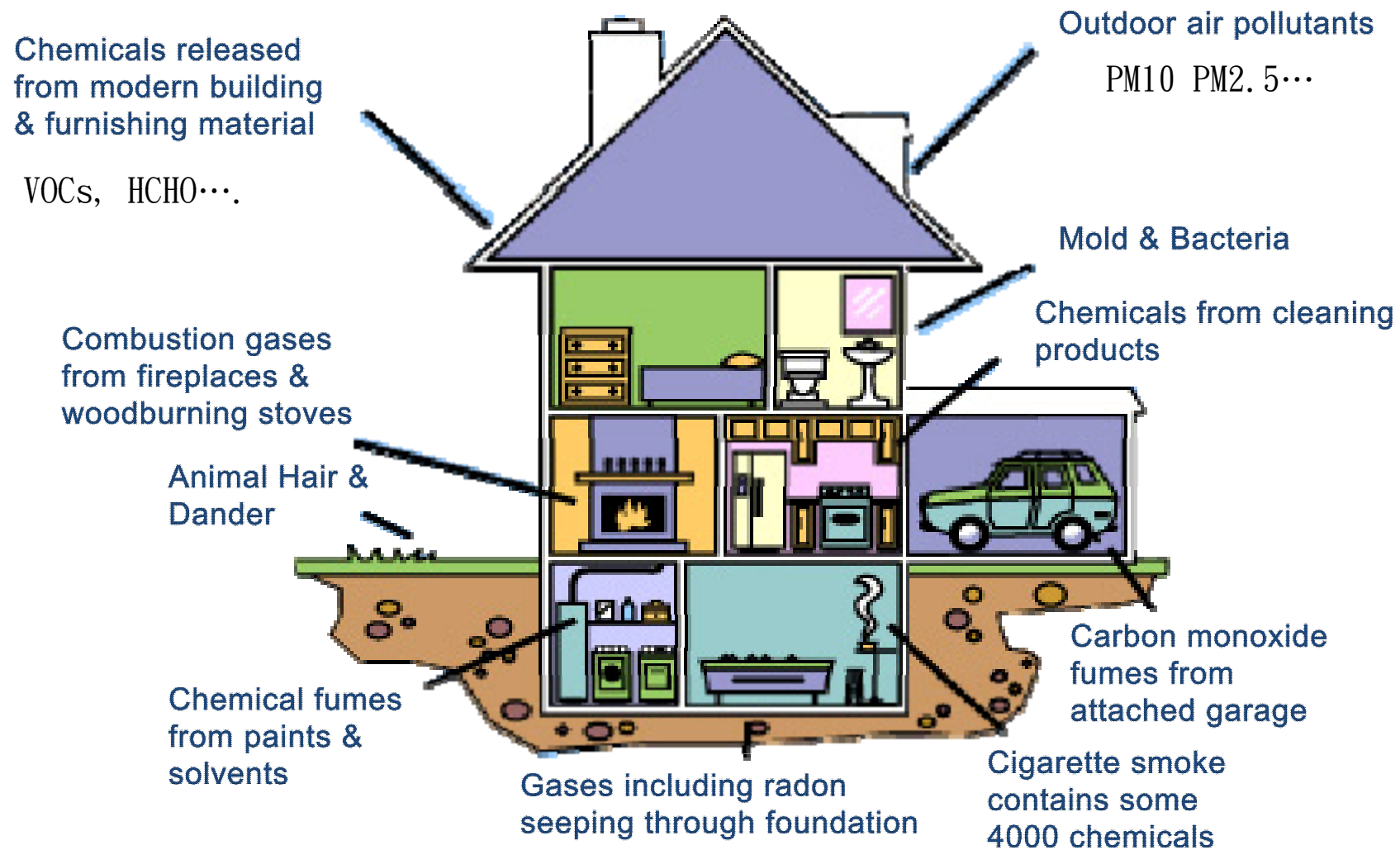
© 2007 Thomson Higher Education

Some primary air pollutants may react with one another or with other chemicals in the air to form secondary air pollutants.

- A 2013 assessment by WHO's International Agency for Research on Cancer (IARC) concluded that **outdoor air pollution is carcinogenic to humans**, with the particulate matter component of air pollution most closely associated with increased cancer incidence, especially cancer of the lung.



## 4. Indoor air pollutant's source is...



Source: Reprinted from US Environmental protection Agency. Air Pollution Control Orientation Course, Indoor Air Pollution.  
Available at: <http://www.epa.gov/air/oaqps/eog/course422/ap4.html>. Accessed March 22,2010



## 5. Key air pollutants

# 1) Particulate Matter

- Particulate matter (PM) is an air pollution term for a mixture of solid particles and liquid droplets found in the air.
- Of particular concern is a class of particles known as fine particulate matter or PM<sub>2.5</sub> that gets deep into the lung
- Particles that are small enough to be inhaled have the potential to cause health effects.
- Children frequently breathe through their mouths, thus bypassing the nasal filtration mechanism

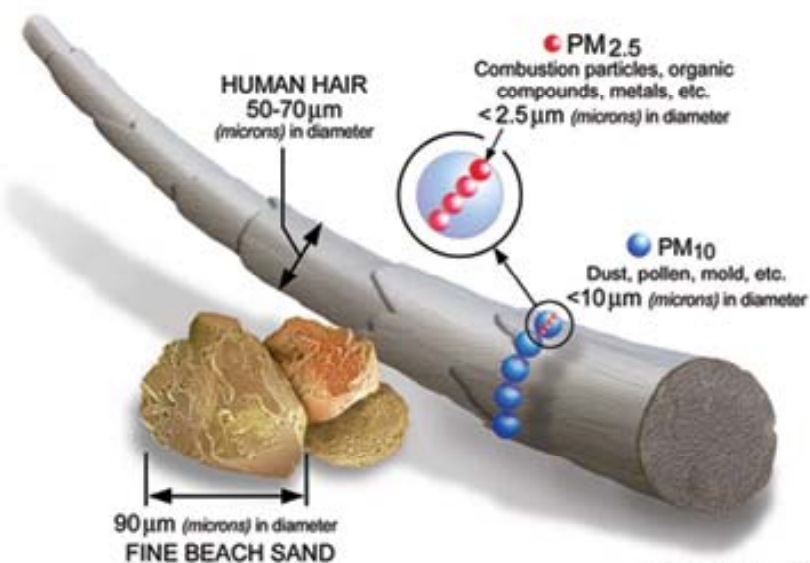
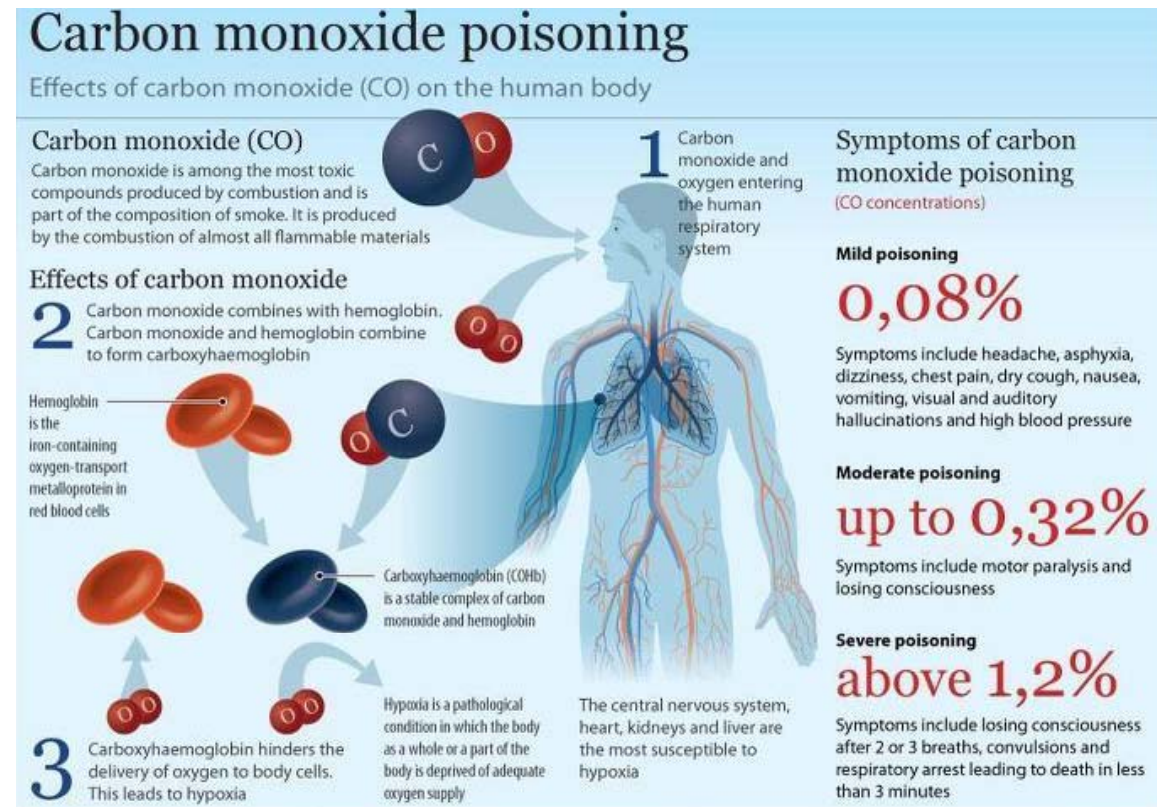


Image courtesy of the U.S. EPA

## 2) Carbon Monoxide

- Incomplete combustion of biomass fuel/ poorly maintained or poorly ventilated cooking or heating appliances
- The binding of CO to hemoglobin: 245 times as strong as that for oxygen
- Brain tissue metabolism is maintained until carboxyhemoglobin approaches 20%.



### Guidelines

100mg/m<sup>3</sup> for 15 mins ; 35mg/m<sup>3</sup> for 1 hr ; 10 mg/m<sup>3</sup> for 8hrs ; 7mg/m<sup>3</sup> for 24 hrs

### 3) Polycyclic Aromatic Hydrocarbons

- A large group of organic compounds with two or more fused aromatic (benzene) rings
- Particle-bound PAHs (multiringed) are considered to be very hazardous to human health

#### Benzo[a]pyrene(B[a]P)

- ✓ As a marker for carcinogenic PAHs as the contribution of B[a]P to the total carcinogenic potential is high
- ✓ Buring of fossil fuels and biofuels for cooking in unvented stove (gas< kerosene<coal<wood<dung cake/wood mixture<dung cake)
- ✓ Human effects  
Their mutagenicity and carcinogenicity by the formation of DNA adducts ► Immunotoxic, decrease fertility, embryotoxicity, reduction in birth weight, increased the risk of asthma symptoms by the age of 1-2 years

## 4) Nitrogen Dioxide and Sulphur Dioxide

---

- Indoor sources of NO<sub>2</sub> Tobacco smoke, burning appliances such as stoves, ovens, heaters, fireplace
- NO<sub>2</sub> is a free radical and it has the potential to deplete tissue antioxidant defenses and cause injury and inflammation
- Indoor level of NO<sub>2</sub> are from both indoor and outdoor sources
- The association of lower respiratory symptoms with indoor NO<sub>2</sub> in children from the Six Cities study and respiratory illness from meta-analysis in 1990s
- Similar respiratory health effect observed with increasing SO<sub>2</sub>

## 5) VOCs AND Formaldehyde

- **Volatile organic compounds (VOCs)** are organic chemicals that have a high vapor pressure at ordinary room temperature
  - cleaning, painting, or hobby supplies that contain them
  - dry clean your clothes with home dry-cleaning products; at a professional dry-cleaners;
  - graphics and crafts materials such as glues, permanent markers, and photographic solutions
- **Formaldehyde** is a naturally-occurring organic compound with the formula  $\text{CH}_2\text{O}$ .
  - Exposed at home if you use unvented.
  - by using household products such as construction materials, latex paints, fingernail polish, cosmetics, disinfectants, glues, lacquers, manufactured pressed wood products, fiberglass, new carpets, permanent press fabrics, paper products, and some cleaners.

## II. Children and Environment :Why children are important?

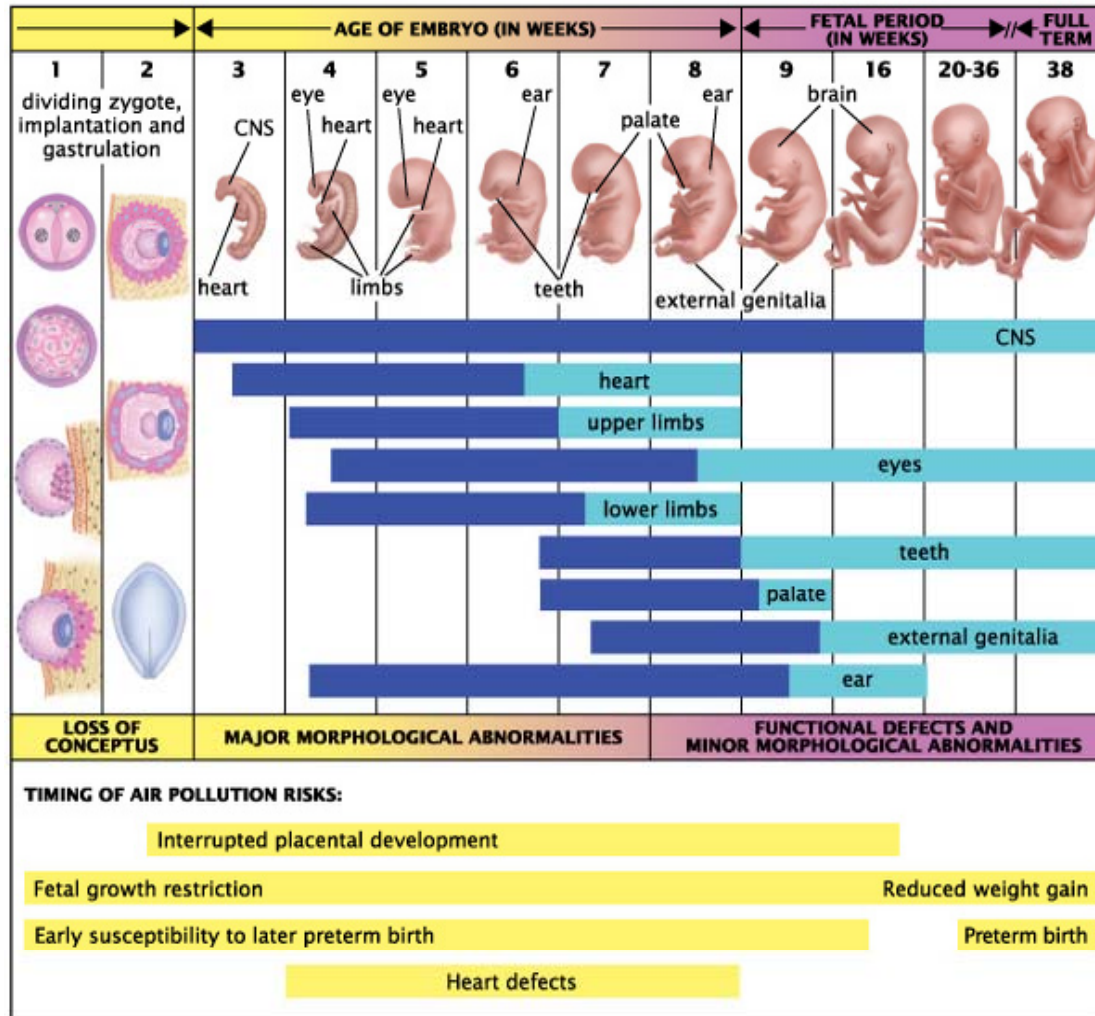


# 1. Children are not “little adults”

- Exposures to toxic chemicals in early life are important causes of disease and dysfunction in children and also in adults.
- Diseases caused by chemicals can successfully be prevented, thus saving lives, enhancing the quality of life, reducing health care and education costs, and in creasing productivity.



## 2. Critical Window of Susceptibility



Note: Blue bars indicate time periods when major morphological abnormalities can occur, while light blue bars correspond to periods at risk for minor abnormalities and functional defects.

Figure 1. Fetal development and timing of air pollution risks.

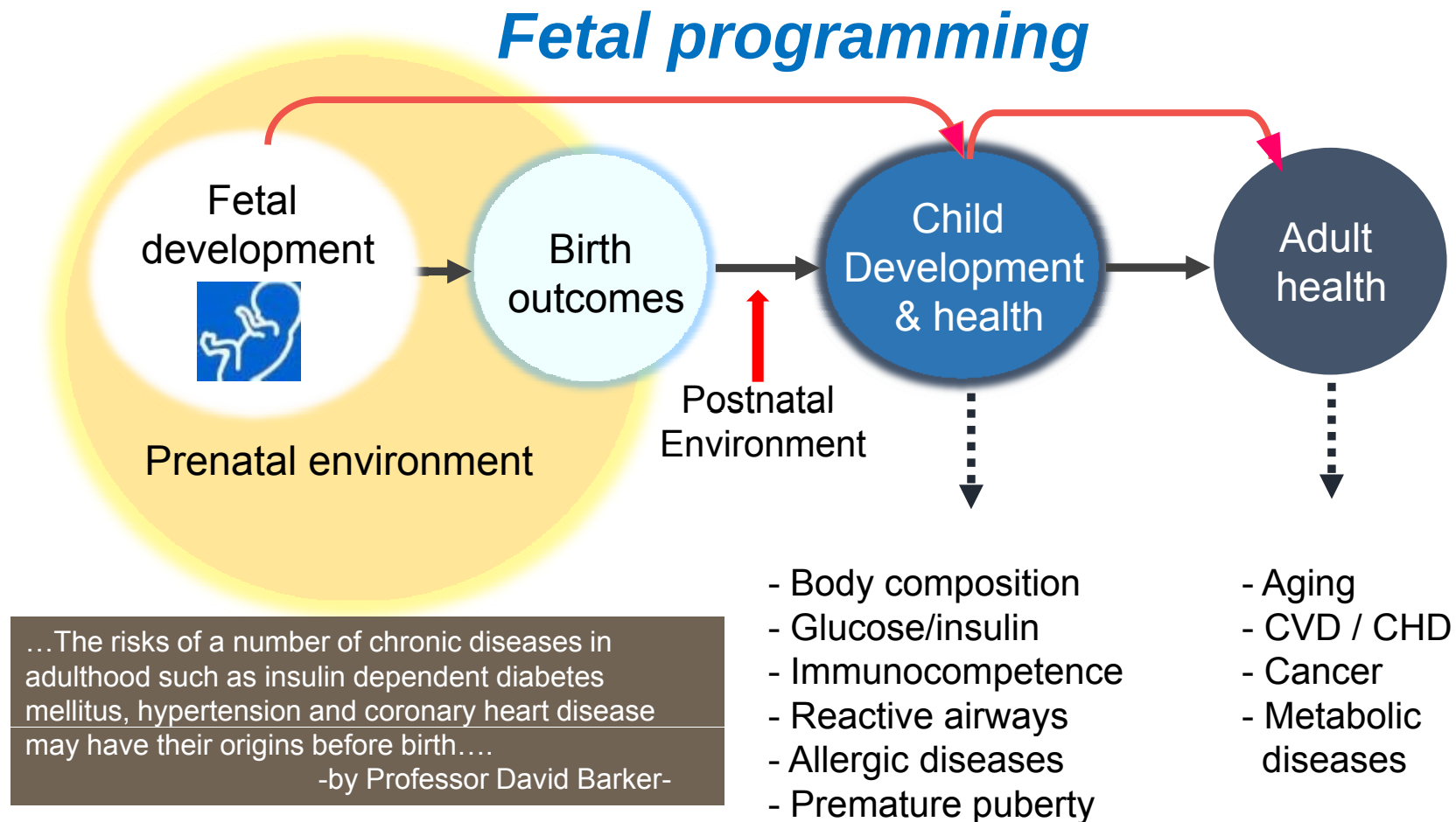
| Stage:<br>Age:       | Newborn<br>0-2 mos                                                                  | Infant/Toddler<br>2 mos-2 yrs | Young Child<br>2-6 yrs                                                              | School-Age Child<br>6-12 yrs | Adolescent<br>12-18 yrs                                                             |
|----------------------|-------------------------------------------------------------------------------------|-------------------------------|-------------------------------------------------------------------------------------|------------------------------|-------------------------------------------------------------------------------------|
|                      |  |                               |  |                              |  |
| Lung development:    | Alveolar development                                                                |                               |                                                                                     |                              |                                                                                     |
|                      | High respiratory rate                                                               |                               | Increasing lung volume                                                              |                              |                                                                                     |
| Air pollution risks: | Respiratory death                                                                   |                               | Chronic cough and bronchitis                                                        |                              |                                                                                     |
|                      |                                                                                     |                               | Reduced lung function                                                               |                              |                                                                                     |
|                      |                                                                                     |                               | Wheezing and asthma attacks                                                         |                              |                                                                                     |
|                      | Respiratory symptoms and illnesses*                                                 |                               | Respiratory-related school absences                                                 |                              |                                                                                     |

\*Air pollution exposure has also been more recently linked to respiratory symptoms and illnesses in early life including cough, bronchitis, wheeze and ear infections

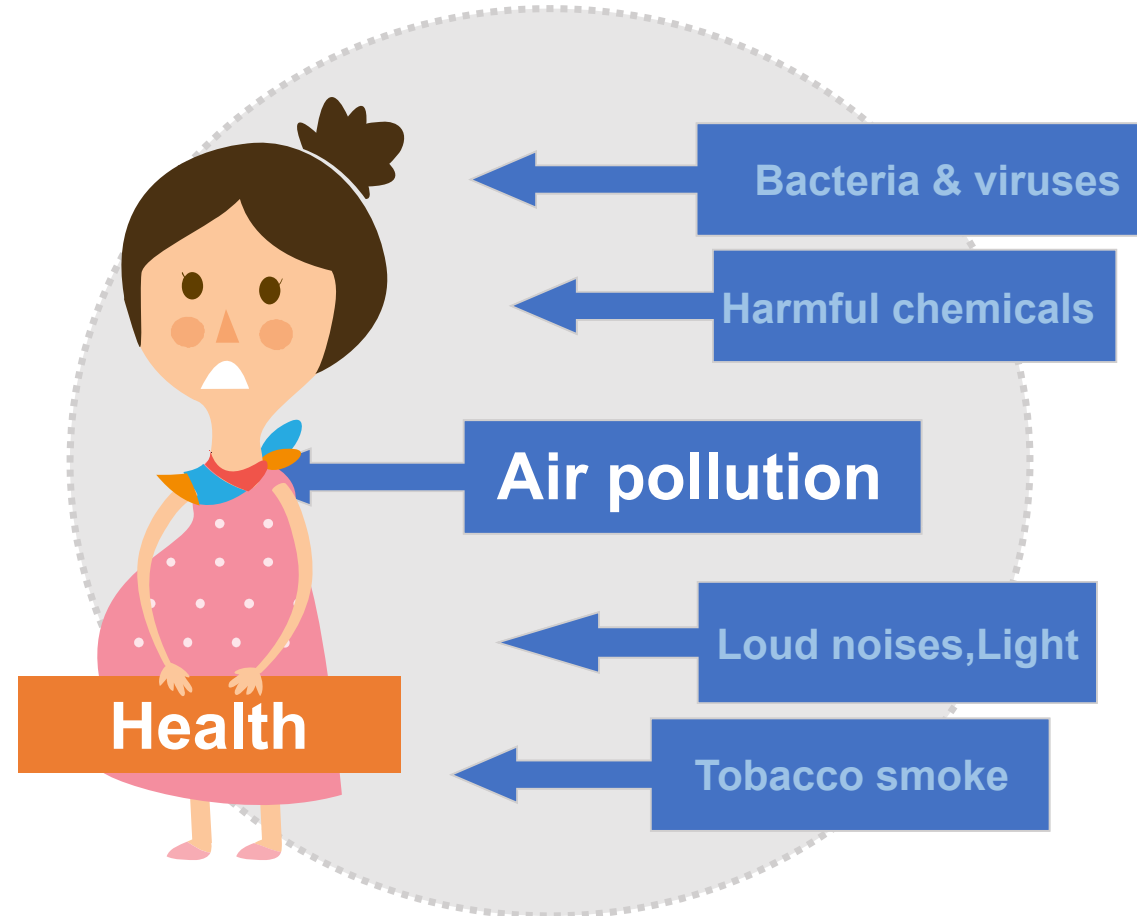
Figure 2. Air pollution effects on the developing respiratory system.

UCLA Institute of the Environment and Sustainability,  
<http://www.environment.ucla.edu/reportcard/article1700.html>

# 3. Developmental Origins of Health & Disease

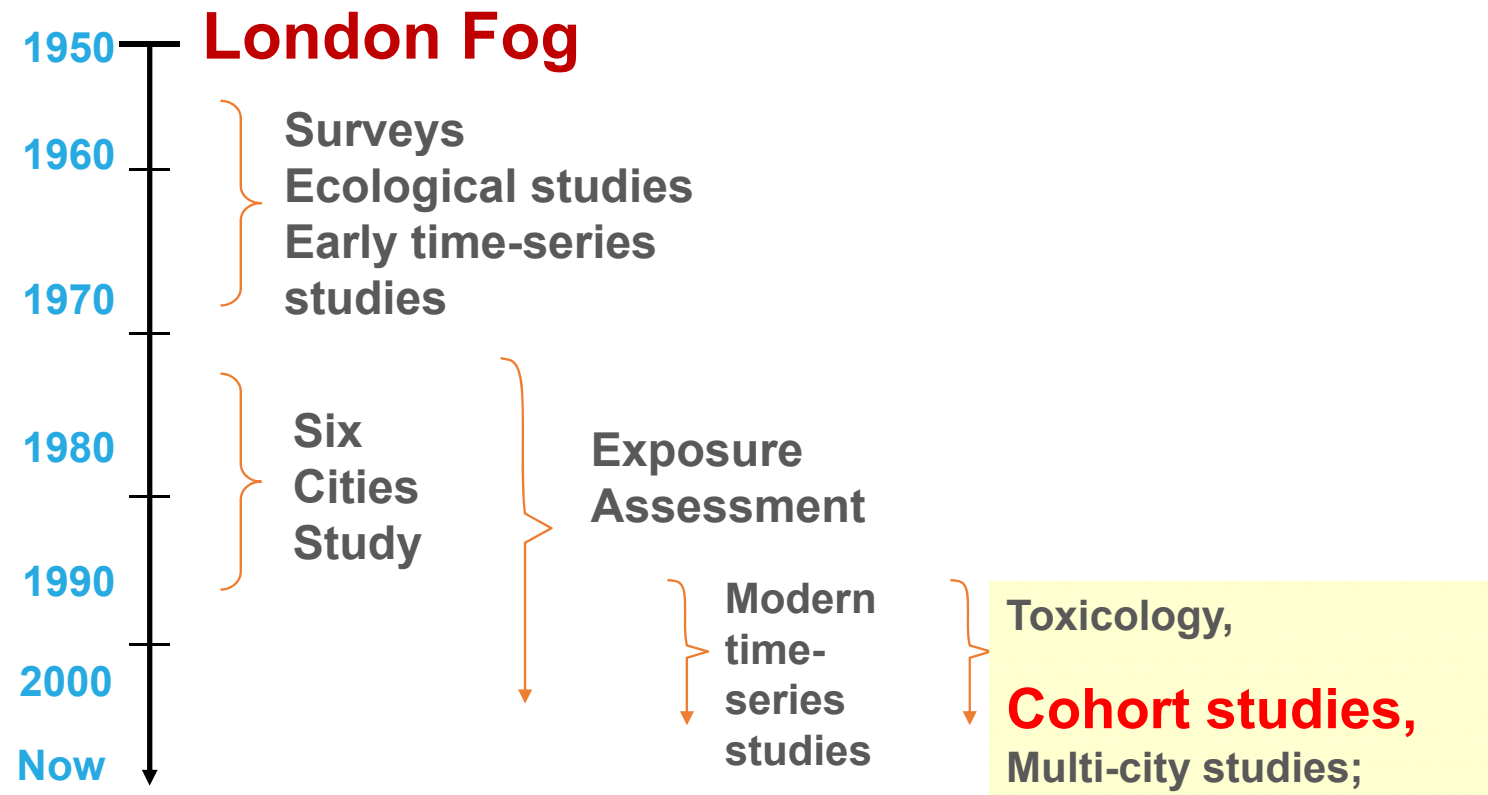


### III. Air pollution and health

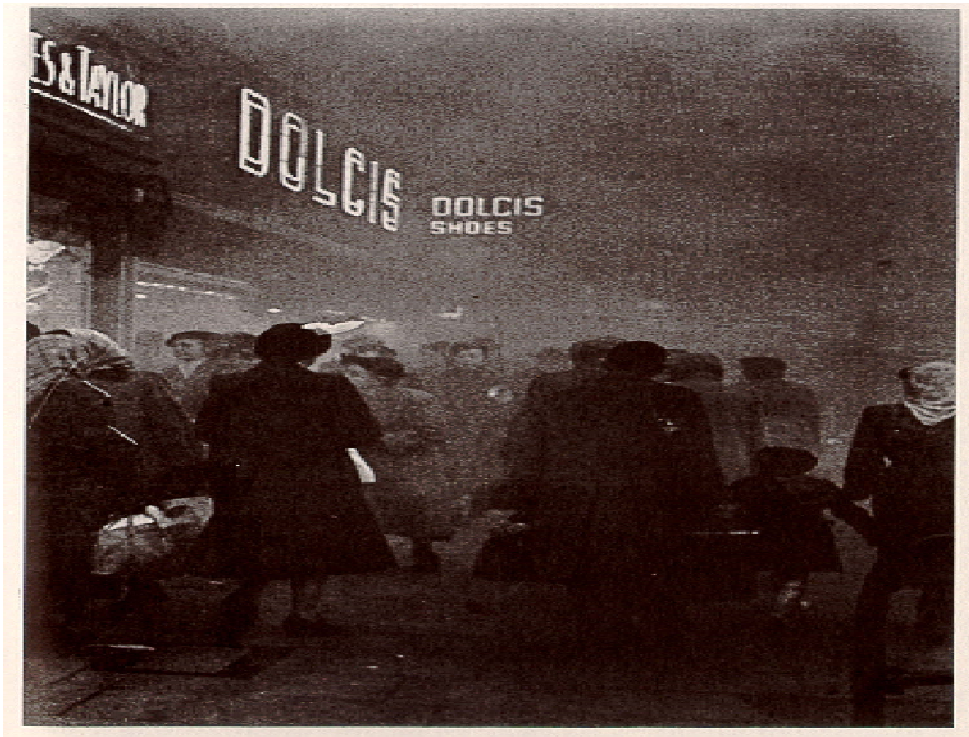


# 1. Historical episode and Study

- 50+ Years of Air Pollution Research



# 1) London, December 1952

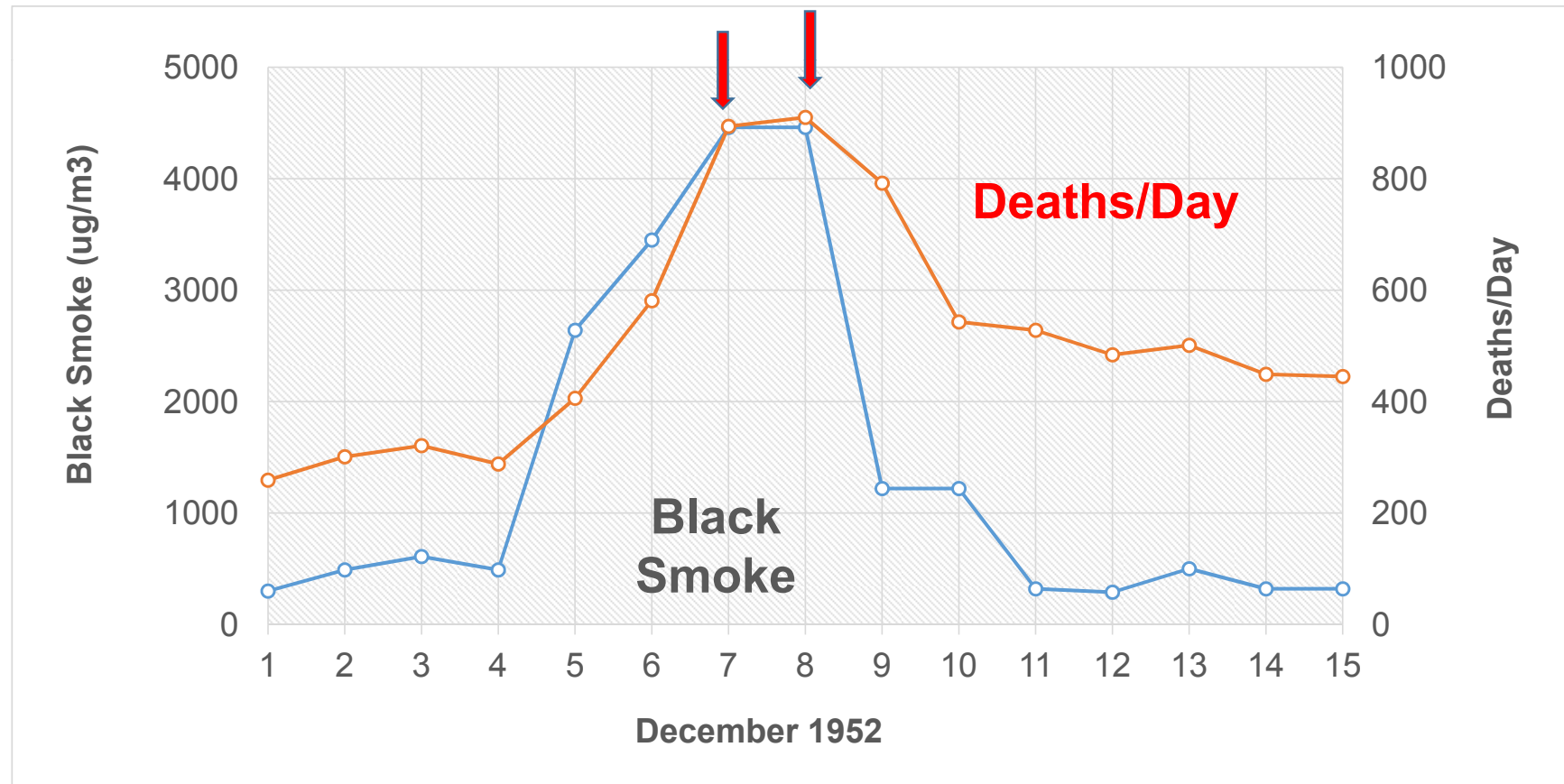


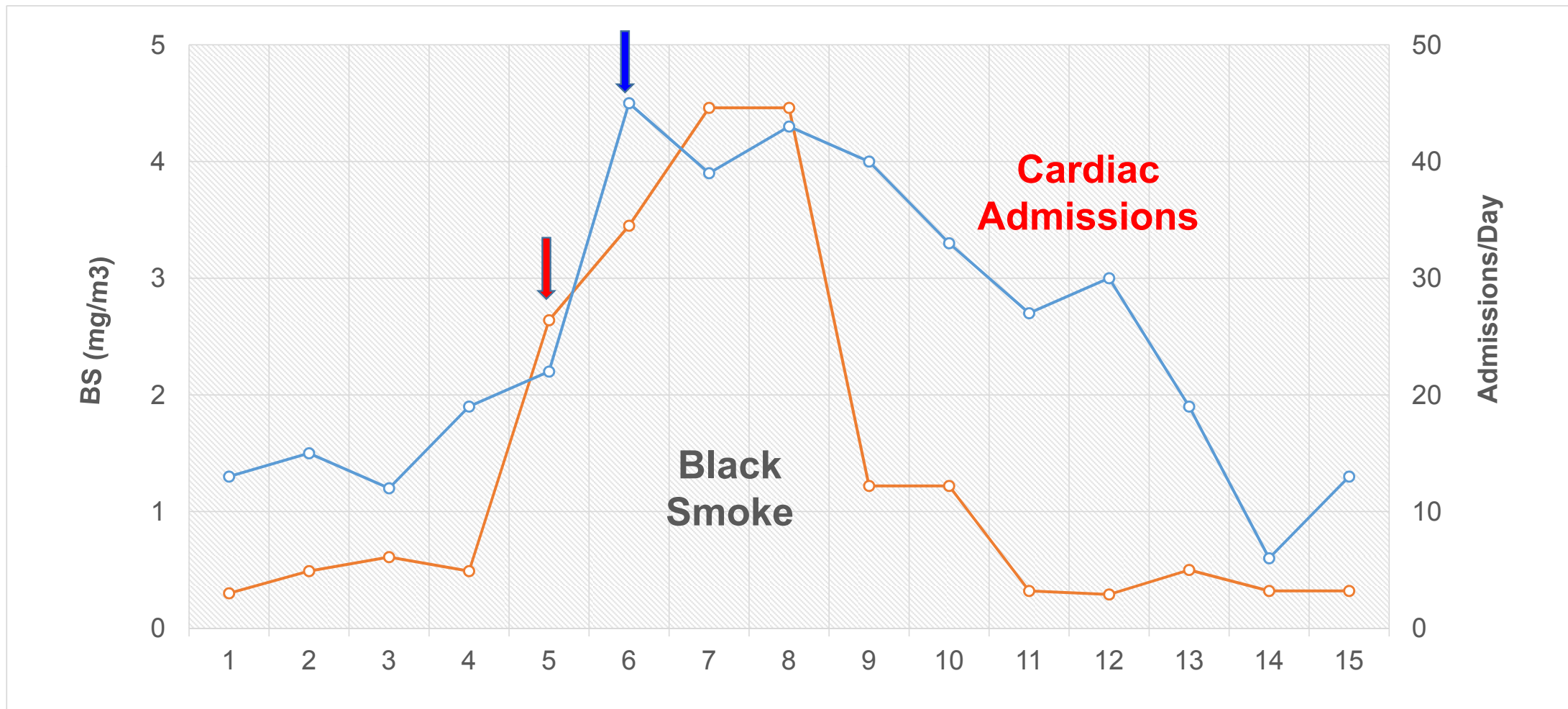
London December 1952,  
Picture Taken at Noon

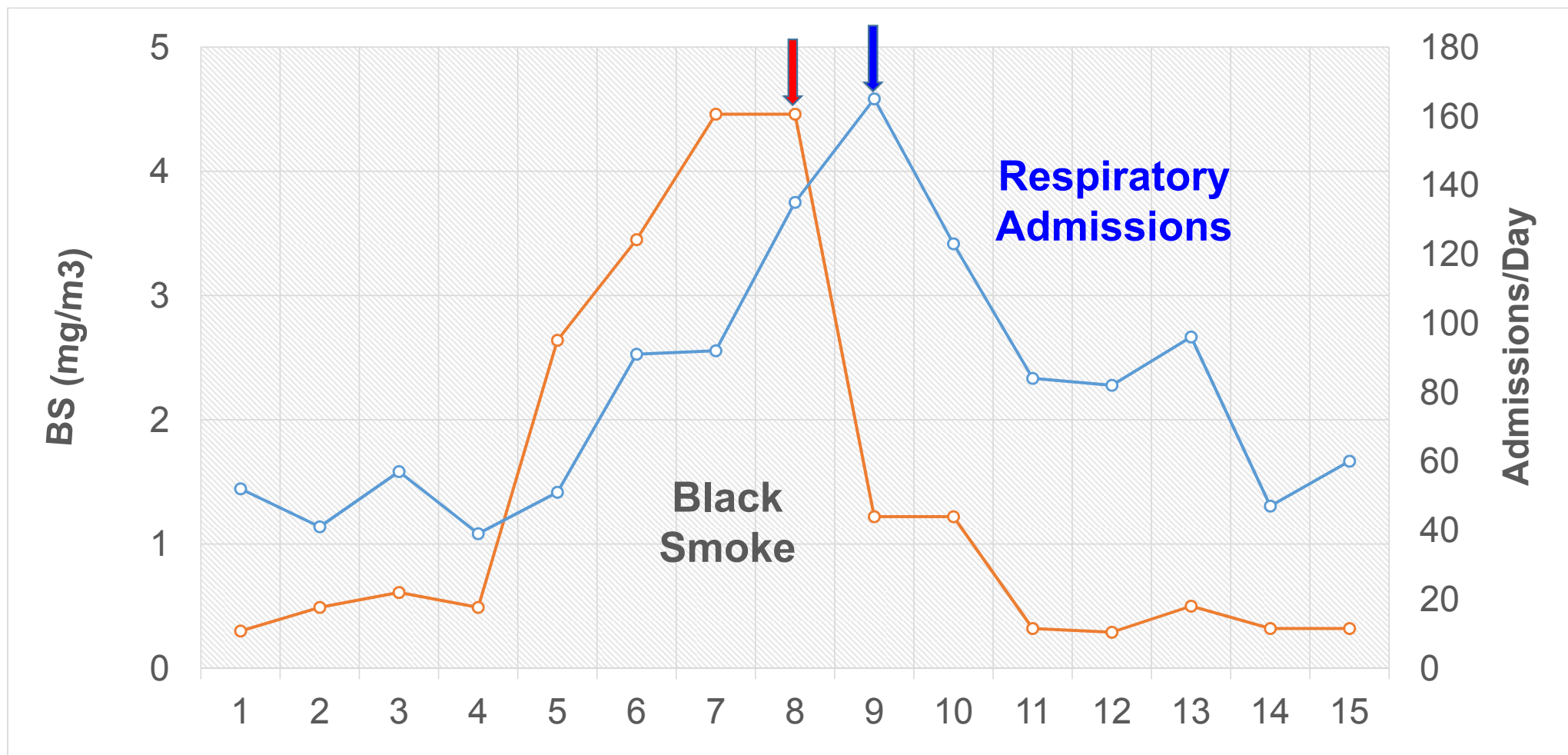


Donora, Pennsylvania Noon,  
October 28, 1948

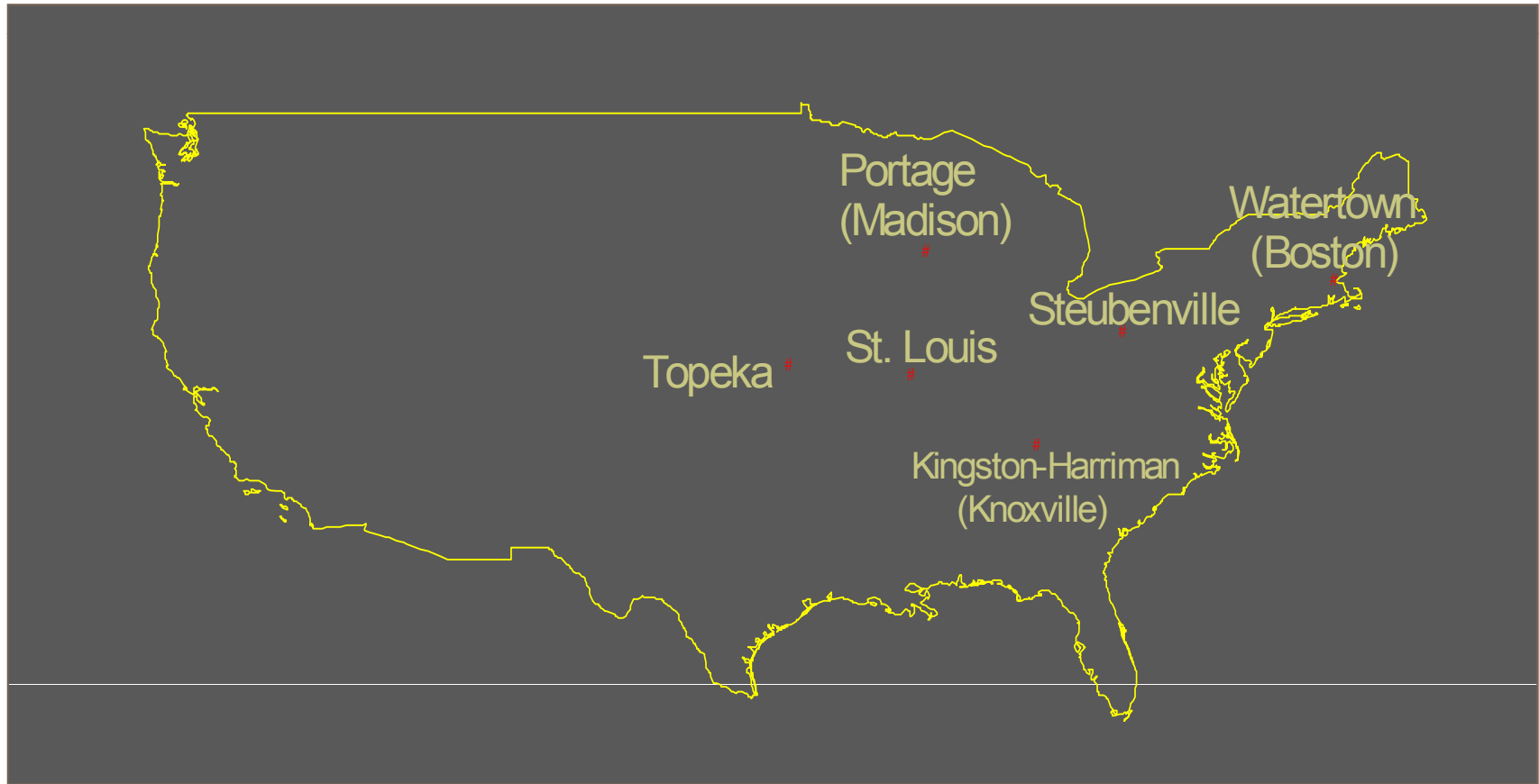
- 1952 London Fog Episode







## 2) Harvard 6 Cities Study



▲ Original  
Six City  
monitors

● EPA  
monitors

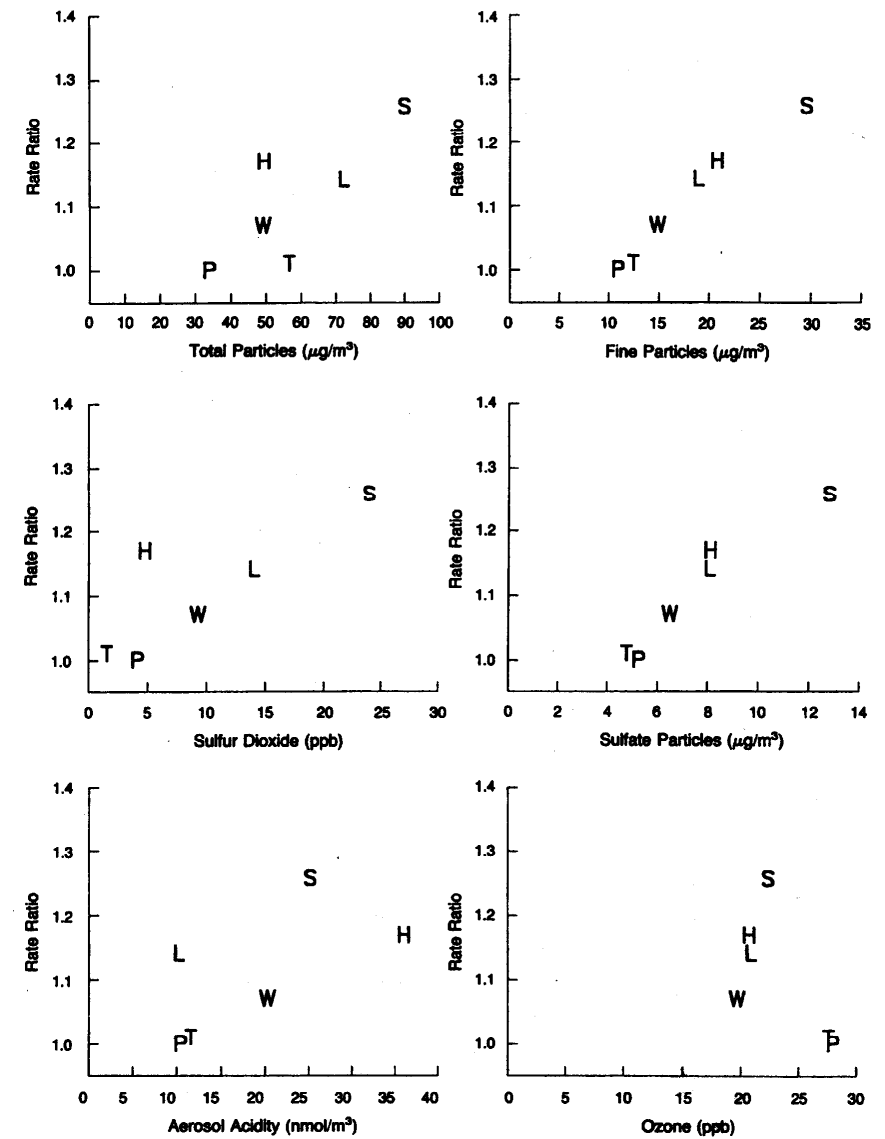
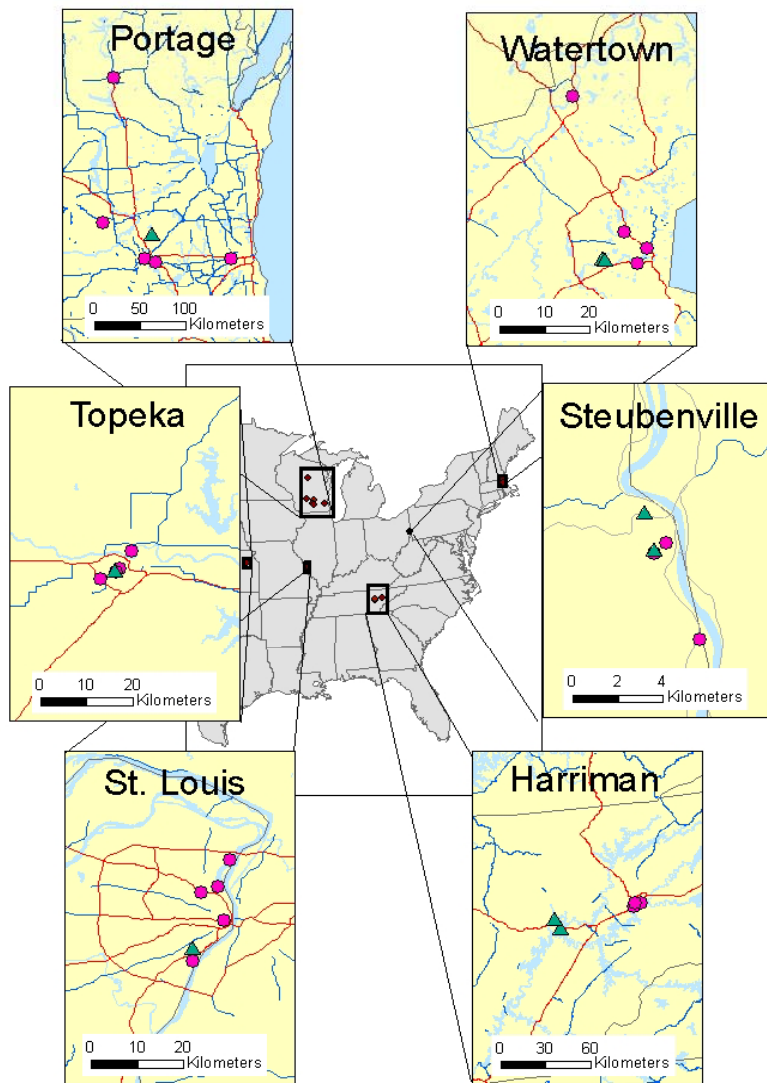
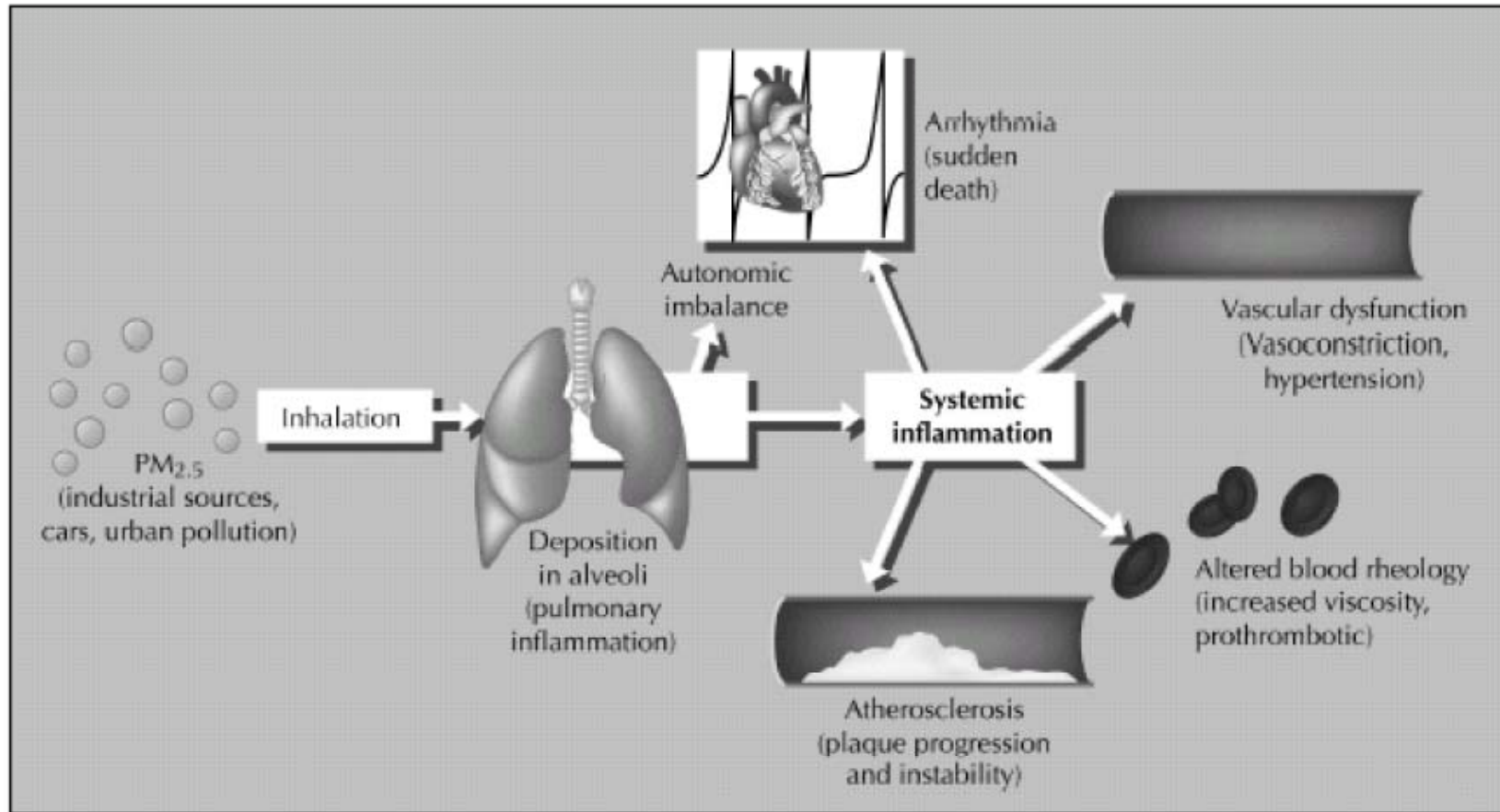
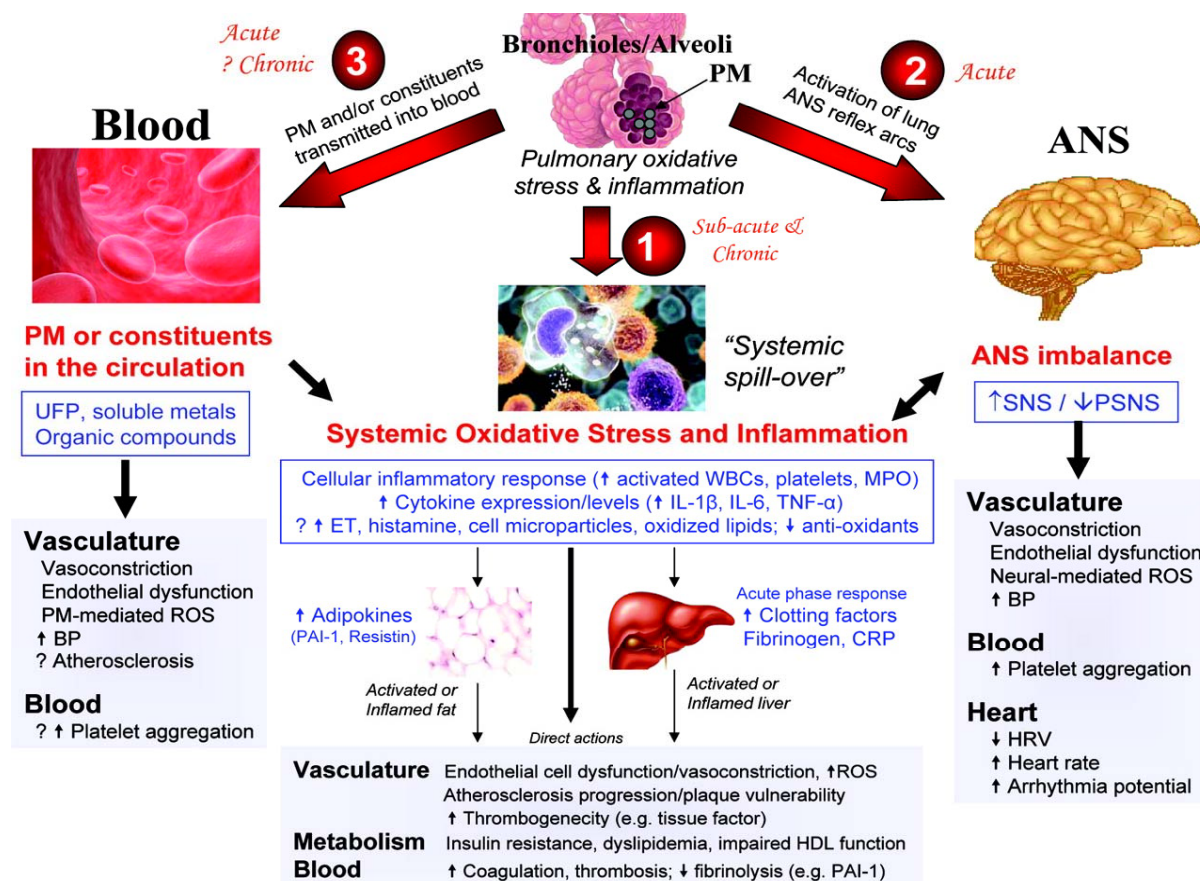


Figure 3. Estimated Adjusted Mortality-Rate Ratios and Pollution Levels in the Six Cities. Mean values are shown for the measures of air pollution. P denotes Portage, Wisconsin; T Topeka, Kansas; W Watertown, Massachusetts; L St. Louis; H Harriman, Tennessee; and S Steubenville, Ohio.

## 2. Mechanisms of pathogenesis



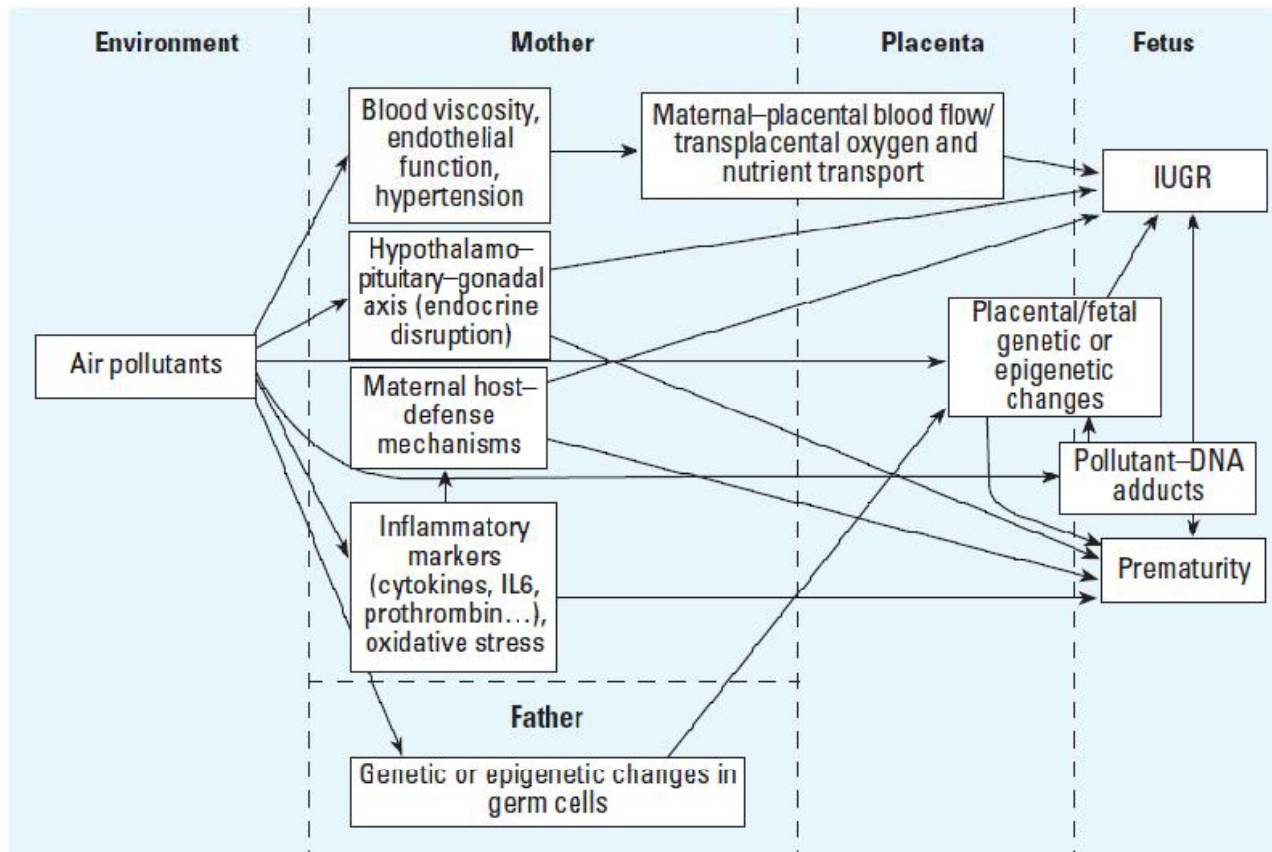
Biological pathways linking PM exposure with CVDs. The 3 generalized intermediary pathways and the subsequent specific biological responses that could be capable of instigating cardiovascular events are shown.



Brook R D et al. *Circulation*. 2010;121:2331-2378

# Possible biological mechanisms by which air pollutants may affect birth outcomes

Source : Slama et al. *Environmental Health Perspective*, 2008, 116(6):7953



- Air pollutants effect many biological reactions on mother, placenta and fetus.
- Fine particles can induce oxidative stress and inflammation.
- That contributes to onset of preterm labor and other adverse perinatal outcomes.



### 3. Air pollutant and Health effect

# 1) SUMMARY MAJOR OUTDOOR POLLUTANTS

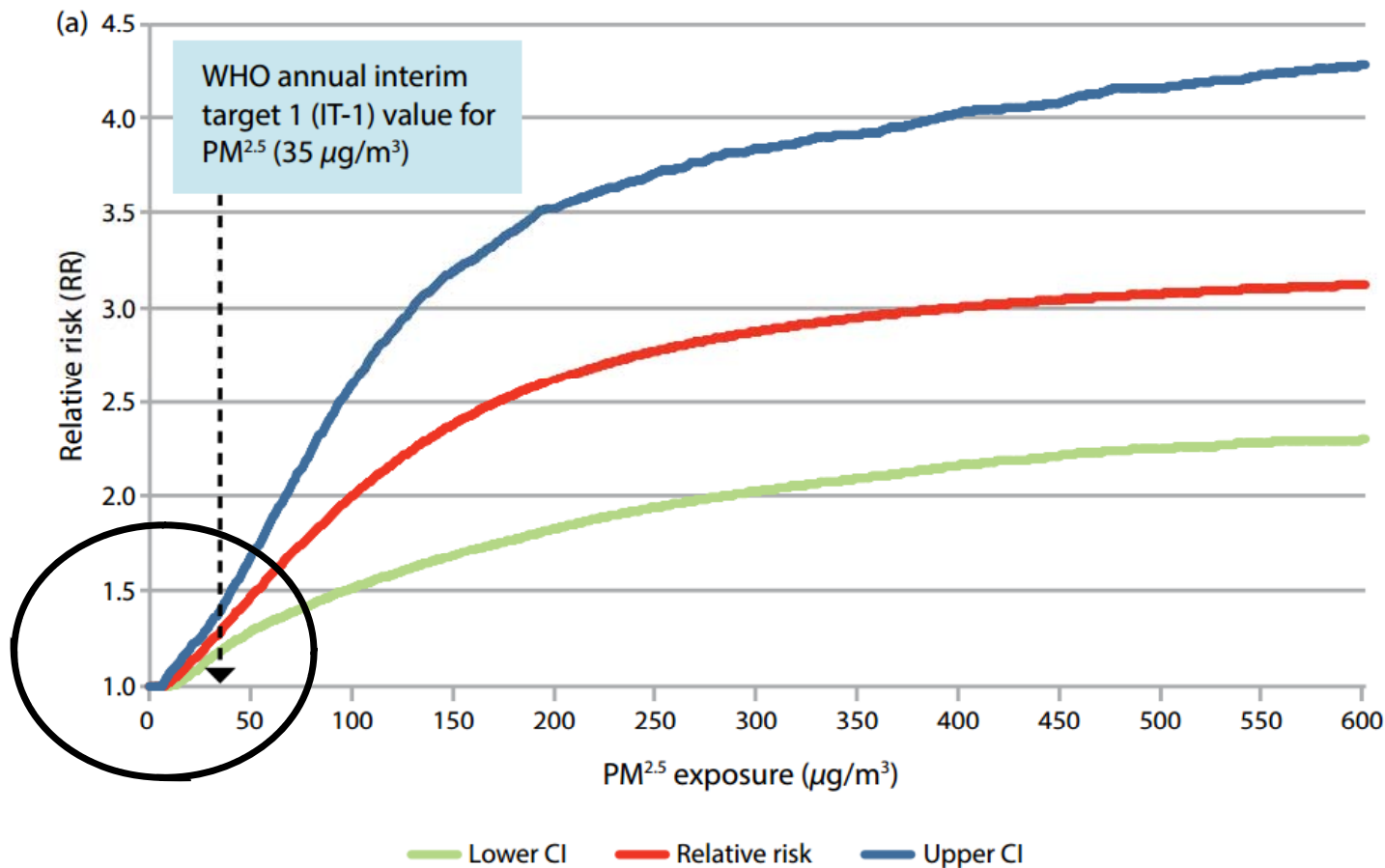
| Pollutant                 | Sources                                                                                                                  | Health Effects                                                                                                                                                            |
|---------------------------|--------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>Particulate Matter</b> | Automobile, bus and truck exhaust, fuel burning (wood stoves, fireplaces), industry, construction.                       | ↑ infant respiratory mortality<br>↓ lung function<br>↓ lung growth<br>↑ symptoms in asthmatics                                                                            |
| <b>Ozone</b>              | Produced when nitrogen oxides (vehicle emissions) and volatile organic compounds (VOC) chemically react under sunlight.  | ↓ lung growth<br>↑ asthma exacerbations<br>↑ all respiratory hospitalization<br>↑ asthma hospitalization<br>↑ asthma ED visit<br>↑ school absence for respiratory illness |
| <b>Nitrogen dioxide</b>   | Results from high temperature fuel combustion and atmospheric reactions.                                                 | ↑ symptoms in asthmatics<br>↓ lung growth                                                                                                                                 |
| <b>Carbon monoxide</b>    | Formed when carbon-containing fuel is not burned completely, emitted by motor vehicles more than any other source.       | ↑ asthma hospitalization<br>↑ clinic visits for lower respiratory tract disease<br>headache                                                                               |
| <b>Sulfur dioxide</b>     | <b>Industrial sites such as smelters, paper mills, power plants and steel manufacturing plants are the main sources.</b> | ↑ asthma hospitalization<br>↑ clinic visits for lower respiratory tract disease                                                                                           |

[http://www.who.int/ceh/capacity/Outdoor\\_air\\_pollution.pdf](http://www.who.int/ceh/capacity/Outdoor_air_pollution.pdf)

## 2) Studies for the relationship between PM and low birth weight

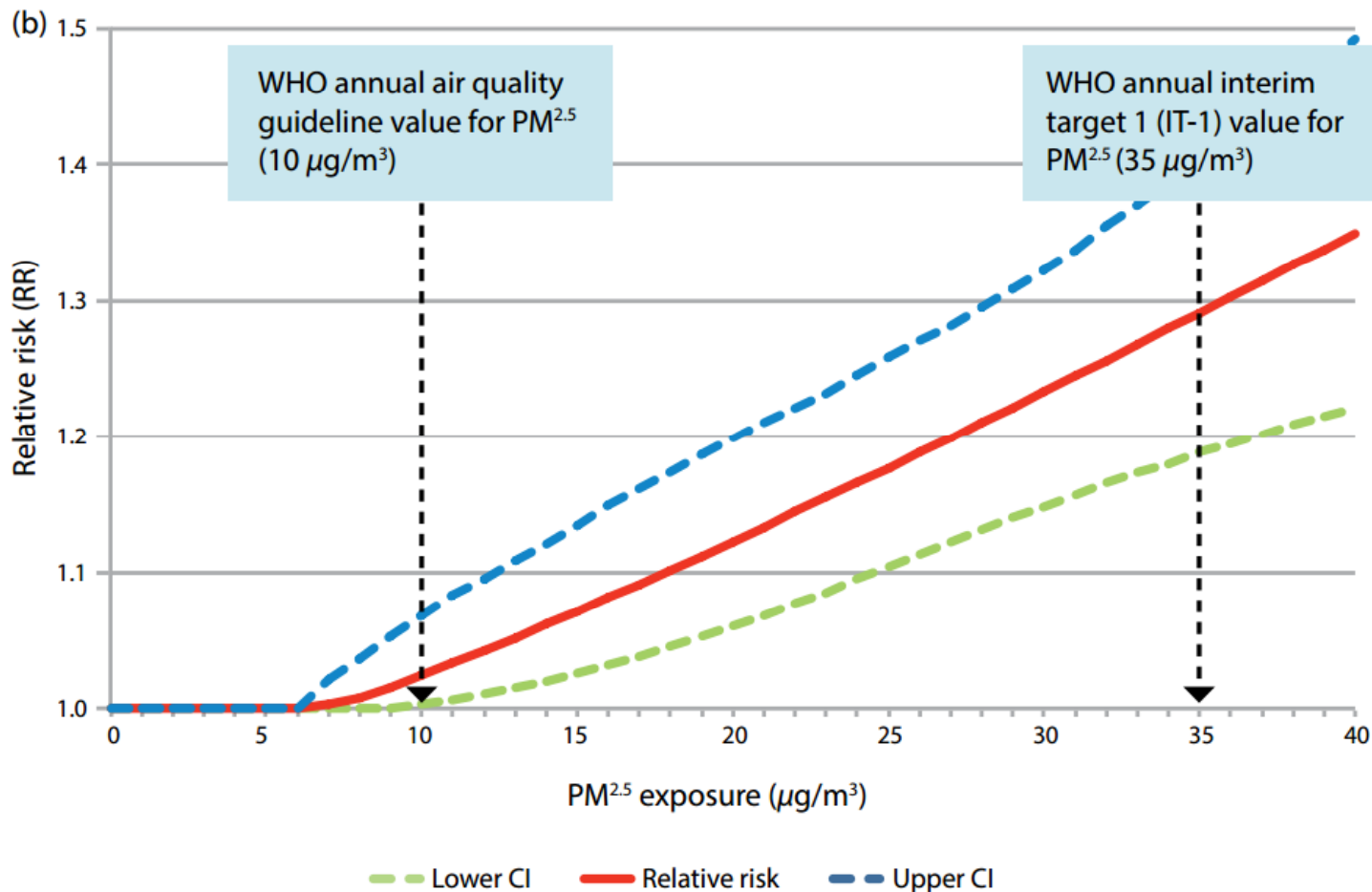
| Author                | Substance  | Published Year | Exposure period (trimester) | Effect size§                                                                                                                                                                                   |
|-----------------------|------------|----------------|-----------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Vinikoor-Imler et al. | PM2.5      | 2014           | Entire                      | Null associations                                                                                                                                                                              |
| Wilhelm et al.        | PM2.5      | 2012           | Entire                      | From gasoline IQR OR 1.07 (0.97,1.18)<br>From diesel IQR OR 1.06 (0.99,1.14)                                                                                                                   |
| Madsen et al.         | PM2.5/PM10 | 2010           | Entire                      | Null associations                                                                                                                                                                              |
| Seo et al.            | PM10       | 2010           | Entire                      | Pusan city OR 1.24 (1.02–1.52)*<br>Daegu city OR 1.19 (1.04–1.37)*                                                                                                                             |
| Morello-Frosch et al. | PM2.5/PM10 | 2010           | Entire                      | An increase of 10 $\mu\text{g}/\text{m}^3$ of PM2.5 lowered 12.8g (11.3g–14.3g) of birth weight<br>An increase of 10 $\mu\text{g}/\text{m}^3$ of PM10 lowered 7.7g (6.6g–7.9g) of birth weight |
| Xu et al.             | PM10       | 2010           | First<br>Second             | IQR AOR 1.13 (1.02, 1.25)*<br>IQR AOR 1.10 (1.00–1.22)*                                                                                                                                        |
| Bell et al.           | PM2.5/PM10 | 2007           | Entire                      | PM2.5 IQR OR 1.05 (1.02–1.09)*<br>PM10 IQR OR 1.03 (0.99–1.06)                                                                                                                                 |
| Slama et al.          | PM2.5      | 2007           | Entire                      | Lowest vs highest quartile PR 1.7 (1.2–2.7)*                                                                                                                                                   |
| Dugandzic et al.      | PM10       | 2006           | First                       | Lowest vs highest quartile RR 1.33 (1.02–1.74)*                                                                                                                                                |
| Salam et al.          | PM10       | 2005           | Third                       | An increase of 20 $\mu\text{g}/\text{m}^3$ lowered 21.7g (1.1g–42.2g)* of birth weight                                                                                                         |
| Gouveia et al.        | PM10       | 2004           | Second                      | Lowest vs highest quartile AOR 1.25 (1.03–1.53)*                                                                                                                                               |
| Lin et al.            | PM10       | 2004           | Entire                      | Null associations                                                                                                                                                                              |
| Lee et al.            | PM10       | 2003           | First<br>Second             | IQR OR 1.03 (1.00–1.07)*<br>IQR OR 1.04 (1.00–1.08)*                                                                                                                                           |

3) The relationship between level of PM<sub>2.5</sub> exposure ( $\mu\text{g}/\text{m}^3$ ) and relative risk (95% CI) of child ALRI, based on the integrated exposure-response (IER) function, for (a) exposure over the range 0–600  $\mu\text{g}/\text{m}^3$



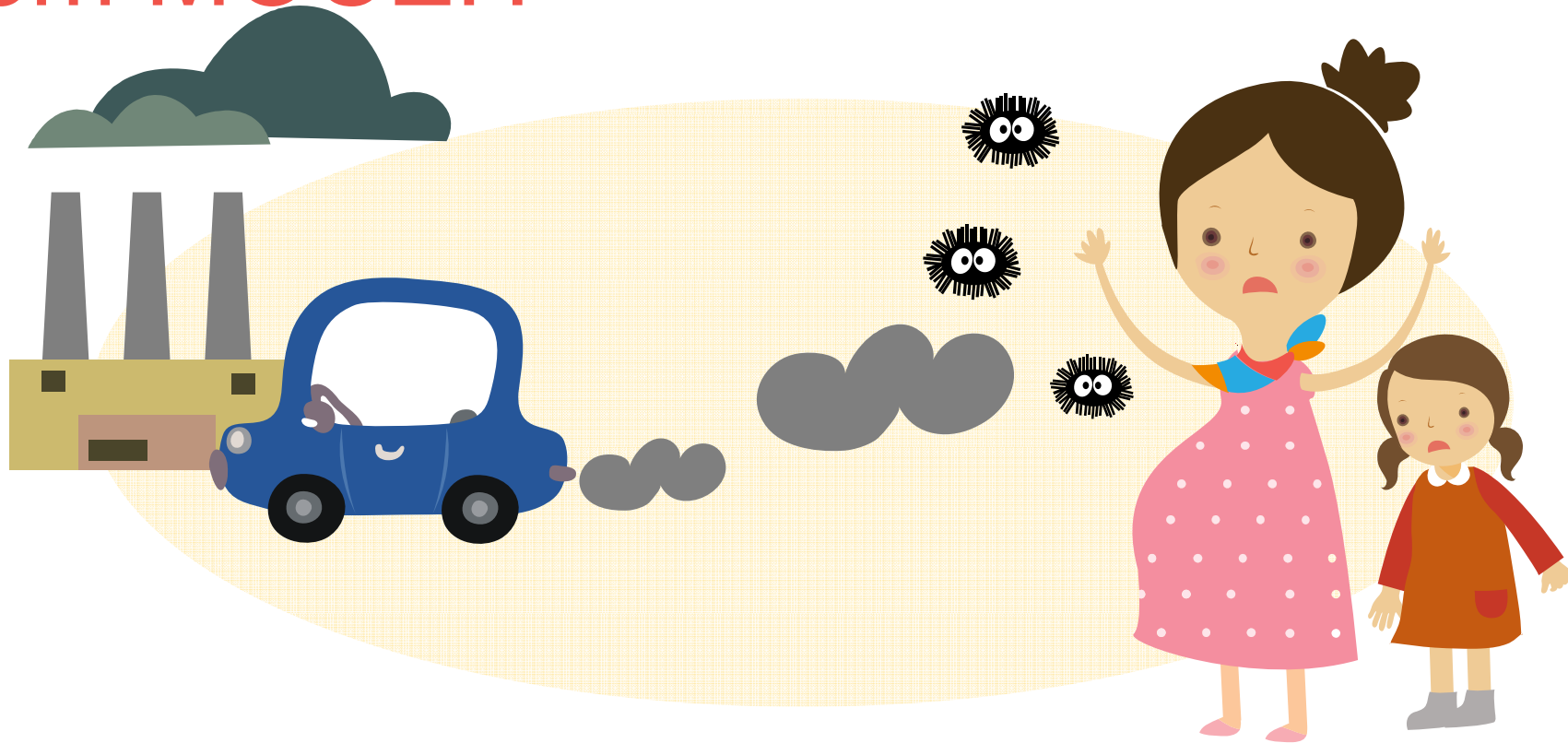
WHO indoor air quality guidelines: household fuel combustion - See more at: <http://apps.who.int/iris/handle/10665/141496#sthash.SyqtYxhA.dpuf>

(b) over the range 0–40  $\mu\text{g}/\text{m}^3$  which spans the WHO annual AQG for PM<sub>2.5</sub>



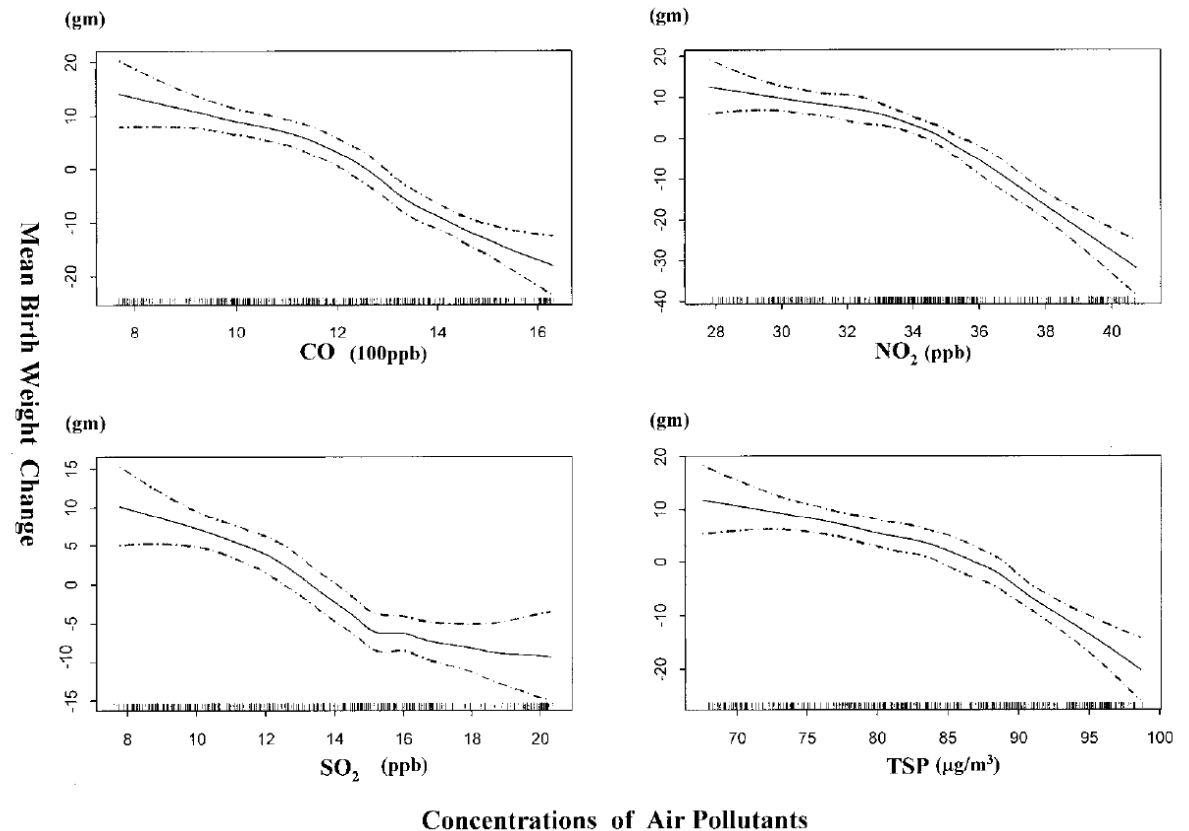
WHO indoor air quality guidelines: household fuel combustion - See more at: <http://apps.who.int/iris/handle/10665/141496#sthash.SyqtYxhA.dpuf>

# IV. Air pollution and Pregnancy outcome and children's health:some evidences from MOCEH



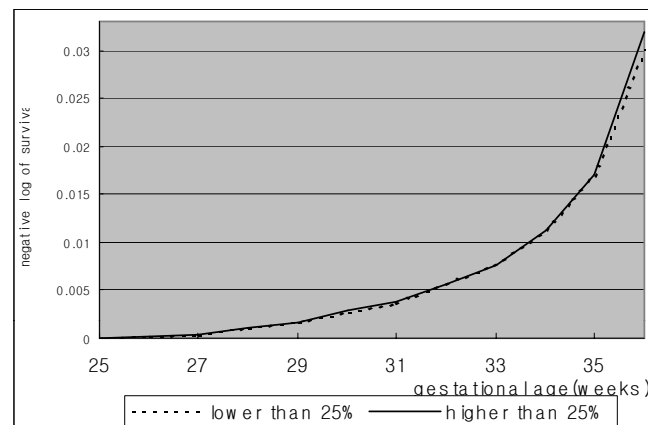
# 1. Mother graph of MOCEH

- Carbon monoxide, nitrogen dioxide, sulfur dioxide, and total suspended particle concentrations in the first trimester of pregnancy period are risk factors for low birth weight.

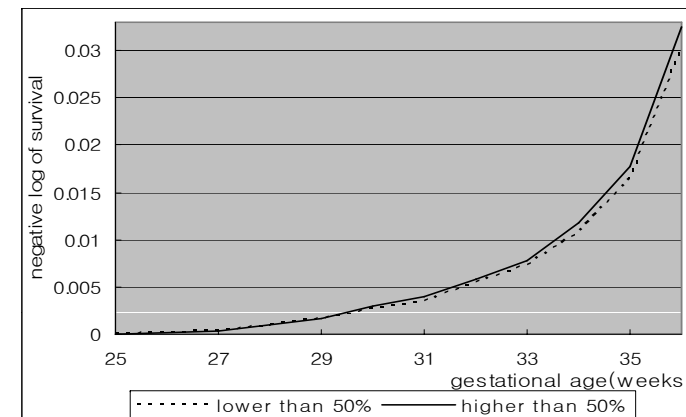


Eun-Hee Ha et al, Is Air Pollution a Risk Factor for Low Birth Weight in Seoul? EPIDEMIOLOGY, 2001

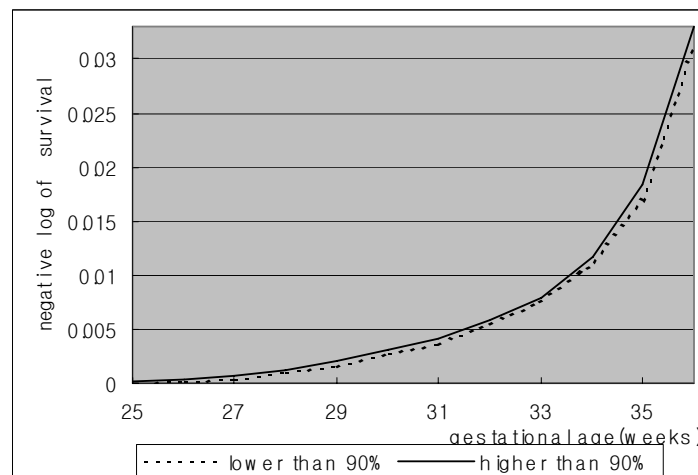
# Difference of cumulative hazard functions between low and high $PM_{10}$ concentration groups during the seventh month of pregnancy



(a) Divided on the basis of 25% value of  $PM_{10}$



(b) Divided on the basis of median of  $PM_{10}$



(c) Divided on the basis of 90% value of  $PM_{10}$

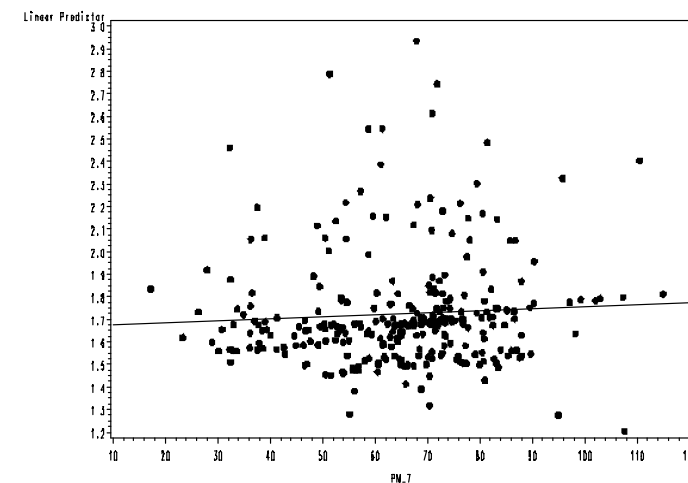
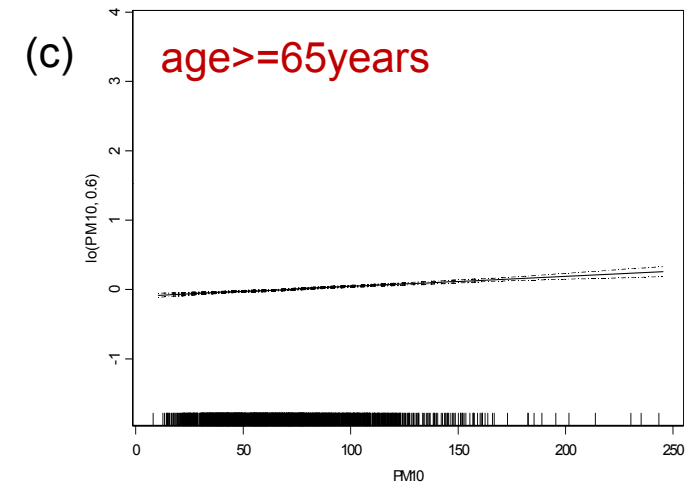
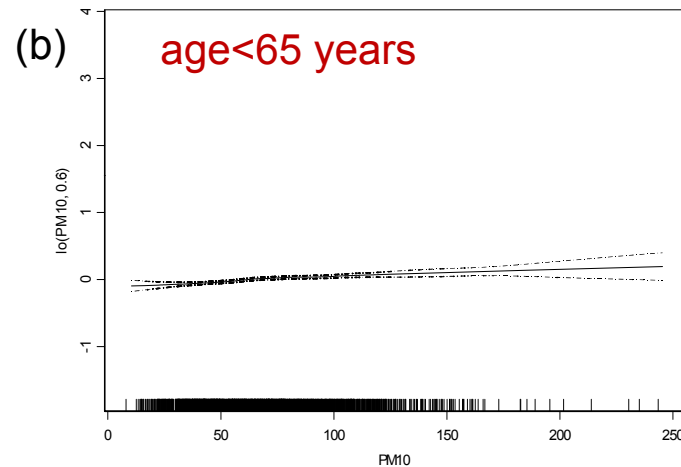
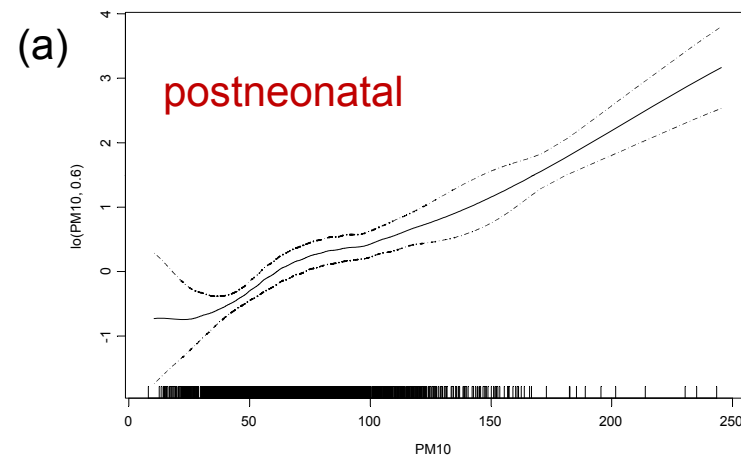


Figure 2. Difference of cumulative hazard functions between low and high  $PM_{10}$  concentration groups during the seventh month of pregnancy



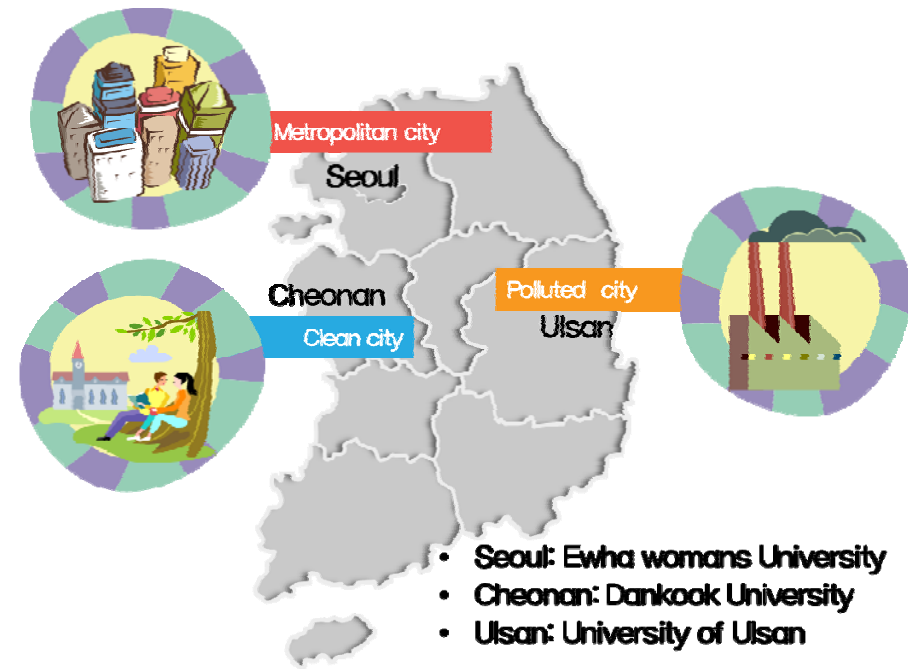
Regarding respiratory mortality, RR for an interquartile change of  $\text{PM}_{10}$  in postneonates (RR: 2.018; 95% CI: 1.784-2.283) was also greater than those in the other groups.

Infant susceptibility of mortality to air pollution in Seoul, South Korea.  
Ha EH et al, Pediatrics 2003

## 2. Brief Introduction to MOCEH

### 1) Scope of research

- In Korea, the National Environmental Health Action Plan (NEHAP, 2006) developed by the Ministry of Environment outlines strategies for eliminating environmental health hazards and reducing their adverse effects on pregnant women and their children.
- In 2006, the Mothers and Children's Environmental Health (MOCEH) study, a multi-center prospective cohort study, began as an initiative of the NEHAP.



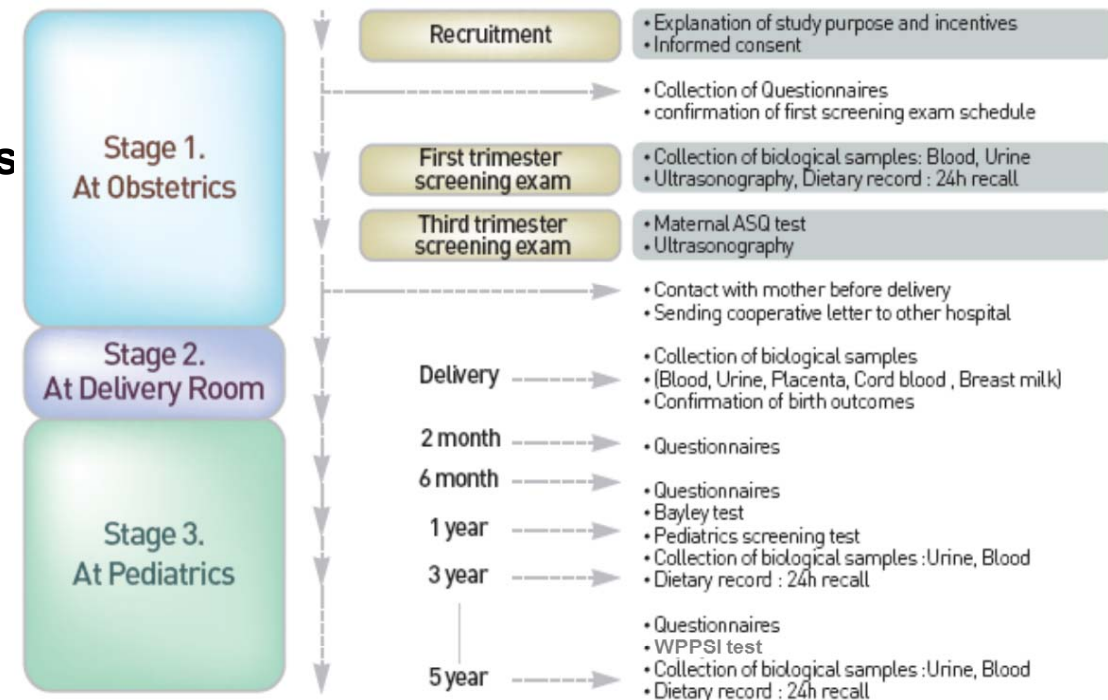
## 2) Study process

### ■ Assessment of determinants

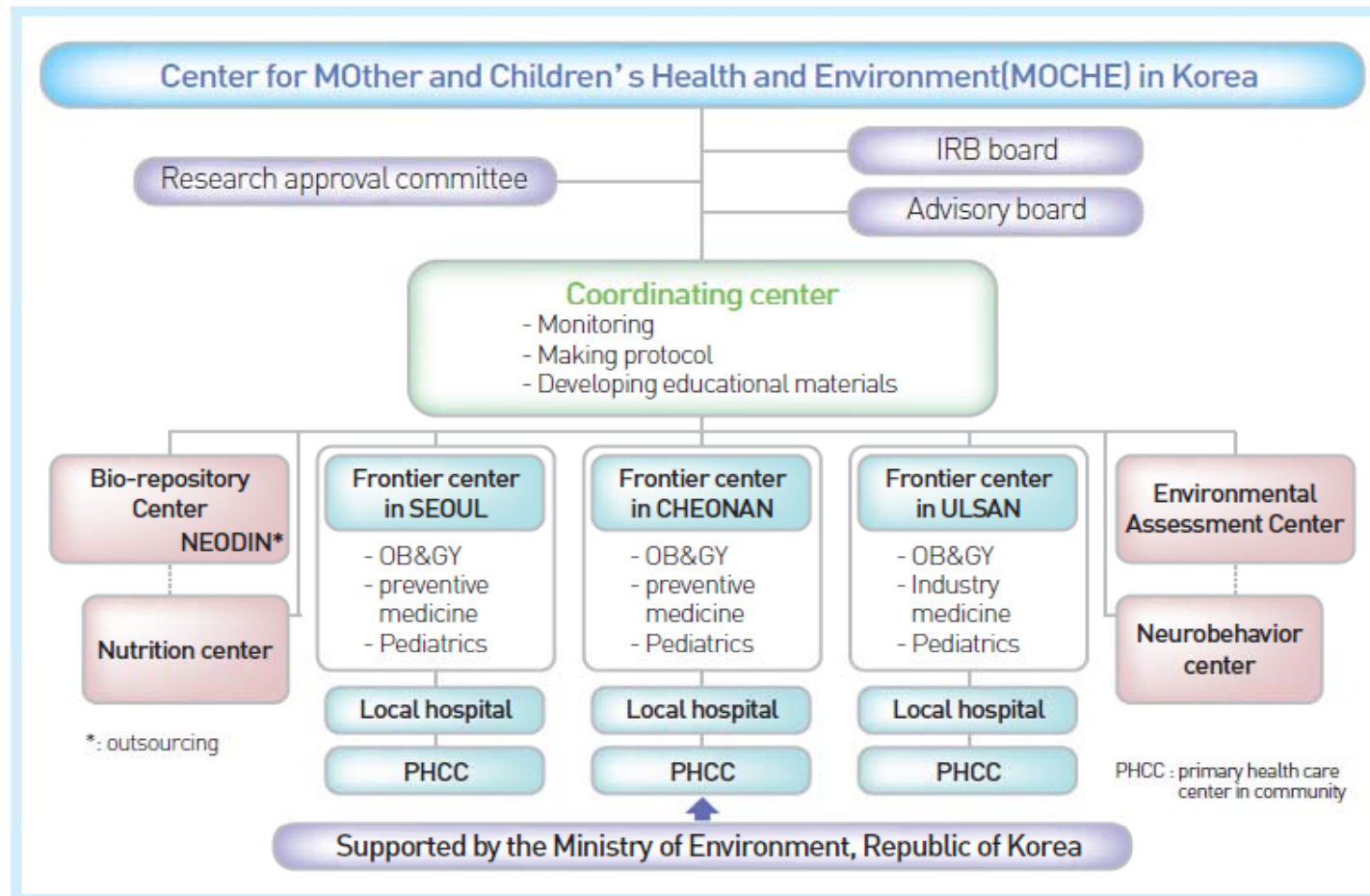
- Questionnaires
- **Measurements of environmental pollutants (Home visiting)**
- Biological samples
- Dietary information

### ■ Assessment of outcomes

- Intrauterine growth
- Birth outcomes
- Growth and development
- Neurocognitive development
- Atopy, allergy, and asthma in children

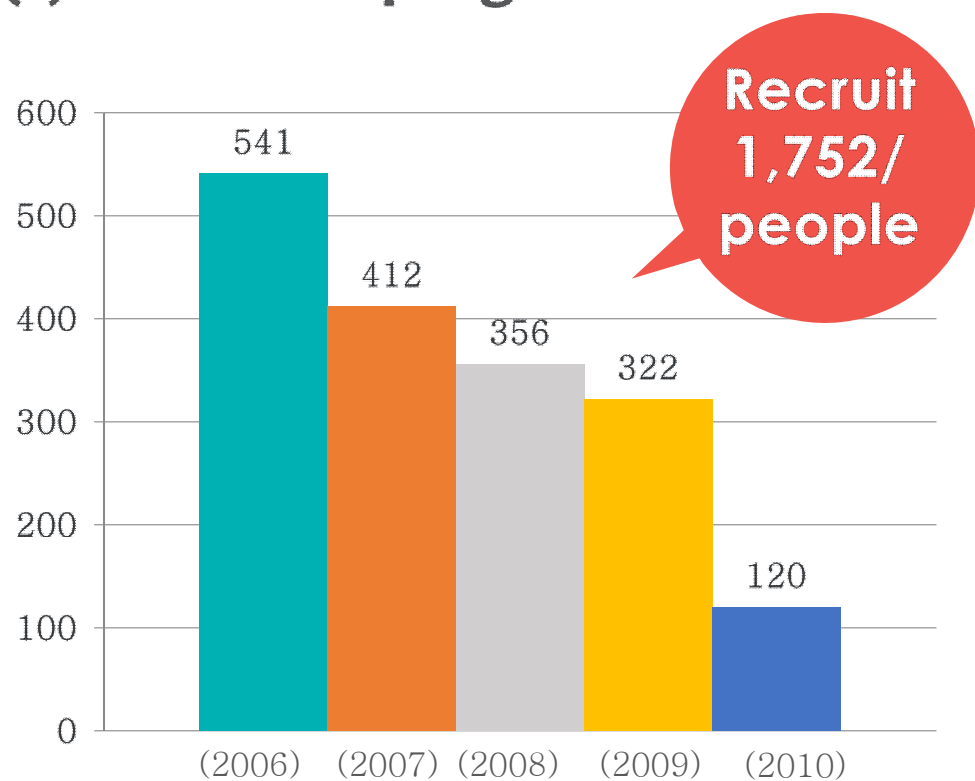


### 3) Community-Hospital based collaborative network

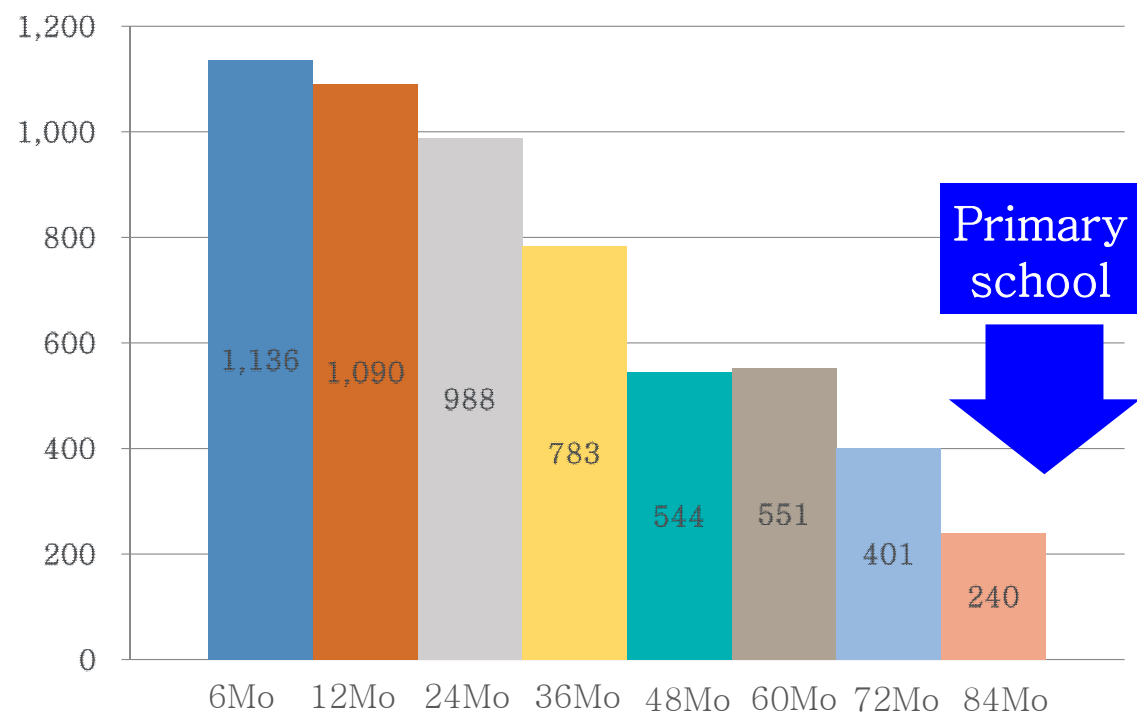


# 4) Recruitment Results

(1) Cohort of pregnant women



(2) Cohort of infants





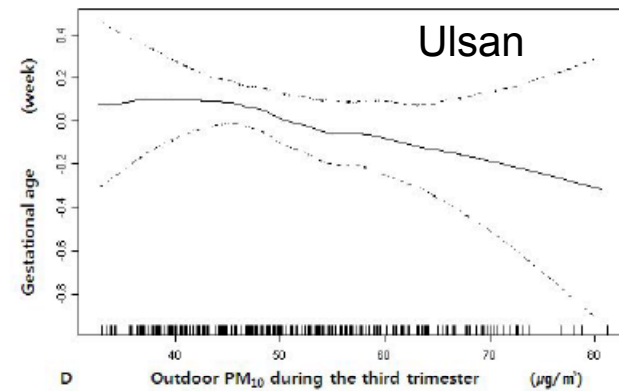
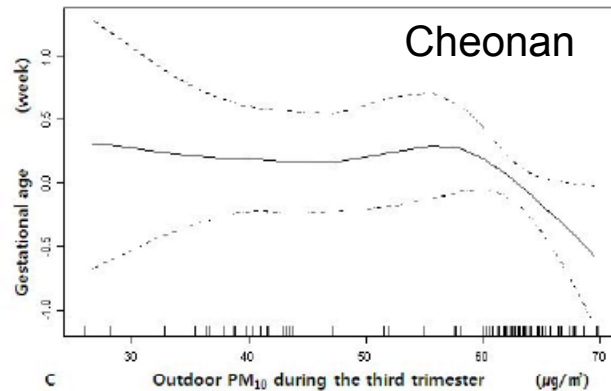
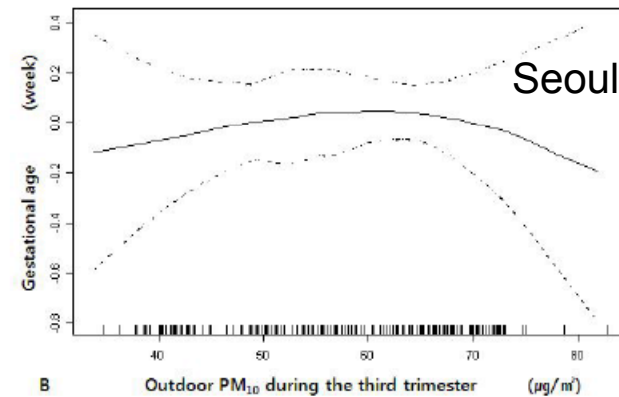
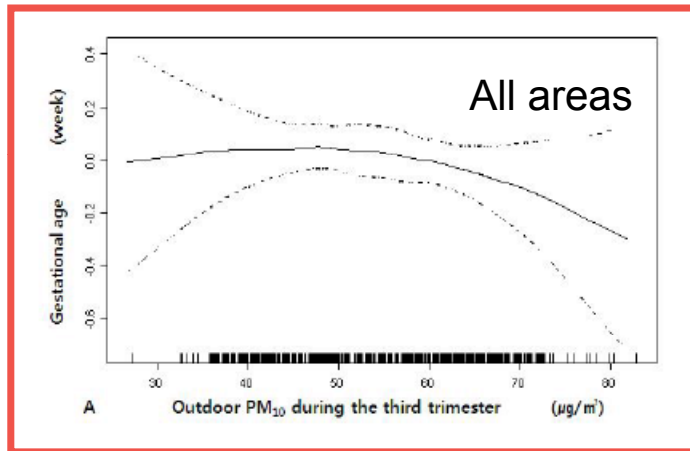
### **3. Air pollution and children's health: Some evidences from MOCEH study**

- Particulate Matter(PM10)**
- VOCs/HCHO**

- **Particulate Matter(PM10)**
- VOCs/HCHO



# 1) Gestational age for third trimester of outdoor PM<sub>10</sub>



\* Adjusted controlling for maternal age, maternal education, baby sex, BMI, house age, and factory

# 2) Personal exposure PM10 and growth development

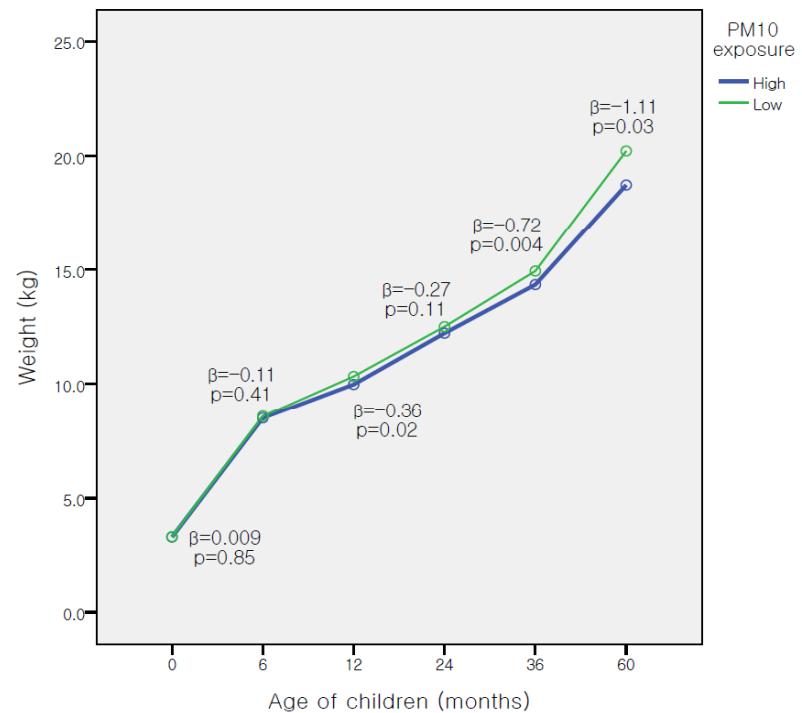
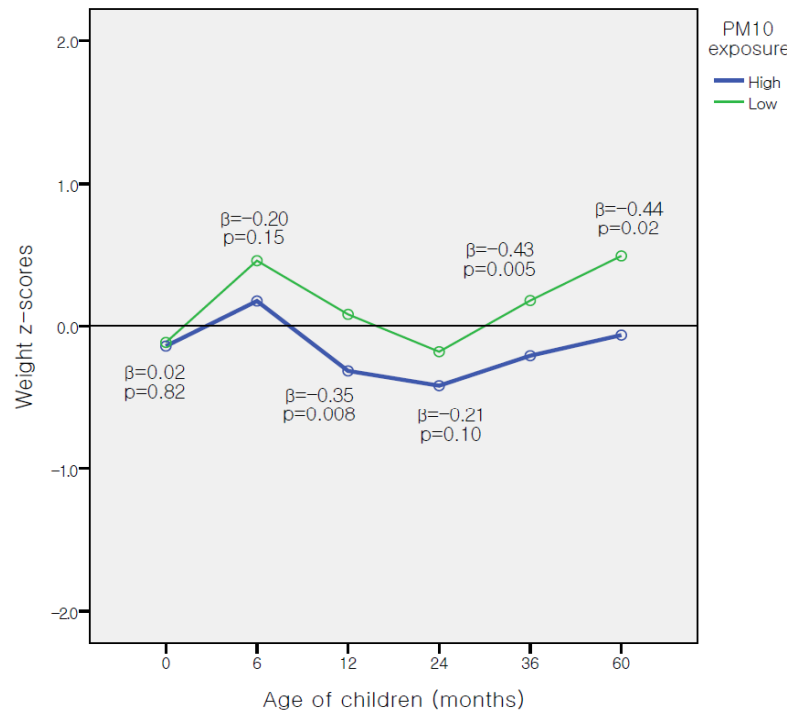
Korean Children's Environmental Health Study

## Association of PM10 exposure with children's weight z-scores (GLM)

\* The model 1 was adjusted for sex, maternal age, gestational age, maternal education, family income, birth weight, and maternal body mass index.  
The model 2 was adjusted for the same covariates in model 1 and feeding method at 6 months.

| Measured period                     |           | Change in weight for age z-scores (95% confidence intervals) |                             |     |                              |
|-------------------------------------|-----------|--------------------------------------------------------------|-----------------------------|-----|------------------------------|
| PM10 (10 $\mu\text{g}/\text{m}^3$ ) | Weight    | N                                                            | Model 1*                    | N   | Model 2*                     |
| Prenatal                            | At birth  | 1,129                                                        | -0.004 (-0.07, 0.06)        | N/A | N/A                          |
|                                     | 6 months  | 812                                                          | -0.02 (-0.12, 0.09)         | 740 | -0.03 (-0.14, 0.08)          |
|                                     | 12 months | 728                                                          | -0.09 (-0.19, 0.007)        | 638 | <b>-0.11 (-0.21, -0.002)</b> |
|                                     | 24 months | 648                                                          | -0.06 (-0.17, 0.05)         | 595 | -0.03 (-0.14, 0.09)          |
|                                     | 36 months | 545                                                          | -0.06 (-0.19, 0.07)         | 481 | -0.06 (-0.20, 0.07)          |
|                                     | 60 months | 387                                                          | -0.12 (-0.29, 0.06)         | 340 | -0.16 (-0.35, 0.03)          |
| 0-6 months                          | 6 months  | 812                                                          | -0.07 (-0.15, 0.01)         | 740 | -0.08 (-0.17, 0.006)         |
|                                     | 12 months | 728                                                          | 0.02 (-0.06, 0.10)          | 638 | 0.002 (-0.08, 0.08)          |
|                                     | 24 months | 648                                                          | -0.01 (-0.10, 0.09)         | 595 | 0.004 (-0.09, 0.10)          |
|                                     | 36 months | 545                                                          | 0.08 (-0.02, 0.19)          | 481 | 0.01 (-0.01, 0.21)           |
|                                     | 60 months | 387                                                          | -0.04 (-0.17, 0.08)         | 340 | -0.08 (-0.22, 0.06)          |
| 7-12 months                         | 12 months | 728                                                          | <b>-0.13 (-0.21, -0.05)</b> | 638 | <b>-0.14 (-0.22, -0.05)</b>  |
|                                     | 24 months | 648                                                          | -0.08 (-0.18, 0.01)         | 595 | -0.06 (-0.15, 0.04)          |
|                                     | 36 months | 545                                                          | <b>-0.15 (-0.25, -0.04)</b> | 481 | <b>-0.16 (-0.37, -0.05)</b>  |
|                                     | 60 months | 387                                                          | <b>-0.17 (-0.29, -0.04)</b> | 340 | <b>-0.19 (-0.34, -0.06)</b>  |
| 13-24 months                        | 24 months | 648                                                          | -0.12 (-0.26, 0.02)         | 595 | -0.07 (-0.23, 0.08)          |
|                                     | 36 months | 545                                                          | -0.03 (-0.19, 0.13)         | 481 | -0.02 (-0.29, 0.16)          |
|                                     | 60 months | 387                                                          | -0.14 (-0.33, 0.06)         | 340 | -0.17 (-0.38, 0.05)          |
| 25-36 months                        | 36 months | 545                                                          | -0.02 (-0.17, 0.14)         | 481 | -0.02 (-0.19, 0.16)          |
|                                     | 60 months | 387                                                          | -0.17 (-0.37, 0.04)         | 340 | -0.20 (-0.45, 0.04)          |
| 37-60 months                        | 60 months | 286                                                          | -0.19 (-0.45, 0.07)         | 248 | -0.22 (-0.51, 0.08)          |

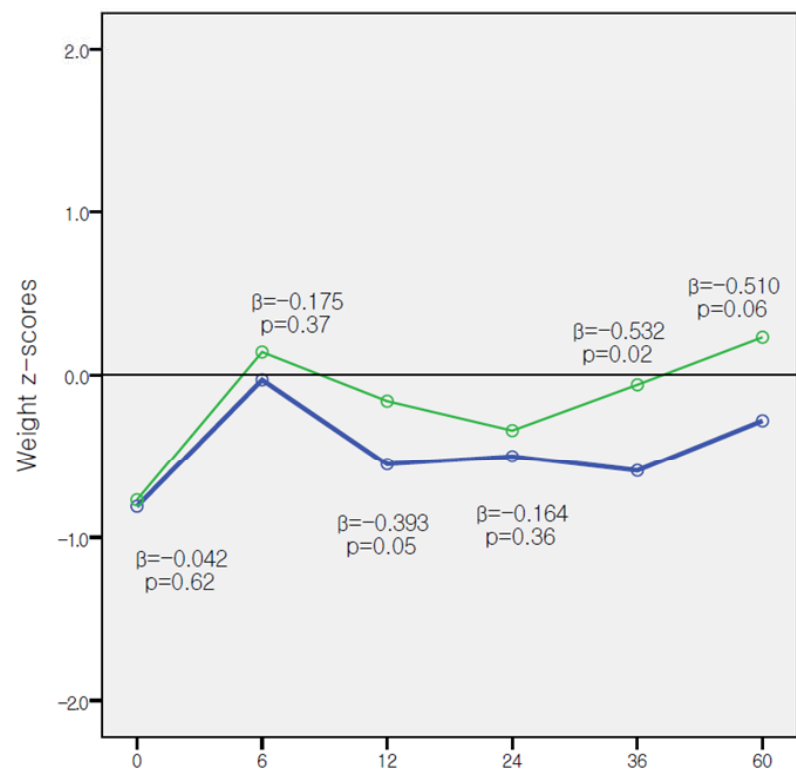
- Difference of average weight for age z scores and weights between children exposed to PM10 under  $50 \mu\text{g}/\text{m}^3$  and over  $50 \mu\text{g}/\text{m}^3$  from pregnancy to 24 months



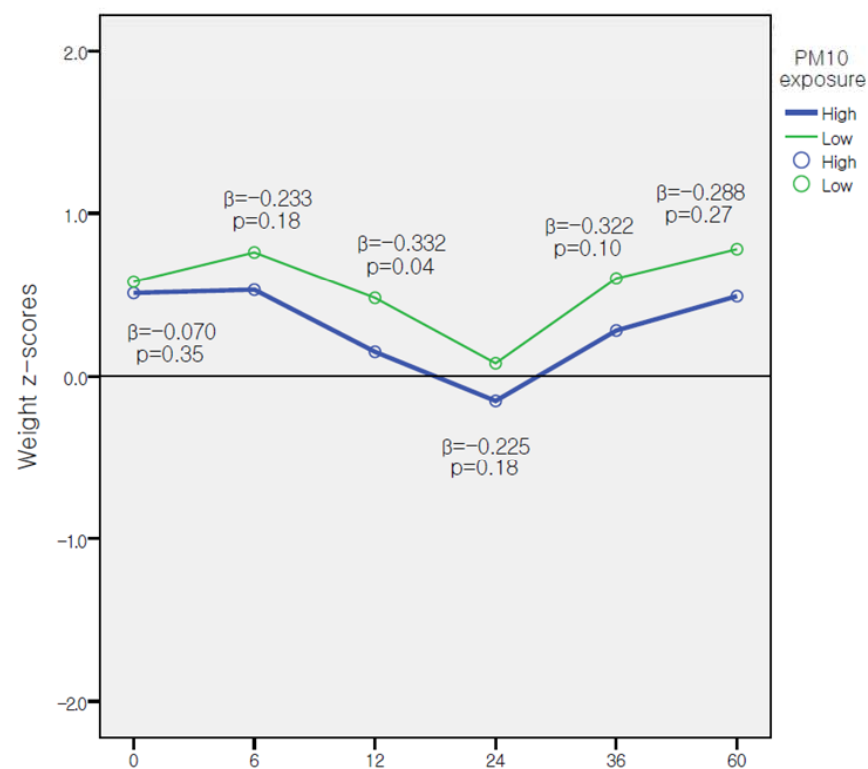
\* Total number of participants in the analysis (the number of participants in high exposure group)

At birth: 380 (278), 6 months: 238 (152), 12 months: 213 (137), 24 months: 199 (124), 36 months: 179 (113), 60 months: 124 (79)

- Difference of average weight z-scores between children exposed to PM10 under  $50 \mu\text{g}/\text{m}^3$  and over  $50 \mu\text{g}/\text{m}^3$  from pregnancy to 24 months in children with birth weight under 3.3 kg and over 3.3 kg



Age of children with birth weight under 3.3kg (months)



Age of children with birth weight over 3.3kg (months)

\* Total number of participants in the analysis (the number of participants in higher exposure group)

Left - At birth: 183 (132), 6 months: 114 (72), 12 months: 102 (67), 24 months: 102 (65), 36 months: 87 (56), 60 months: 62 (42)

Right - At birth: 197 (146), 6 months: 124 (80), 12 months: 111 (70), 24 months: 97 (59), 36 months: 97 (57), 60 months: 62 (37)

### 3) Air pollutant and neurodevelopment

Association of prenatal exposures to PM<sub>10</sub> and NO<sub>2</sub> with children's neurodevelopment over the period of 24 months of age by general linear model and GEEs.<sup>a</sup>

**Table 4**

Association of prenatal exposures to PM<sub>10</sub> and NO<sub>2</sub> with children's neurodevelopment over the period of 24 months of age by the general linear model and GEEs<sup>a</sup> — sub-analysis adjusted for maternal IQ.

|                                                          |                | n                | MDI                  |         | PDI                   |         |
|----------------------------------------------------------|----------------|------------------|----------------------|---------|-----------------------|---------|
|                                                          |                |                  | β (LCI, UCI)         | p-Value | β (LCI, UCI)          | p-Value |
| <i>PM<sub>10</sub> (10 µg/m<sup>3</sup>)<sup>b</sup></i> |                |                  |                      |         |                       |         |
| GLM <sup>b</sup>                                         | 6 months       | 240              | −4.91 (−7.69, −2.12) | <0.001  | −8.85 (−12.14, −5.57) | <0.001  |
|                                                          | 12 months      | 184              | −3.96 (−8.42, 0.51)  | 0.08    | −1.30 (−5.53, 2.93)   | 0.68    |
|                                                          | 24 months      | 184              | 1.05 (−2.69, 4.79)   | 0.58    | 1.93 (−1.74, 5.60)    | 0.91    |
| GEEs <sup>c</sup>                                        | 6 to 24 months | 240 <sup>e</sup> | −2.81 (−5.17, −0.46) | 0.02    | −3.57 (−5.96, −1.18)  | 0.004   |
| GEEs <sup>d</sup>                                        | 6 to 24 months | 240 <sup>e</sup> | −3.41 (−5.88, −0.94) | 0.007   | −3.93 (−6.37, −1.48)  | 0.002   |
| <i>NO<sub>2</sub> (10 ppb)<sup>b</sup></i>               |                |                  |                      |         |                       |         |
| GLM <sup>b</sup>                                         | 6 months       | 240              | −4.11 (−6.26, −1.95) | <0.001  | −5.57 (−8.17, −2.97)  | <0.001  |
|                                                          | 12 months      | 184              | −0.85 (−4.86, 3.16)  | 0.55    | −0.91 (−4.68, 2.86)   | 0.63    |
|                                                          | 24 months      | 184              | −0.18 (−3.23, 2.88)  | 0.30    | −2.39 (−5.37, 0.60)   | 0.12    |
| GEEs <sup>c</sup>                                        | 6 to 24 months | 240 <sup>e</sup> | −2.08 (−4.09, −0.08) | 0.04    | −3.48 (−5.64, −1.32)  | 0.002   |
| GEEs <sup>d</sup>                                        | 6 to 24 months | 240 <sup>e</sup> | −1.66 (−3.70, 0.38)  | 0.11    | −3.19 (−5.36, −1.02)  | 0.004   |

<sup>a</sup> Models were adjusted for sex, birth weight, maternal age, gestational age, maternal education, family income, and maternal IQ.

<sup>b</sup> General linear model analysis was used to see the effect of prenatal exposure on neurodevelopment at 6, 12, and 24 months.

<sup>c</sup> General estimating equations were used to see the effect of prenatal exposure on neurodevelopment over the 24 months with repeated outcome measures.

<sup>d</sup> General estimating equations without adjusting for maternal IQ were performed on the same subanalysis group to compare the result with the previous model of sub-analysis.

<sup>e</sup> Total number of mothers who underwent Korean Wechsler Adult Intelligence Scale.

# 4) Air pollutant and Allergy disease

Association of prenatal exposures to PM10 and NO2 on pneumonia, bronchitis, and bronchiolitis at 6 and 12 months of age

| Period       | Measures      | MODEL 1 <sup>a</sup> |       |      |            |       |      |               |       |      | MODEL 2 <sup>b</sup> |       |      |            |       |      |               |       |      |
|--------------|---------------|----------------------|-------|------|------------|-------|------|---------------|-------|------|----------------------|-------|------|------------|-------|------|---------------|-------|------|
|              |               | Pneumonia            |       |      | Bronchitis |       |      | Bronchiolitis |       |      | Pneumonia            |       |      | Bronchitis |       |      | Bronchiolitis |       |      |
|              |               | n                    | OR    | P    | n          | OR    | P    | n             | OR    | P    | n                    | OR    | P    | n          | OR    | P    | n             | OR    | P    |
| At 6 months  |               |                      |       |      |            |       |      |               |       |      |                      |       |      |            |       |      |               |       |      |
| Prenatal     | PM10 Indoor   | 128                  | 1.002 | 0.86 | 128        | 1.058 | 0.03 | 128           | 1.007 | 0.41 | 128                  | 1.003 | 0.82 | 128        | 1.087 | 0.02 | 128           | 1.007 | 0.42 |
|              | PM10 outdoor  | 61                   | 0.999 | 0.94 | 61         | 1.014 | 0.39 | 61            | 0.985 | 0.24 | 56                   | 1.001 | 0.96 | 56         | 1.026 | 0.24 | 56            | 0.959 | 0.08 |
|              | PM2.5 Indoor  | 128                  | 0.993 | 0.78 | 128        | 1.094 | 0.03 | 128           | 0.978 | 0.30 | 128                  | 0.996 | 0.87 | 128        | 1.148 | 0.02 | 128           | 0.978 | 0.28 |
|              | PM2.5 outdoor | 62                   | 0.908 | 0.30 | 62         | 1.029 | 0.15 | 62            | 0.965 | 0.21 | 57                   | 0.896 | 0.25 | 57         | 1.061 | 0.22 | 57            | 0.943 | 0.07 |
| postnatal    | NO2 Indoor    | 130                  | 0.925 | 0.35 | 130        | 0.987 | 0.87 | 130           | 1.013 | 0.70 | 114                  | 0.898 | 0.25 | 114        | 1.008 | 0.93 | 114           | 1.014 | 0.71 |
|              | NO2 personal  | 102                  | 0.934 | 0.35 | 102        | 0.971 | 0.68 | 102           | 1.029 | 0.37 | 92                   | 0.917 | 0.30 | 92         | 0.981 | 0.82 | 92            | 0.897 | 0.19 |
|              |               |                      |       |      |            |       |      |               |       |      |                      |       |      |            |       |      |               |       |      |
| At 12 months |               |                      |       |      |            |       |      |               |       |      |                      |       |      |            |       |      |               |       |      |
| Prenatal     | PM10 Indoor   | 114                  | 0.988 | 0.33 | 114        | 1.01  | 0.33 | 114           | 0.998 | 0.82 | 114                  | 0.989 | 0.39 | 114        | 1.01  | 0.31 | 114           | 0.997 | 0.81 |
|              | PM10 outdoor  | 59                   | 0.996 | 0.78 | 59         | 1.014 | 0.34 | 59            | 0.96  | 0.10 | 54                   | 0.992 | 0.70 | 54         | 1.064 | 0.08 | 54            | 0.918 | 0.14 |
|              | PM2.5 Indoor  | 114                  | 0.982 | 0.46 | 114        | 1.03  | 0.14 | 114           | 0.974 | 0.34 | 114                  | 0.985 | 0.56 | 114        | 1.043 | 0.08 | 114           | 0.971 | 0.31 |
|              | PM2.5 outdoor | 60                   | 0.989 | 0.64 | 60         | 1.021 | 0.21 | 60            | 0.963 | 0.33 | 55                   | 0.985 | 0.61 | 55         | 1.103 | 0.05 | 55            | 0.972 | 0.55 |
| postnatal    | NO2 Indoor    | 127                  | 0.932 | 0.16 | 127        | 0.976 | 0.65 | 127           | 1.028 | 0.41 | 112                  | 0.94  | 0.25 | 112        | 0.974 | 0.68 | 112           | 1.052 | 0.18 |
|              | NO2 personal  | 106                  | 0.911 | 0.11 | 106        | 0.983 | 0.74 | 106           | 1.028 | 0.42 | 94                   | 0.917 | 0.08 | 94         | 0.977 | 0.72 | 94            | 1.088 | 0.06 |

<sup>a</sup> Model 1 was adjusted for birth weight, infant sex, gestational age, maternal age, maternal history of allergic diseases, and paternal history of allergic diseases.

<sup>b</sup> Model 2 was adjusted for birth weight, infant sex, gestational age, maternal age, maternal history of allergic diseases, paternal history of allergic diseases, maternal education, and family income.

# Association of prenatal exposures to PM10 and NO2 asthma, allergic diseases, and atopic dermatitis at 6 and 12 months of age

<sup>a</sup> Model 1 was adjusted for birth weight, infant sex, gestational age, maternal age, maternal history of allergic diseases, and paternal history of allergic diseases.

<sup>b</sup> Model 2 was adjusted for birth weight, infant sex, gestational age, maternal age, maternal history of allergic diseases, paternal history of allergic diseases, maternal education, and family income.

| Period       | Measures      | MODEL 1 <sup>a</sup> |       |      |                   |       |      |                   |       |      | MODEL 2 <sup>b</sup> |       |      |                   |       |      |                   |       |      |
|--------------|---------------|----------------------|-------|------|-------------------|-------|------|-------------------|-------|------|----------------------|-------|------|-------------------|-------|------|-------------------|-------|------|
|              |               | Asthma               |       |      | Allergic diseases |       |      | Atopic dermatitis |       |      | Asthma               |       |      | Allergic diseases |       |      | Atopic dermatitis |       |      |
|              |               | n                    | OR    | P    | n                 | OR    | P    | n                 | OR    | P    | n                    | OR    | P    | n                 | OR    | P    | n                 | OR    | P    |
| At 6 months  |               |                      |       |      |                   |       |      |                   |       |      |                      |       |      |                   |       |      |                   |       |      |
| Prenatal     | PM10 indoor   | 128                  | 1.364 | 0.41 | 139               | 1.011 | 0.26 | 159               | 0.996 | 0.44 | 128                  | 1.217 | 0.37 | 139               | 1.012 | 0.26 | 159               | 0.996 | 0.48 |
|              | PM10 outdoor  | 61                   | 1.144 | 0.76 | 66                | 1.019 | 0.05 | 69                | 0.995 | 0.45 | 56                   | 1.124 | 0.77 | 61                | 1.031 | 0.02 | 63                | 0.996 | 0.56 |
|              | PM2.5 indoor  | 128                  | 1.532 | 0.20 | 139               | 1.025 | 0.23 | 159               | 1.009 | 0.42 | 128                  | 0.997 | 0.27 | 139               | 1.029 | 0.20 | 159               | 0.998 | 0.29 |
|              | PM2.5 outdoor | 62                   | 1.194 | 0.73 | 67                | 1.024 | 0.05 | 70                | 0.994 | 0.56 | 57                   | 1.165 | 0.82 | 62                | 1.035 | 0.02 | 64                | 0.995 | 0.62 |
| postnatal    | NO2 indoor    | 130                  | 2.561 | 0.53 | 145               | 1.012 | 0.80 | 161               | 1.034 | 0.10 | 114                  | 1.013 | 0.99 | 129               | 1.021 | 0.66 | 139               | 1.046 | 0.06 |
|              | NO2 personal  | 102                  | 1.226 | 0.63 | 111               | 1.023 | 0.63 | 127               | 1.031 | 0.15 | 92                   | 1.572 | 0.86 | 101               | 1.102 | 0.14 | 111               | 1.042 | 0.10 |
| At 12 months |               |                      |       |      |                   |       |      |                   |       |      |                      |       |      |                   |       |      |                   |       |      |
| Prenatal     | PM10 indoor   | 114                  | 0.998 | 0.82 | 114               | 1.026 | 0.05 | 136               | 0.997 | 0.61 | 114                  | 1.415 | 0.81 | 114               | 1.028 | 0.04 | 136               | 1.012 | 0.79 |
|              | PM10 outdoor  | 59                   | 0.96  | 0.10 | 59                | 1.01  | 0.41 | 64                | 1     | 0.96 | -                    | -     | -    | 54                | 1.072 | 0.09 | 57                | 1.003 | 0.77 |
|              | PM2.5 indoor  | 114                  | 0.974 | 0.34 | 114               | 1.014 | 0.61 | 136               | 0.994 | 0.65 | -                    | -     | -    | 114               | 1.019 | 0.52 | 136               | 1.003 | 0.86 |
|              | PM2.5 outdoor | 60                   | 0.963 | 0.33 | 60                | 1.015 | 0.34 | 65                | 1     | 0.97 | -                    | -     | -    | 55                | 1.077 | 0.04 | 58                | 1.004 | 0.78 |
| postnatal    | NO2 indoor    | 127                  | 1.09  | 0.62 | 127               | 0.953 | 0.43 | 151               | 1.028 | 0.21 | 112                  | 2.727 | 0.54 | 112               | 0.946 | 0.40 | 130               | 1.043 | 0.11 |
|              | NO2 personal  | 106                  | 1.044 | 0.76 | 106               | 0.84  | 0.07 | 126               | 1.029 | 0.18 | 94                   | 2.189 | 0.71 | 94                | 0.848 | 0.09 | 109               | 1.055 | 0.06 |

- The relationships between PM2.5 and house dust mite allergen or eosinophil number in group of case-control

| Case                     | Crude model |      |      | Adjusted model** |      |      |
|--------------------------|-------------|------|------|------------------|------|------|
|                          | $\beta$     | se   | P    | $\beta$          | se   | P    |
| Indoor PM2.5             |             |      |      |                  |      |      |
| House dust mite allergen | 0.29        | 0.13 | 0.03 | 0.30             | 0.13 | 0.03 |
| Eosinophil number        | 19.60       | 7.54 | 0.02 | 23.93            | 8.95 | 0.02 |
| Outdoor PM2.5            |             |      |      |                  |      |      |
| House dust mite allergen | 0.03        | 0.25 | 0.91 | 0.09             | 0.30 | 0.78 |
| Eosinophil number        | 7.36        | 5.77 | 0.22 | 8.58             | 7.68 | 0.29 |
| Control                  | Crude model |      |      | Adjusted model** |      |      |
|                          | $\beta$     | se   | P    | $\beta$          | se   | P    |
| Indoor PM2.5.            |             |      |      |                  |      |      |
| House dust mite allergen | 0.08        | 0.13 | 0.54 | -0.03            | 0.13 | 0.83 |
| Eosinophil number        | -0.01       | 2.09 | 1.00 | -1.22            | 2.29 | 0.61 |
| Outdoor PM2.5            |             |      |      |                  |      |      |
| House dust mite allergen | -0.01       | 0.17 | 0.97 | -0.02            | 0.17 | 0.91 |
| Eosinophil number        | -1.92       | 1.71 | 0.28 | -1.99            | 1.75 | 0.28 |

- PM10
- **VOCs/HCHO**



# 1) TVOCs/HCHO and Pregnancy Outcome

- Associations between TVOC and HCHO exposure and birth weight

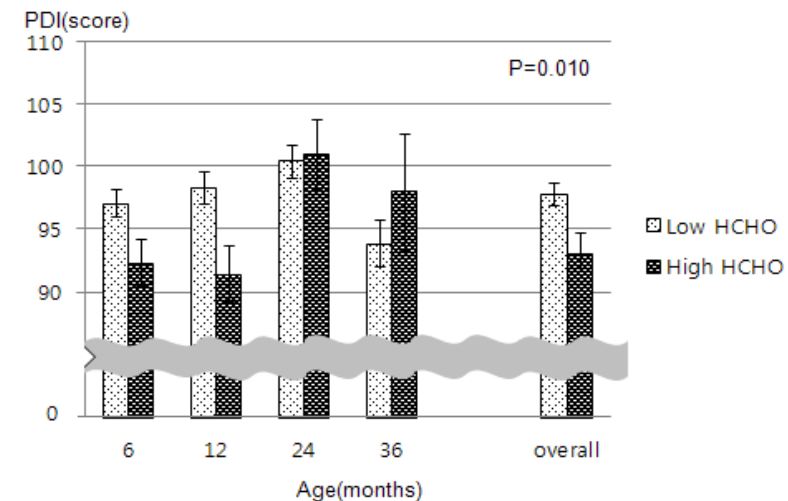
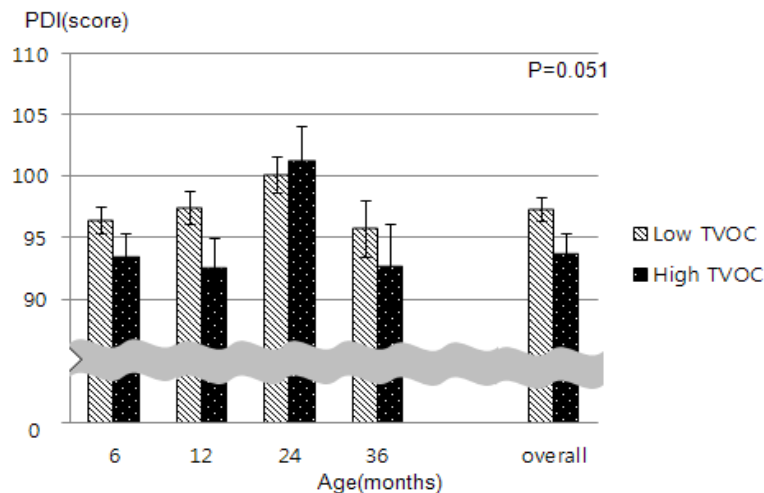
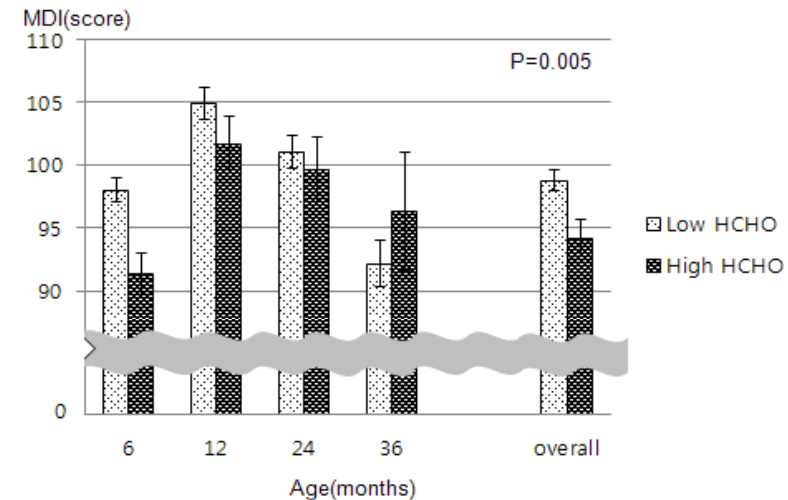
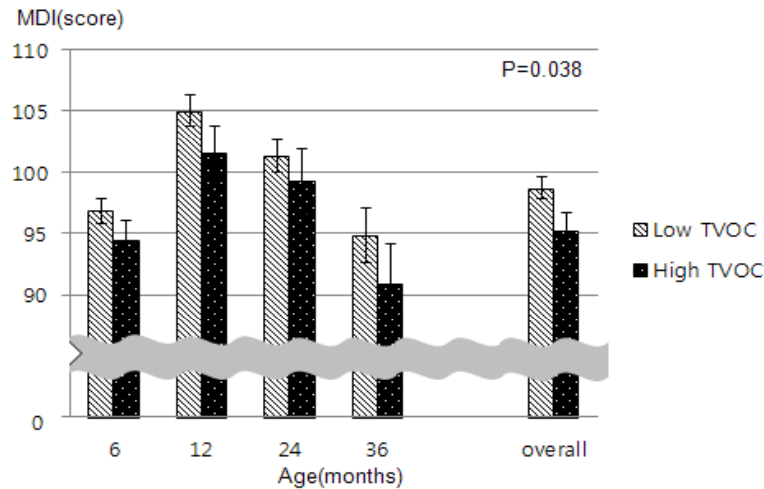
| Birth weight<br>(g) | Crude model |       |      | Model 1†‡ |       |       |
|---------------------|-------------|-------|------|-----------|-------|-------|
|                     | $\beta$     | SE    | P    | $\beta$   | SE    | P     |
| TVOC(log)           | -30.13      | 21.25 | 0.16 | -45.89    | 22.36 | 0.04* |
| HCHO(log)           | -11.21      | 36.43 | 0.76 | -37.98    | 39.55 | 0.34  |

\*P<0.05

†Adjusted for maternal age, gestational age at delivery, pre pregnancy BMI, educational level, birth order, and infant's gender.

‡A adjusted for plus,  
separated kitchen (TVOC); air cleaner and house age (HCHO).

## 2) Associations between ambient personal TVOC exposure during pregnancy and neuro-behavioral development(BSID-II) in infants



### 3) Indoor total volatile organic compounds exposure at 6 months followed by atopic dermatitis

**Table 4** Associations between the level of total VOC exposure and atopic dermatitis

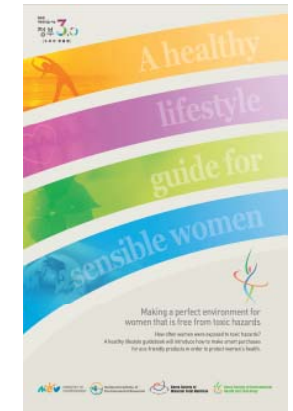
| VOCs exposure            | No. of total subjects | No. of atopic dermatitis | Unadjusted analysis odd ratio (95% CI) | Adjusted analysis odd ratio (95% CI) |
|--------------------------|-----------------------|--------------------------|----------------------------------------|--------------------------------------|
| Low TVOCs (<75th) group  | 71                    | 20 (27.8%)               | 1.0                                    | 1.0                                  |
| High TVOCs (≥75th) group | 34                    | 16 (48.5%)               | 2.447 (1.010–5.757)                    | 3.116 (1.041–9.323)                  |

These odds ratios and 95% CIs adjusted for allergy history of parents, household income, and breastfed longer than 6 month.

VOCs, volatile organic compounds.

High and low VOCs divided by upper 75% of the VOCs distribution of the study subjects (242.1  $\mu\text{g}/\text{m}^3$ ).

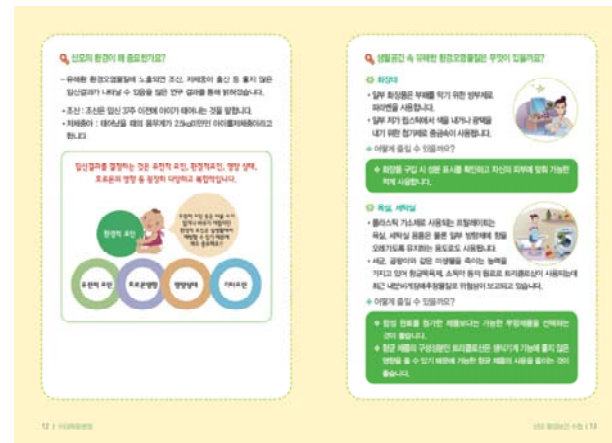
# Guidelines



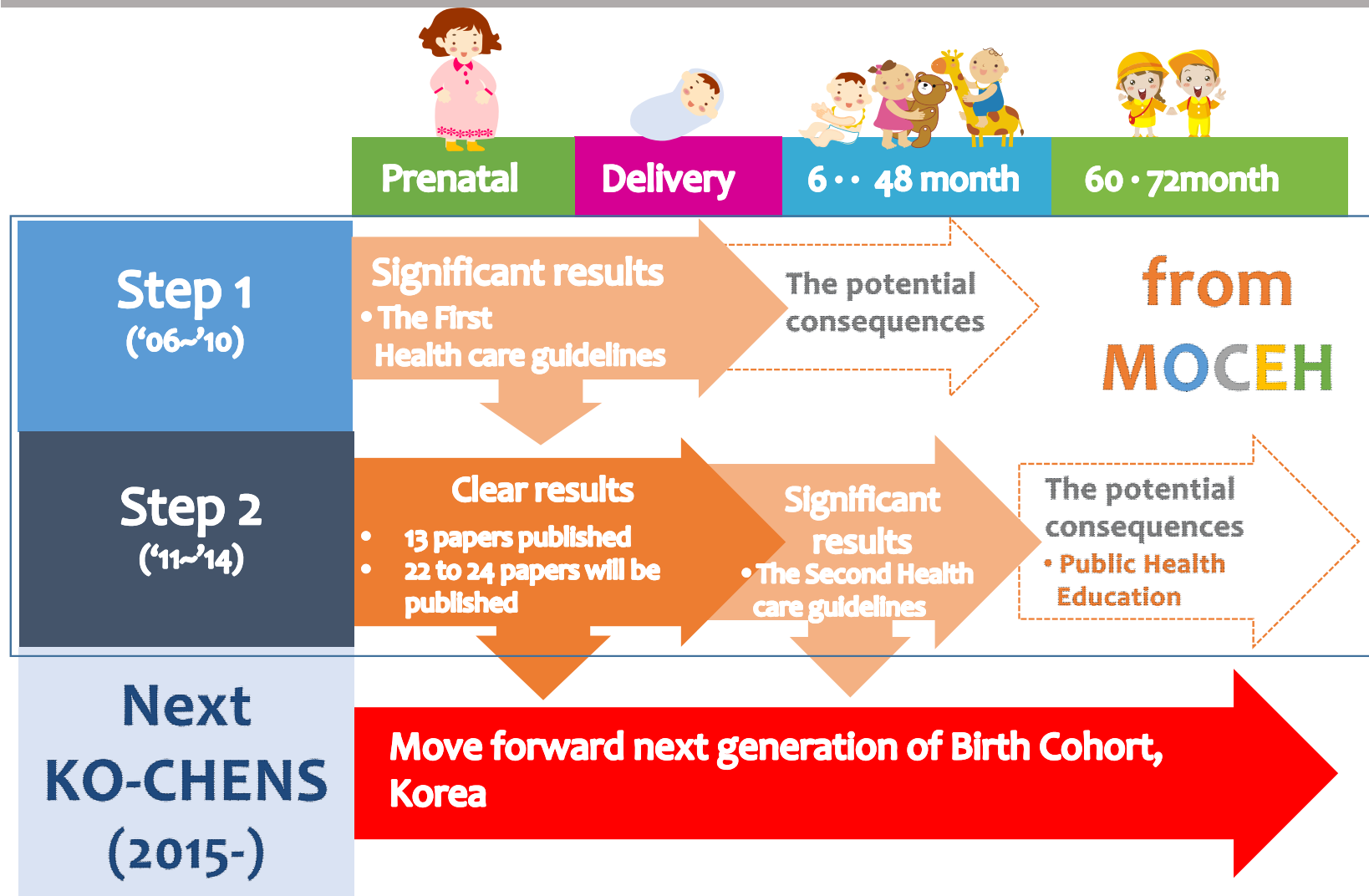
# Application

# Pamphlet

# Booklets



## Linkage of existing research and national initiatives



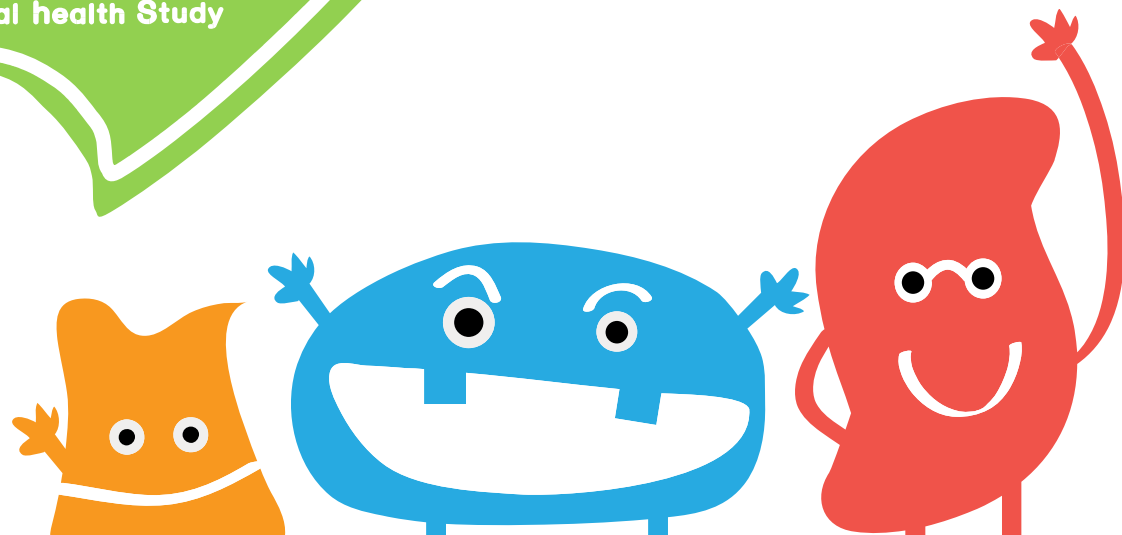
# Korean Children's ENVIRONMENTAL health Study

Hello!!

We are **Koby, Chenchen, and Stu!**

Let us introduce our study

**Ko-CHENS!**  
Korean Children's  
ENVIRONMENTAL health Study





## An introduction to



- Korean CHildren's ENvironmental health Study (Ko-CHENS) is a long-term prospective pregnancy and children cohort study which is planning to include approximately 100,000 pregnancies in the years between 2015 and 2019.
- The study includes the prenatal period up to an age of 18 years and follow-up activities for monitored children into adulthood using biological samples and questionnaires focused on environmental exposures.



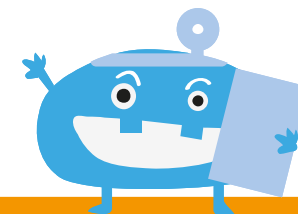
MINISTRY OF  
ENVIRONMENT



National Institute of  
Environmental Research



## Study Design



- Ko-CHENS will use two data collection methods; Main study and Core study.

|                      | Main study                                                           | Core study                                                       |
|----------------------|----------------------------------------------------------------------|------------------------------------------------------------------|
| Study period         | 2015~2036<br>(22years, from the prenatal period until the age of 18) |                                                                  |
| Recruitment period   | 2015~2019 (5years)                                                   | 2015~2017 (3years)                                               |
| Target participants  | 65,000 pregnant women                                                | 5,000 pregnant women                                             |
| Recruiting Institute | Public health care centers<br>/Obstetric units in the hospitals      | Obstetric units in the hospitals<br>/ Public health care centers |
| Follow-up period     | Through linkage to national health registries                        | 20 years through hospital visits                                 |

# Join *Ko-CHENS!* NOW!

Participant  
requirement



All pregnant women (Main study) /  
Pregnant women at less than 20  
weeks of gestation (Core study)

Way to  
Register



You can join Ko-CHENS at  
designated hospitals and centers

Questions  
about  
registration

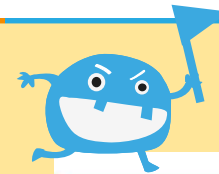


Information of designated centers,  
website, hotline of  
Ko-CHENS are at the bottom

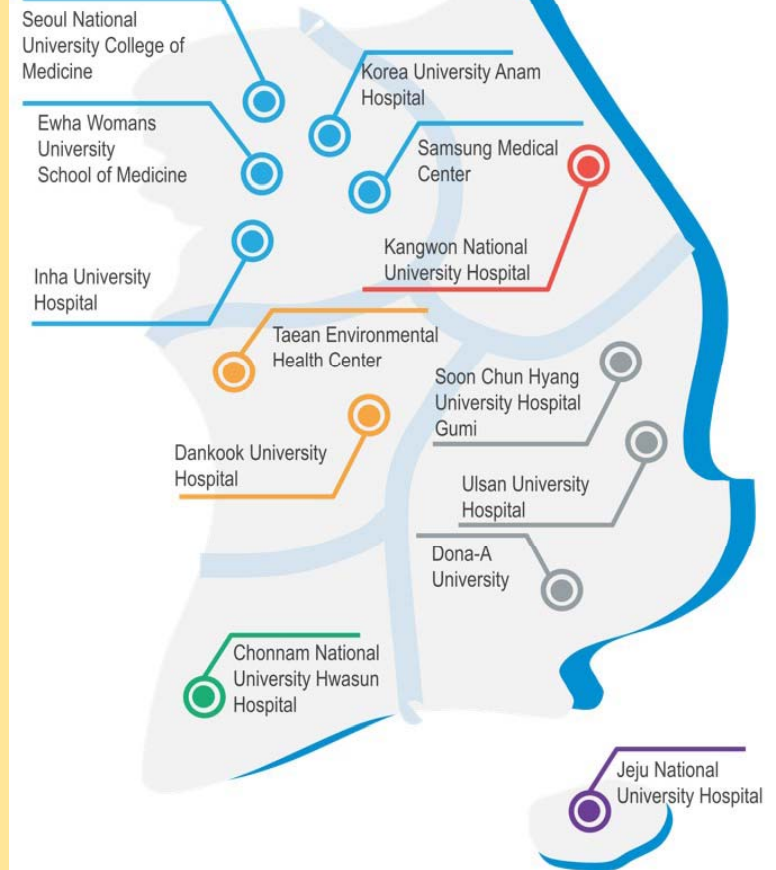
Official website : [cafe.naver.com/environmentforchild](http://cafe.naver.com/environmentforchild)

Mobile ver. Website : [environmentforchild.modoo.at](http://environmentforchild.modoo.at)

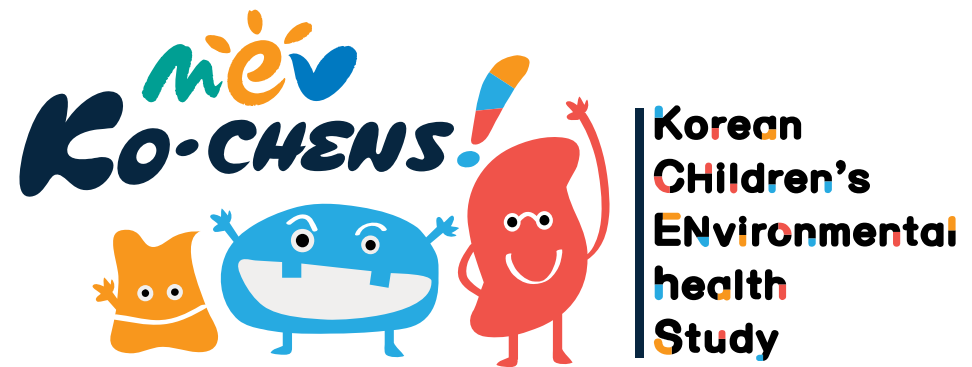
Official hotline : 1544-6701



## Map of Study Locations

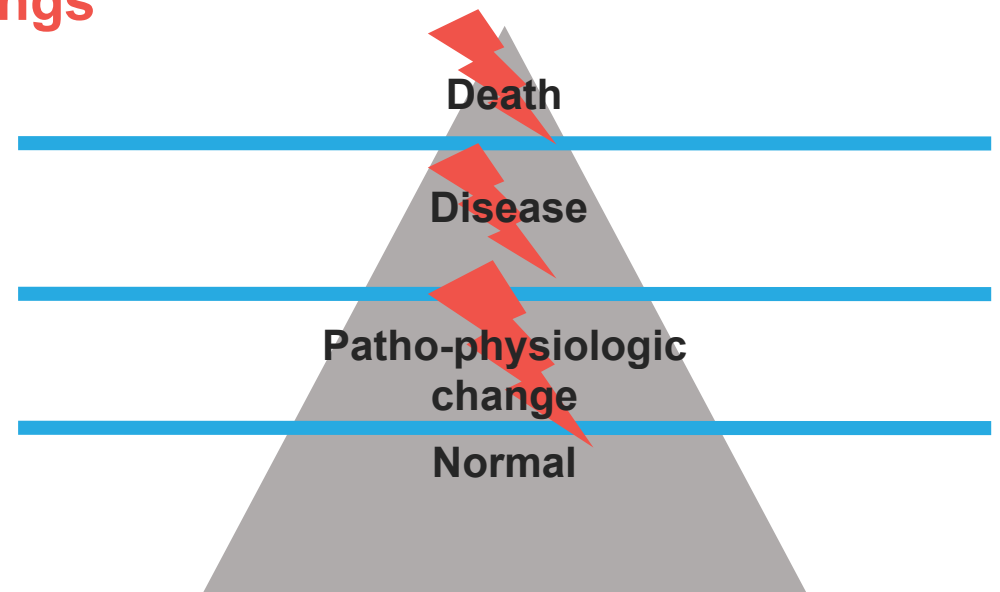


## V. Conclusion



# 1. Is it really that ambient air pollution is a risky?

- “YES”
- “Consistent” and “coherent” findings
- “Consistent findings” over
  - areas/cities/countries
  - populations
  - study designs





## **2. Who are in danger?**

- **Children and the elderly**
- **Persons having a chronic disease**

## **3. What else do we need to pursue?**

- **Physico-chemical characteristics of particles**
  - **Accountability**
  - **Reduction strategies of adverse health effects from air pollution exposure**
- 

- What is “Precautionary Principle”?

*“We must act on facts, and on the most accurate interpretation of them, using the best scientific information. That does not mean that we must sit back until we have 100% evidence about everything. Where the state of the health of the people is at stake, the risks can be so high and the costs of corrective action so great, that prevention is better than cure. We must analyze the possible benefits and costs of action and inaction. **Where there are significant risks of damage to the public health, we should be prepared to take action to diminish those risks, even when the scientific knowledge is not conclusive**, if the balance of likely costs and benefits justifies it.”*

(Horton R. The new public health of risk and radical engagement. Lancet 1998;352:251-2)



Thank you

