

Air pollution and human health

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Disclosures

- Grant funding: NHLBI, Department of Veteran's Affairs
- Consultant, Partner Therapeutics, unrelated to the content of this presentation



Some questions

- Why do we care about air pollution?
 - How does air pollution affect health?
 - Does changing pollution levels lead to clinically important changes in human health?
- What do we know about biological responses to pollution? What organ systems are involved?
- What kinds of air pollution are important in Utah?
 - Where does it come from?
 - What are we doing about it?
- Relevance to COVID-19?

Photo: S. Hoch (Atmos. Sci)

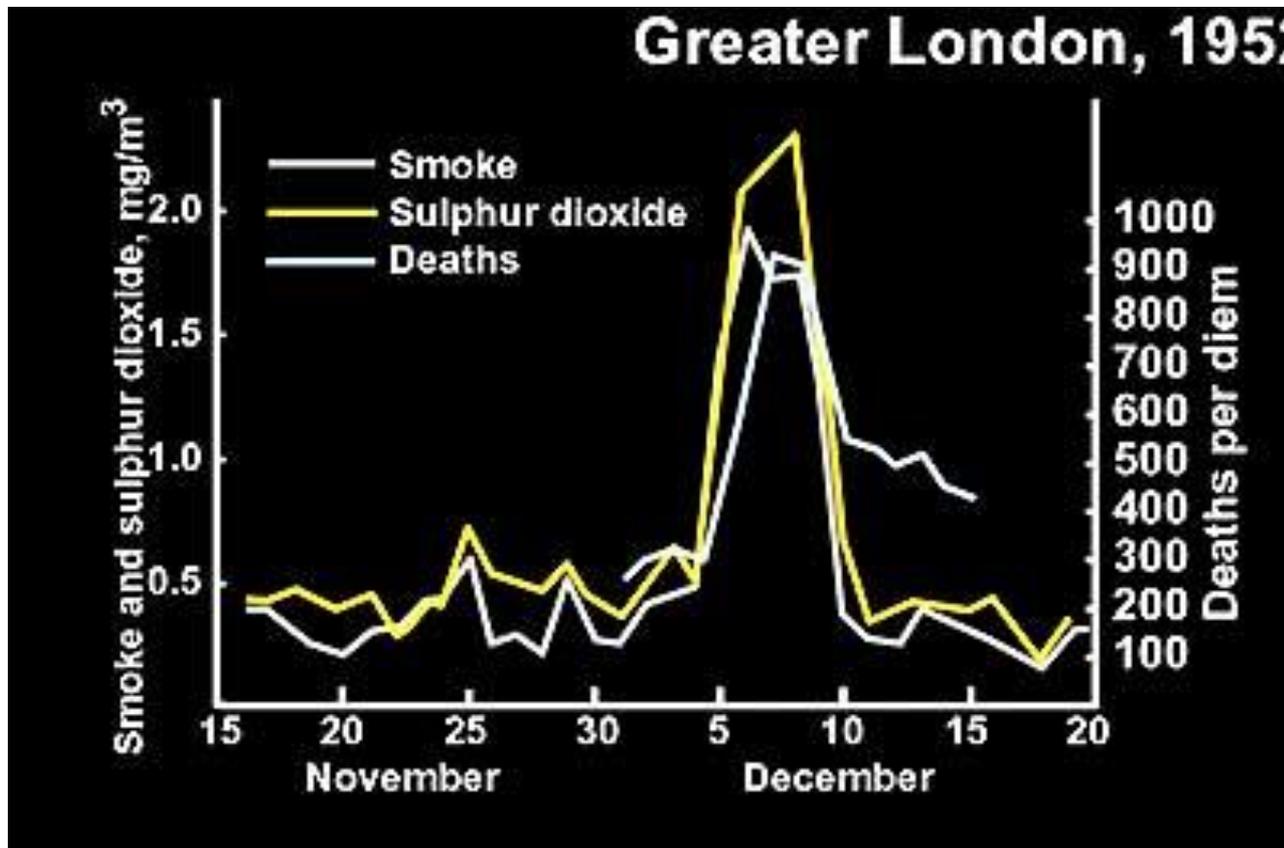


London, December 1952



London

December 5-9, 1952

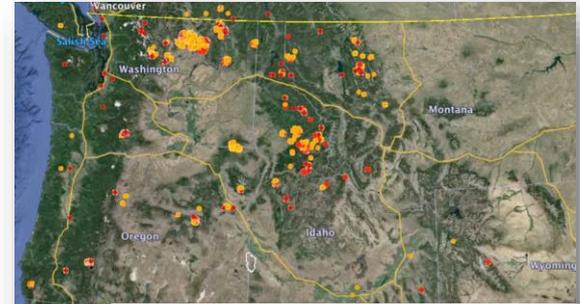


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Worldwide causes of poor air quality

- Transportation
- Solid-fuel burning
- Power generation
- Municipal waste management
- Waste incineration
- Agricultural burning and wildfires
- Meteorology and topography



EPA defined “Criteria” pollutants

- Particulate matter
 - PM_{2.5}
- Ozone
- Lead
- Carbon monoxide
- Sulfur oxides
- Nitrogen dioxide
- Air toxics



PM_{2.5} - The Key Aerodynamic Diameter to Reach the Gas Exchange Portion of The Lung

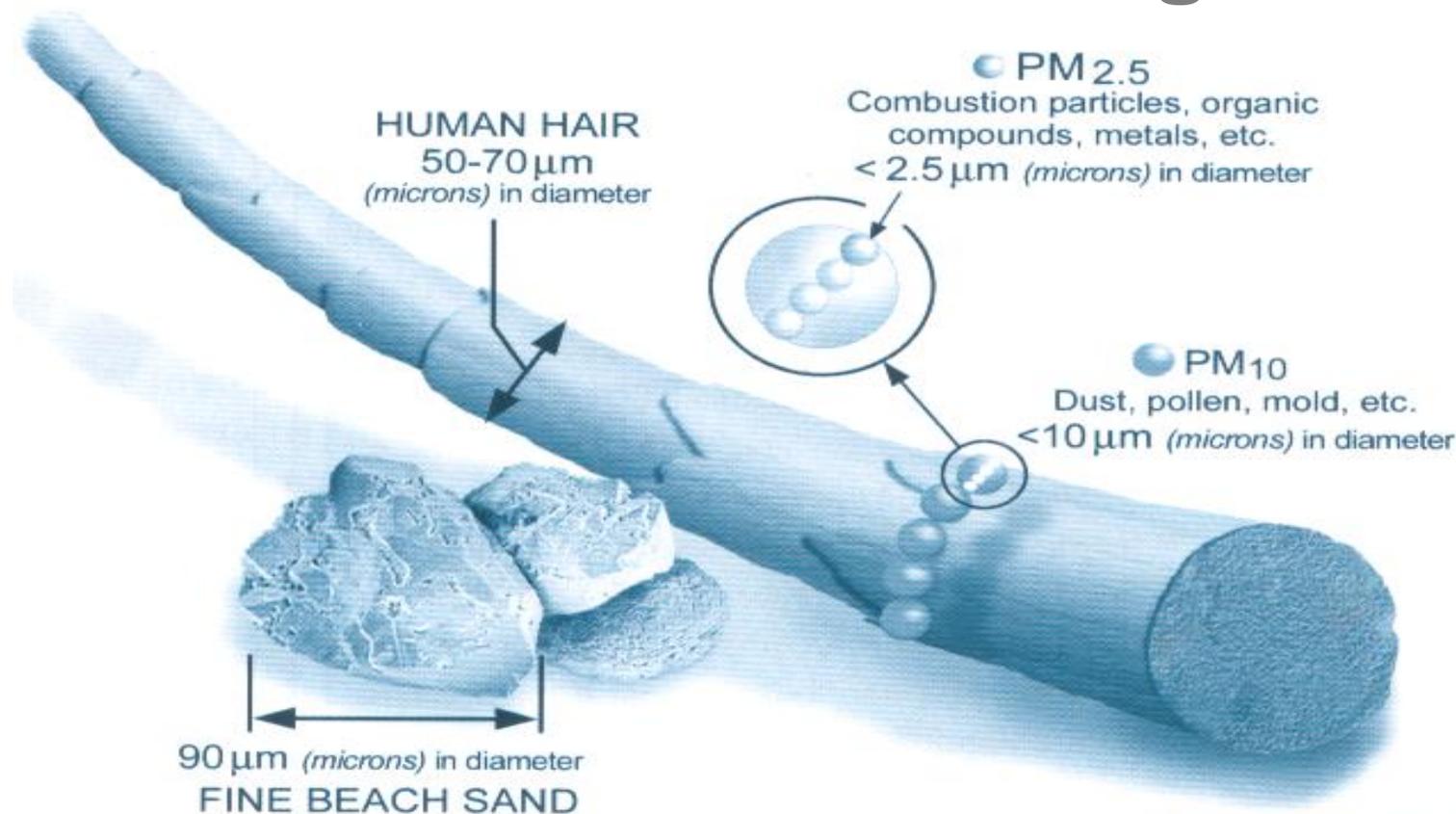


Image courtesy of the U.S. EPA

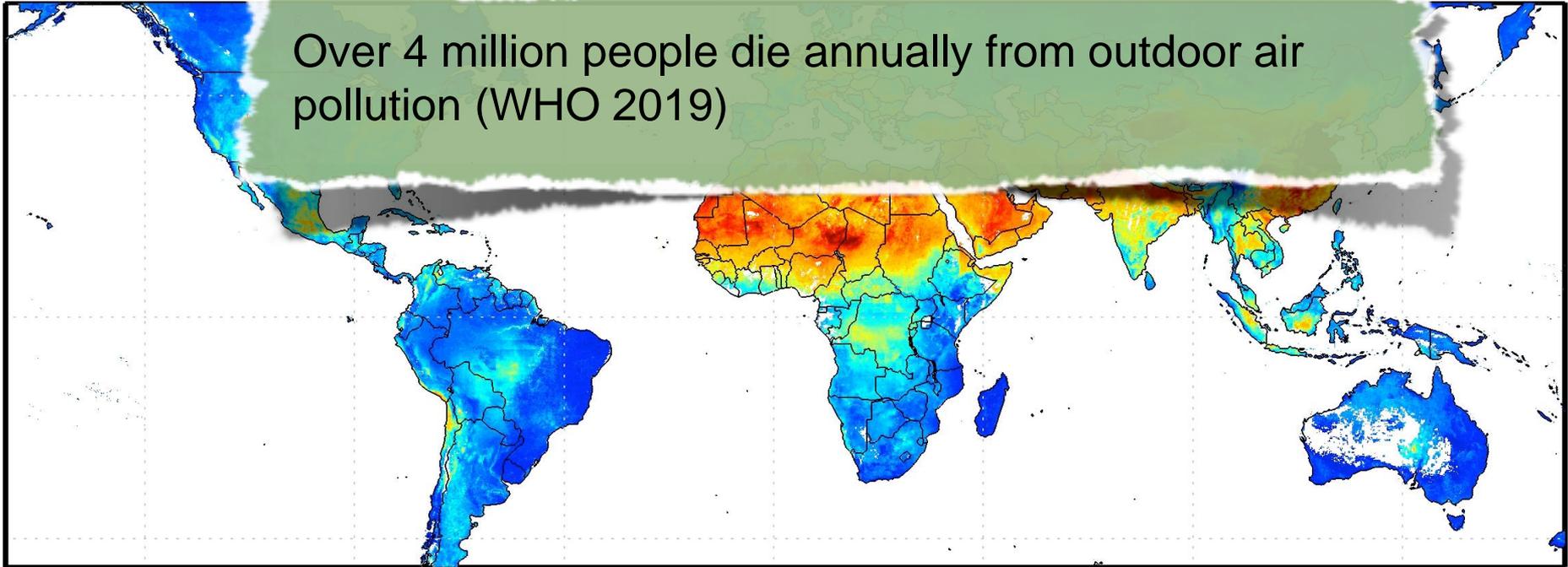


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Global annual PM_{2.5} levels

Over 4 million people die annually from outdoor air pollution (WHO 2019)



Satellite-Derived PM_{2.5} [$\mu\text{g}/\text{m}^3$]

Nasa.gov



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South Africa



Guatemala



India



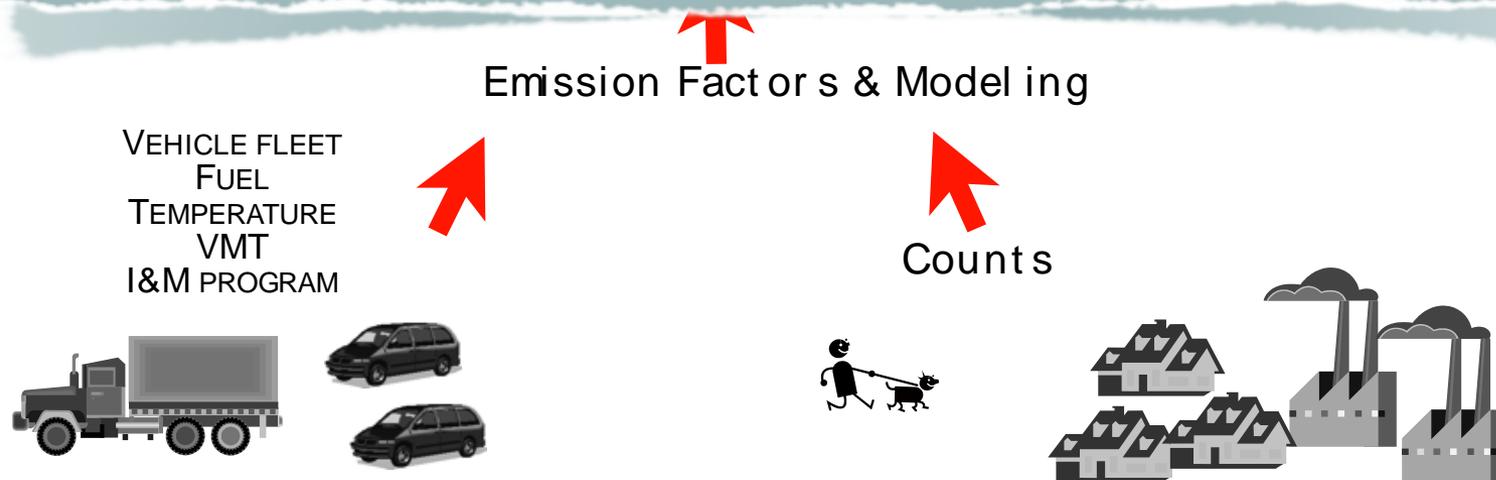
What are our Sources of Pollution?

Mobile sources responsible for 54% of emissions and are decreasing

Areas sources are responsible for about 33% of emissions and are increasing

Point sources are responsible for about 12% of emissions and are increasing

Data for Salt Lake County from DAQ SIP website, September 16, 2018

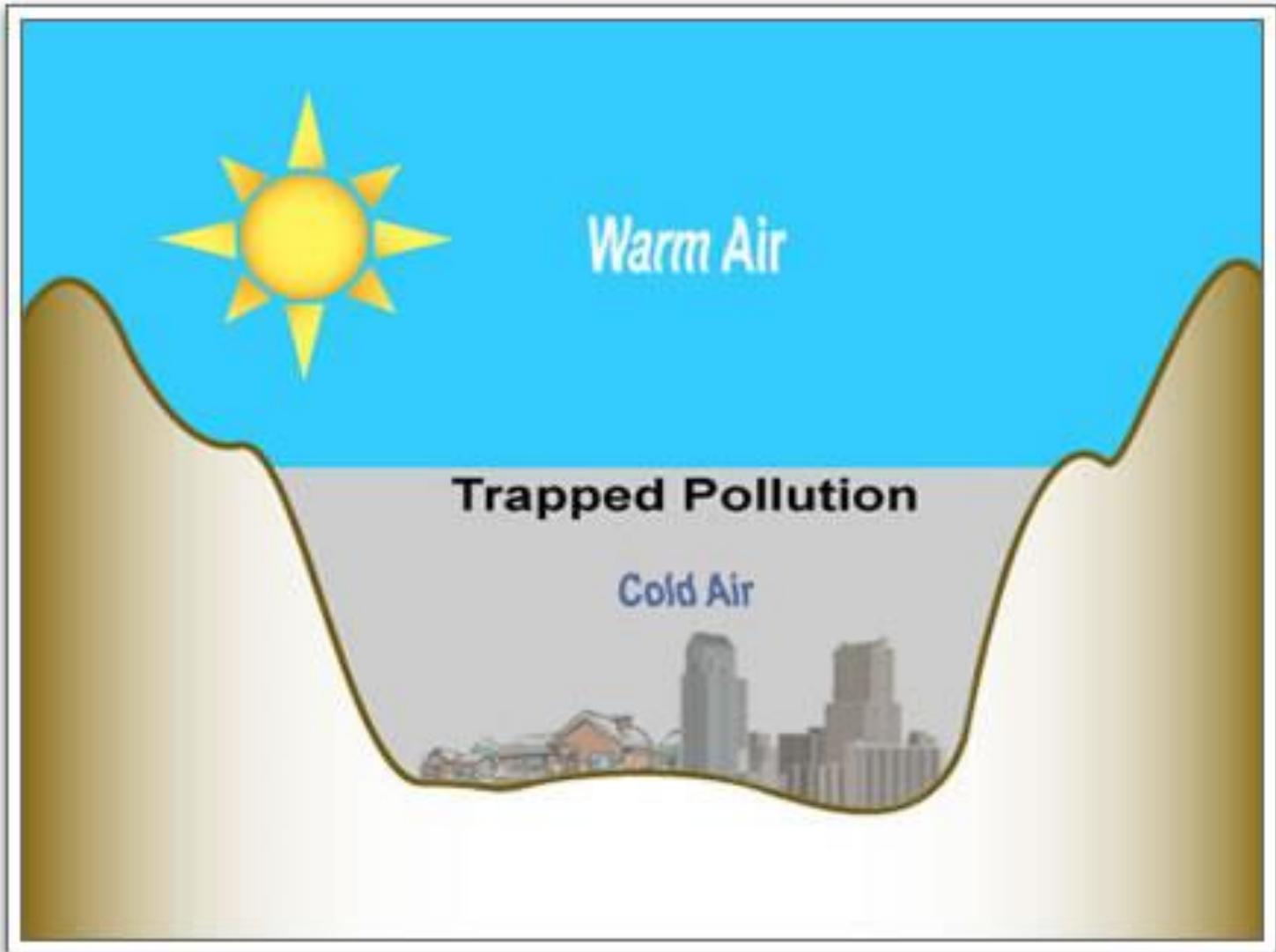


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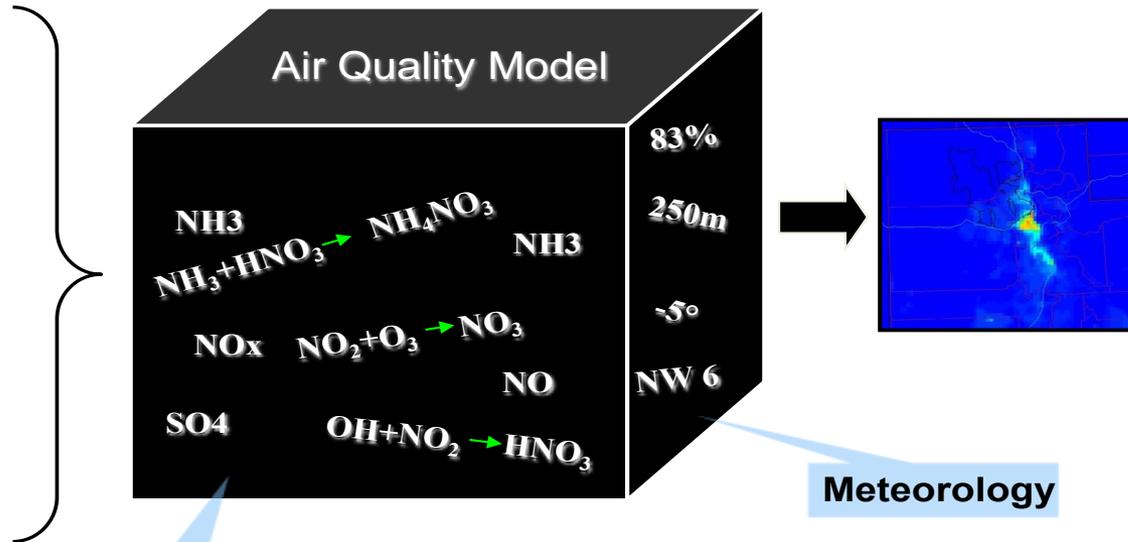


Wintertime inversions



Where does the PM come from?

Emissions



Chemistry

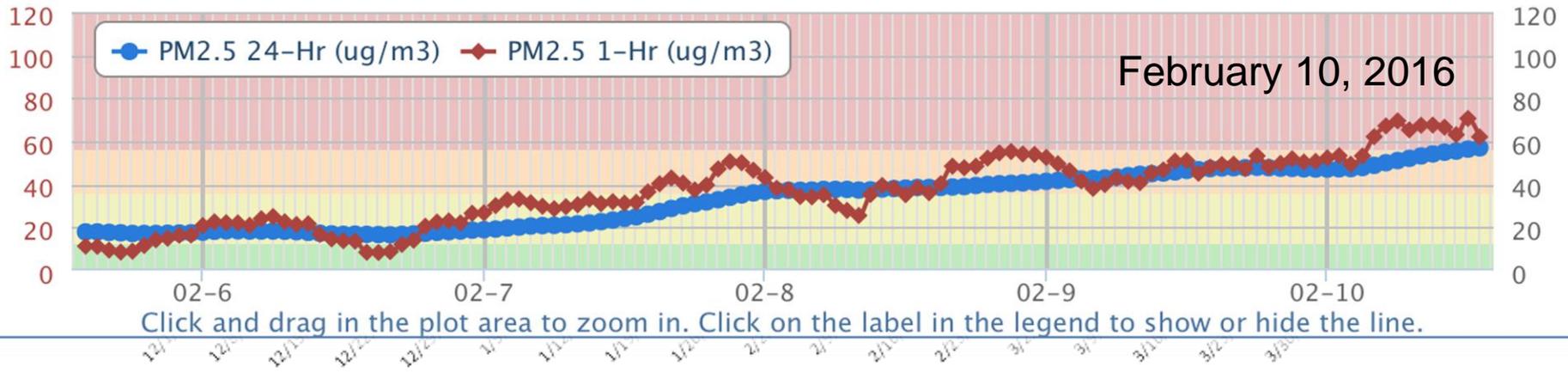
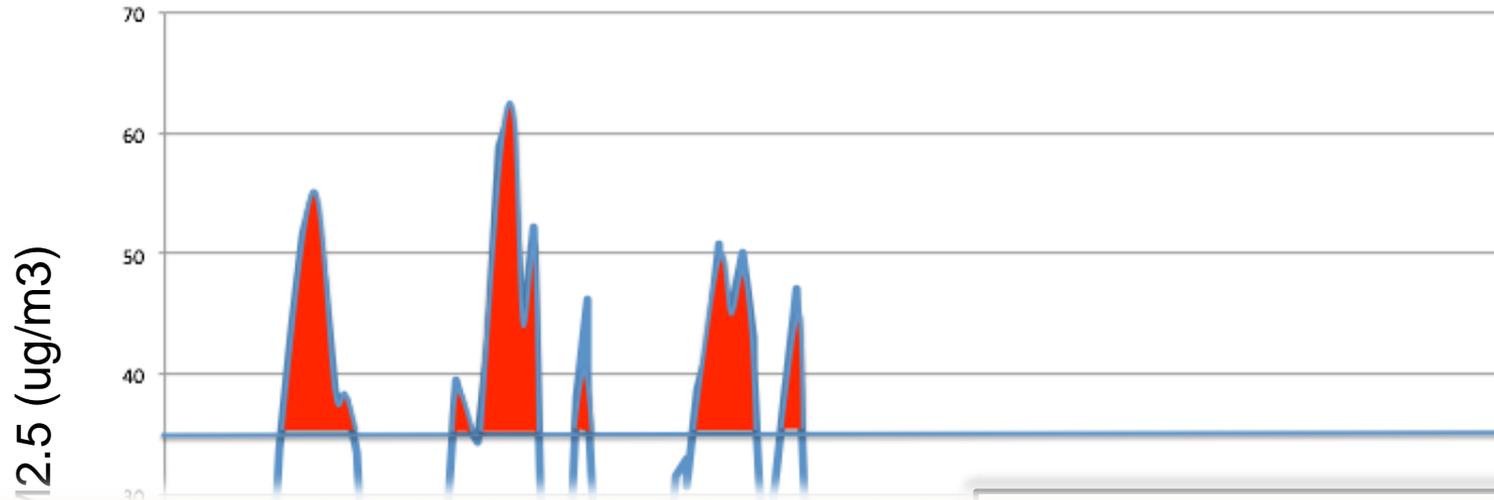
Meteorology

Directly emitted particles 1/3

Indirect reactions 2/3

We contribute to both direct & indirect PM

Wintertime PM_{2.5} levels



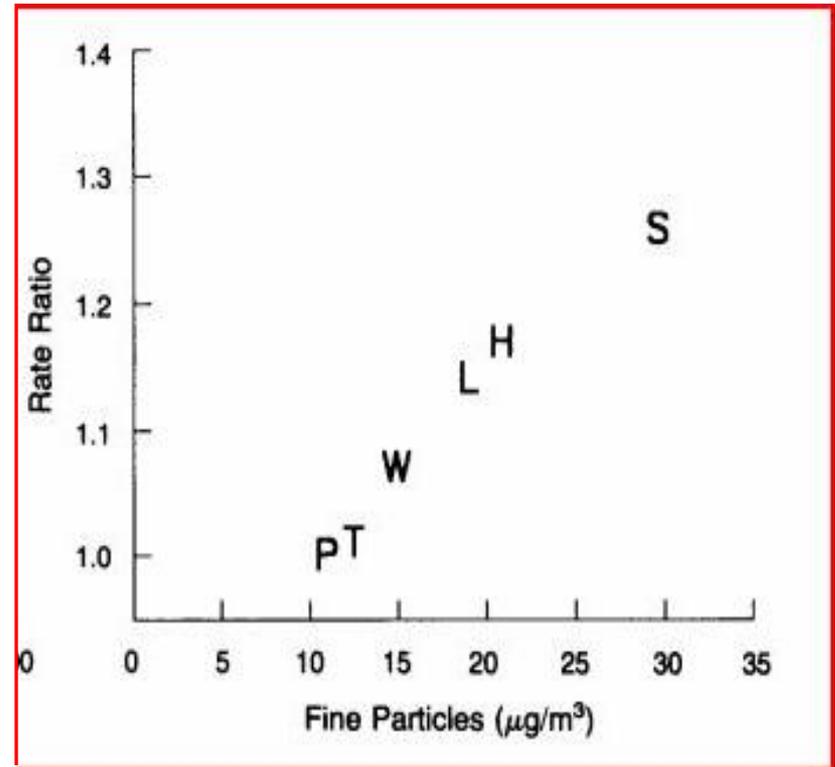
Epidemiologic data concerning the impact of air pollution

- Extensive epidemiologic evidence linking exposure to air pollution to effects on human health
 - Total mortality/life expectancy
 - The Global Burden of Disease study estimated that pollution caused 9 million deaths in 2019, of which more than 60% were due to cardiovascular disease
 - Cardiovascular events (MI and ACS, strokes, sudden death)
 - Asthma
 - Worsening of symptoms for patients with COPD
 - Cancer



Correlation of mortality with air quality - Harvard Six Cities Study

- Prospective study with enrollment in 1979 and follow-up through 1990's
- City specific mortality during early ('80's) and late ('90's) periods and over entire study
- Mortality inversely related to level of fine particulates during each period
- Similar correlations both for overall mean exposure and for exposure in the year prior to death



S- Steubenville, H- Harriman, T- Topeka,
W- Watertown, L- St Louis, P- Portage



Air pollution impacts disease in multiple organ systems

Respiratory disease mortality

Respiratory disease morbidity

Lung cancer

Pneumonia

Upper and lower respiratory symptoms

Airway inflammation

Decreased lung function

Decreased lung growth

Insulin resistance

Type 2 diabetes

Type 1 diabetes

Bone metabolism

High blood pressure

Endothelial dysfunction

Increased blood coagulation

Systemic inflammation

Deep venous thrombosis

Stroke

Neurological development

Mental health

Neurodegenerative diseases

Cardiovascular disease mortality

Cardiovascular disease morbidity

Myocardial infarction

Arrhythmia

Congestive heart failure

Changes in heart rate variability

ST-segment depression

Skin ageing

Premature birth

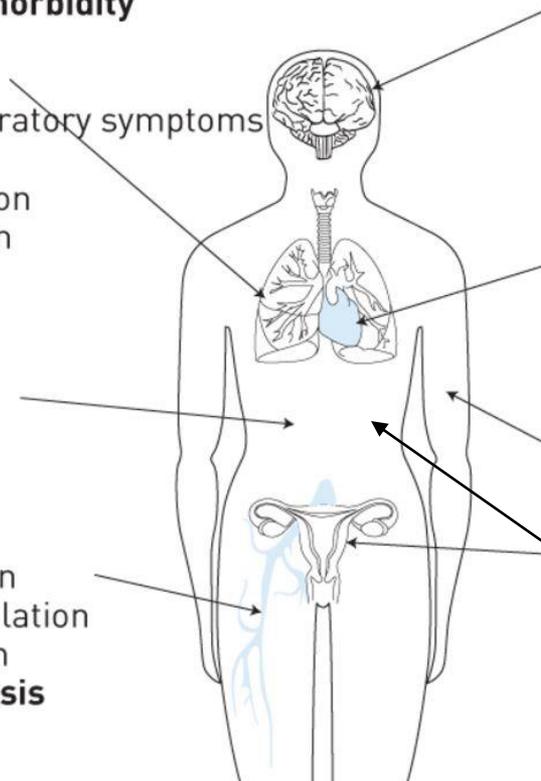
Decreased birthweight

Decreased fetal growth

Intrauterine growth retardation

Decreased sperm quality

Pre-eclampsia



Chronic kidney disease



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Thurston et al. Eur Resp J 2017

Long and short term effects on cardiovascular disease

- Long term:
 - Large national cohort studies demonstrate direct relationships being PM2.5 local exposure and both total mortality and cardiovascular mortality
 - CVD risk increases 10% for each 10 $\mu\text{g}/\text{M}^3$ increase in PM2.5
 - Thurston et al. Environmental Health Perspectives 2016
- Short term:
 - Local studies from Utah have demonstrated increased risk of heart attacks during 1-3 days after onset of increased PM2.5 pollution
 - 4% increased rate for each 10 $\mu\text{g}/\text{M}^3$ increase in PM2.5
 - Pope et al. Circulation 2006



Oxford Street



PM_{2.5} = 28.3

Assessed pollutant exposure,
changes in pulmonary function;
airway inflammation

Randomized cross over trial
60 adults with mild to moderate asthma
taking two leisurely 2-hr walks in London

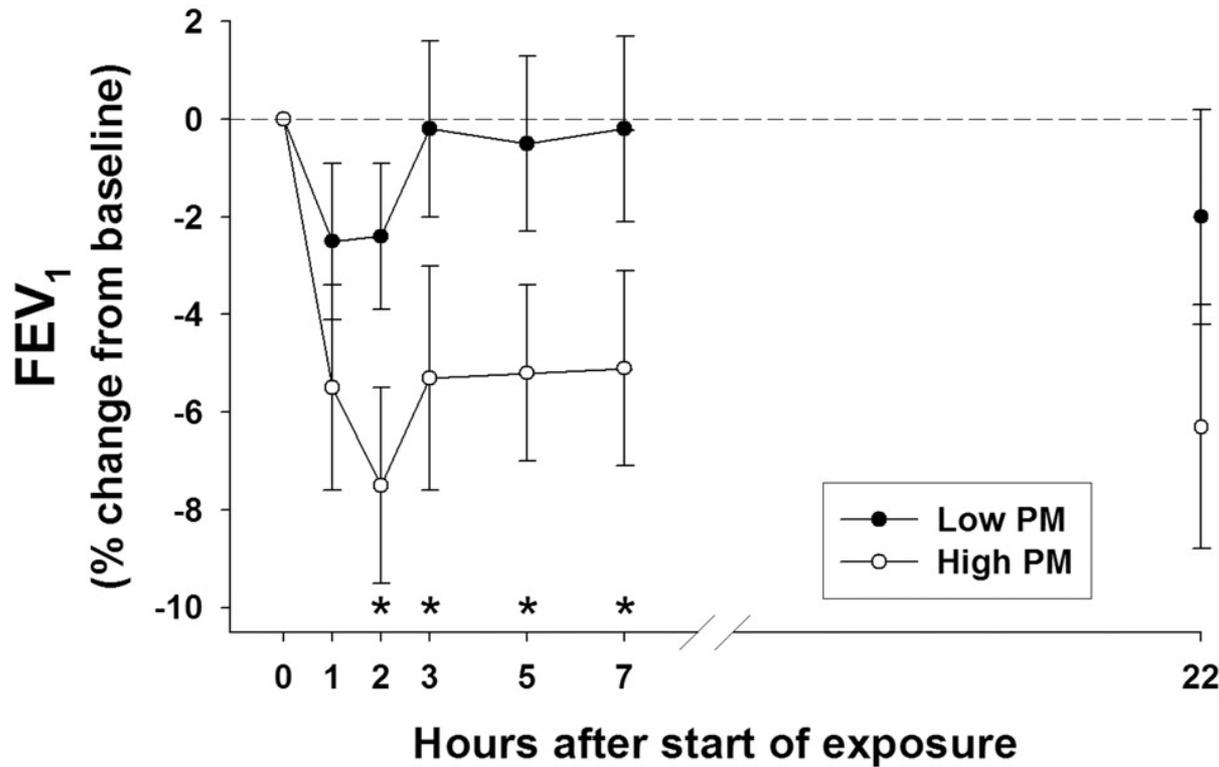


Hyde Park

PM_{2.5} = 11.9



Changes in pulmonary function during and after Exposure on Oxford Street and in Hyde Park



- Key correlations were with PM_{2.5}, ultrafine particles, elemental carbon and nitrogen dioxide
- Significant increases in markers of airway inflammation

McCreanor et al.
NEJM 2007; 357:2348



Impact of air pollution on cognitive function

- Nurses' Health Study
 - 19,000 women aged 70-81
 - Followed over time with estimates of PM_{2.5} and coarse particle exposure
 - Decline in cognitive function on sophisticated testing associated with increased pollution
 - An increment of 10 $\mu\text{g}/\text{m}^3$ long term PM exposure was cognitively equivalent to aging by approximately 2 years



Effects of Air Pollution on Children

- Incidence of asthma
- Acute asthma exacerbations
- Lung growth
- Lost IQ points

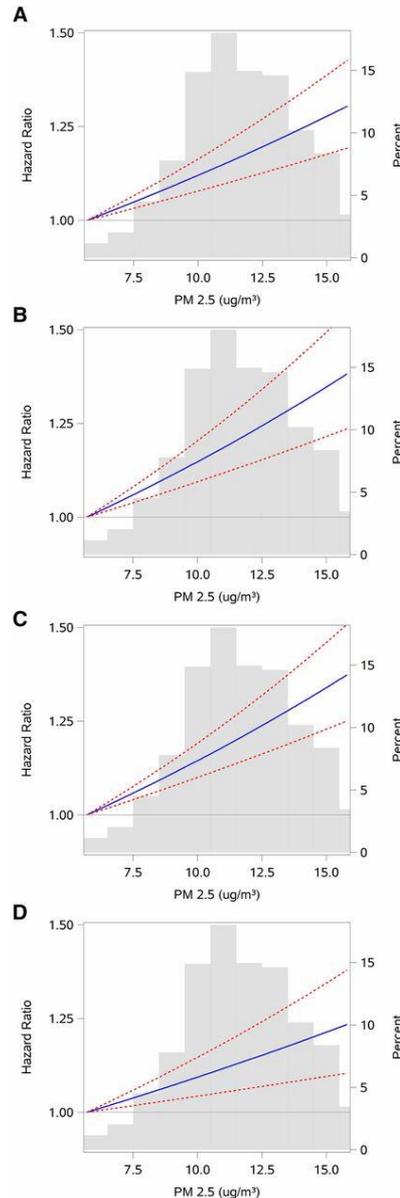
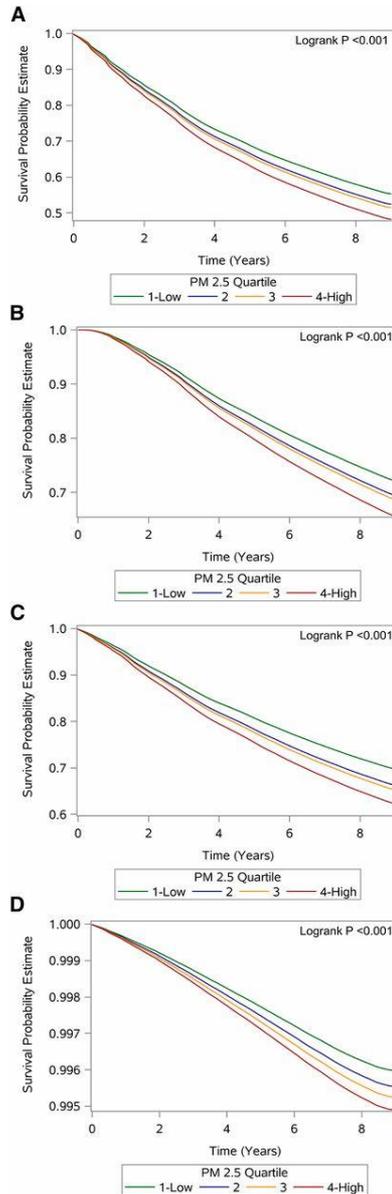


Is there a risk of CKD associated with increased exposure to PM2.5?

- PM2.5 exposure is associated with
 - Hypertension
 - Endothelial cell dysfunction
 - Oxidative stress
- Bowe et al. JASN 2018
 - Observational cohort of 2482737 Veterans followed 8.52 years
 - Ground level PM2.5 monitoring and NASA satellite data



Analyses of risk of renal outcomes by PM2.5 concentrations (PM2.5 of 5.7 $\mu\text{g}/\text{m}^3$ served as a reference) with PM2.5 probability distribution in the background.



Risks:

eGFR < 60 ml/min/1.73 M²

Incident CKD

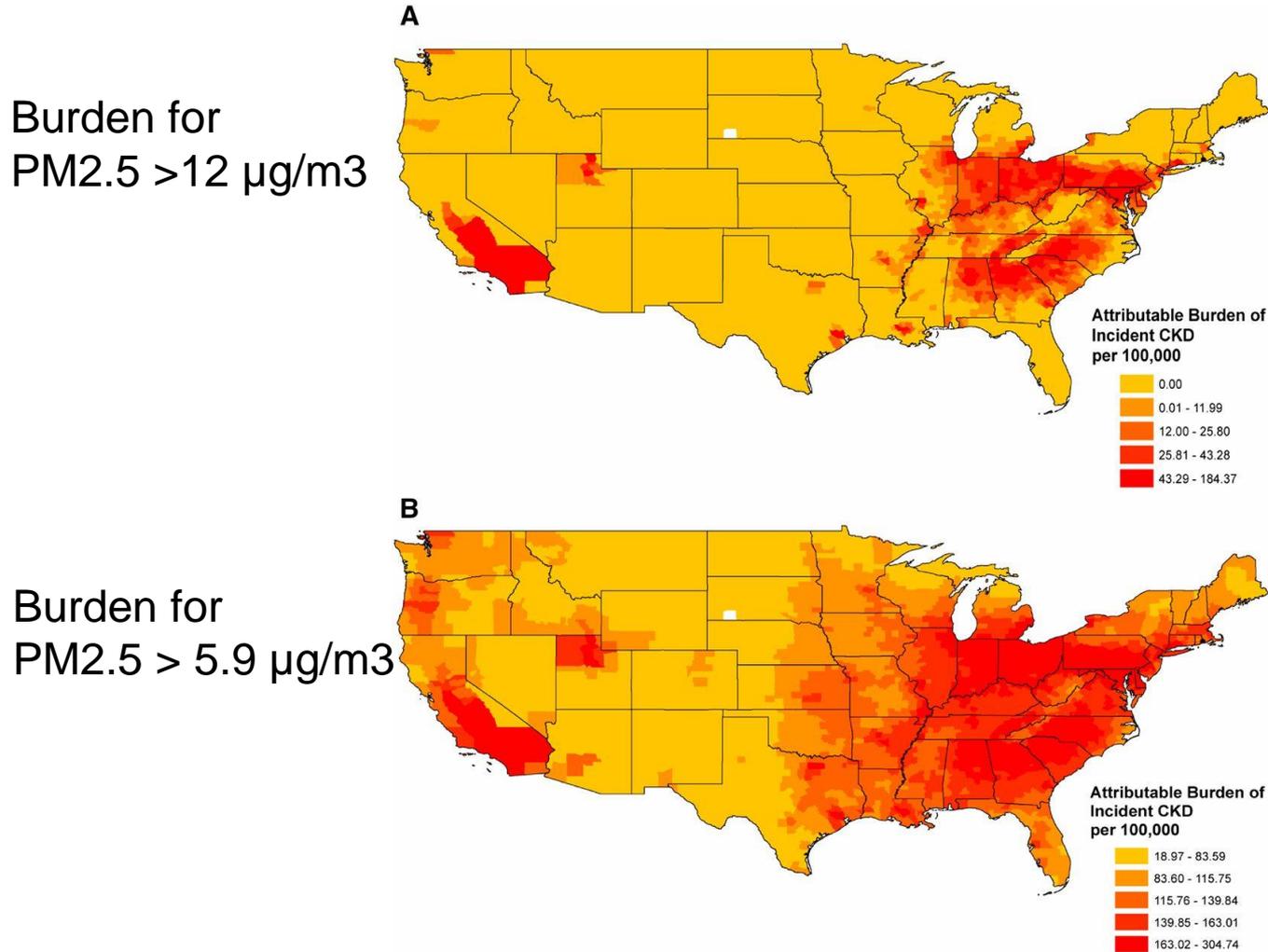
eGFR decline > 30%

ESRD

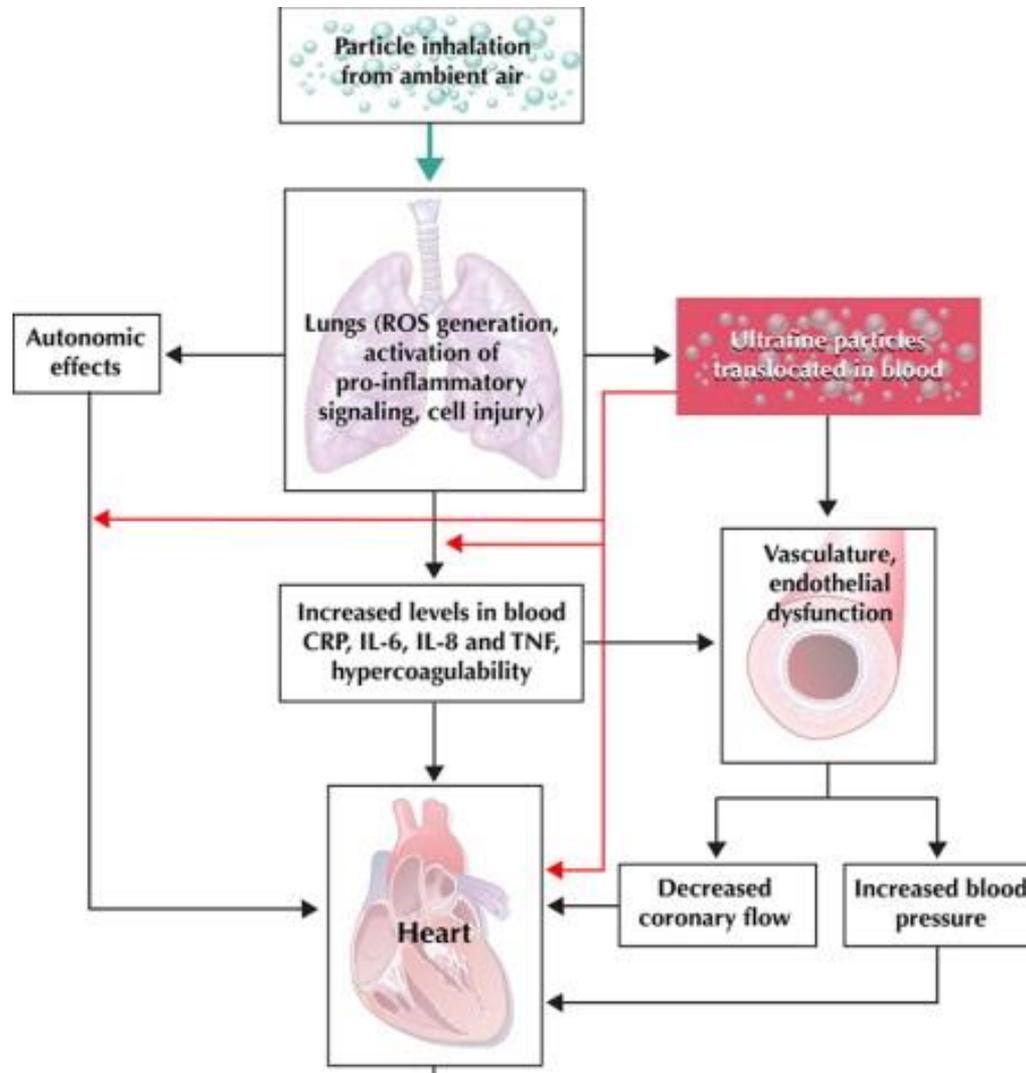
Benjamin Bove et al.
JASN 2018;29:218-230



Geographic distribution of the national burden of incident CKD attributable to air pollution in the United States.



Benjamin Bowe et al. JASN 2018;29:218-230



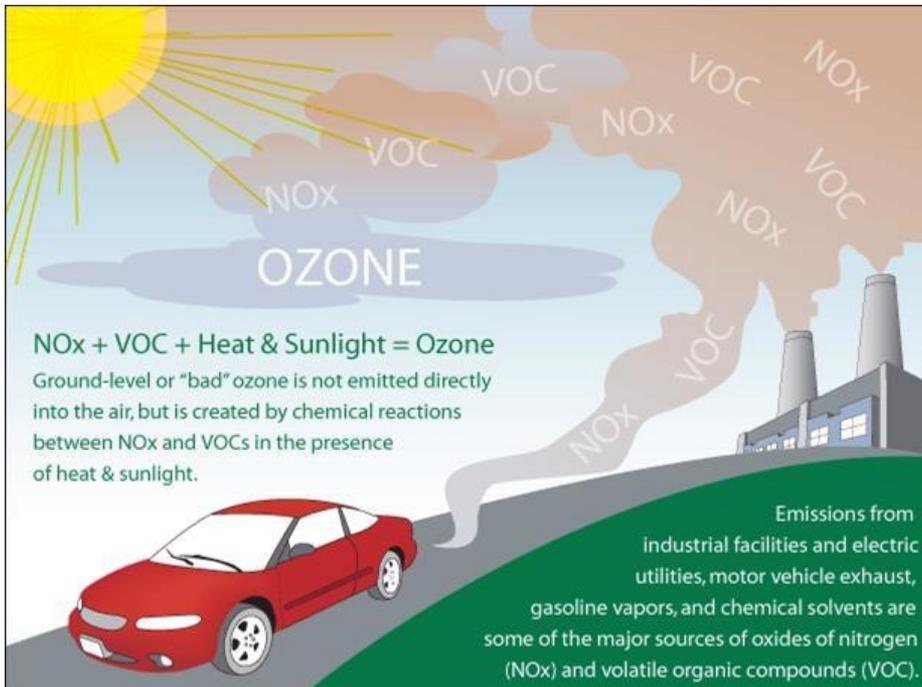
Simkhovich et al. J Am Coll Cardiol 2008



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Ground level ozone



EPA 8 hr standard < 70 ppb

Summertime ozone is a challenge along the Wasatch Front

Wintertime ozone is a challenge in the Uinta Basin



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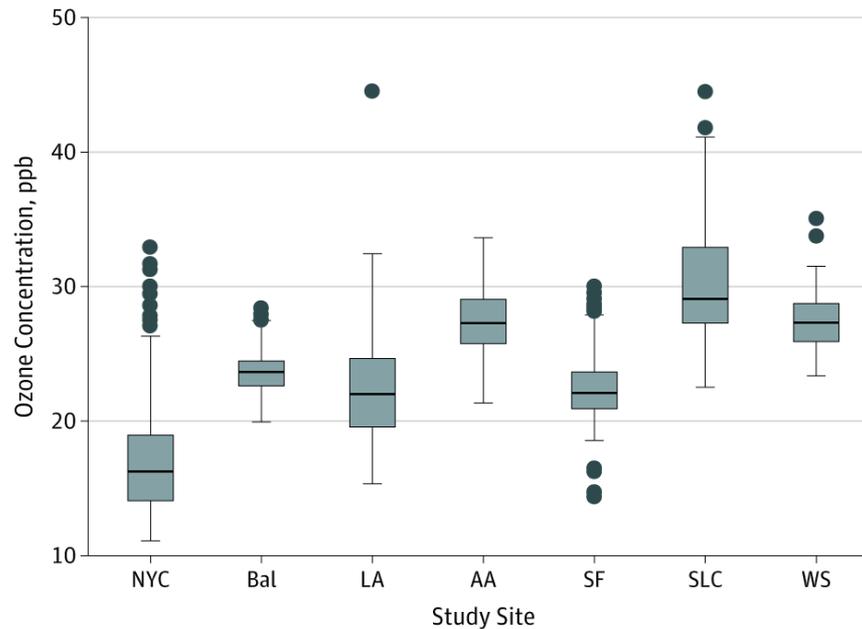
Short-term ozone exposure

- EPA threshold of 8-hr level of 70 ppb
- Association with
 - Acute exacerbations of asthma
 - Acute exacerbations of COPD
- Less information about longer term effects of ozone
 - No annual threshold designated by EPA



From: **Association of Long-term Ambient Ozone Exposure With Respiratory Morbidity in Smokers**

JAMA Intern Med. 2020;180(1):106-115. doi:10.1001/jamainternmed.2019.5498



Distribution of 10-Year Historical Ozone Concentration by Study Site

Date of download: 10/20/2021

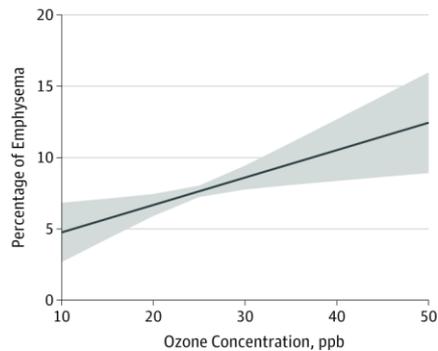
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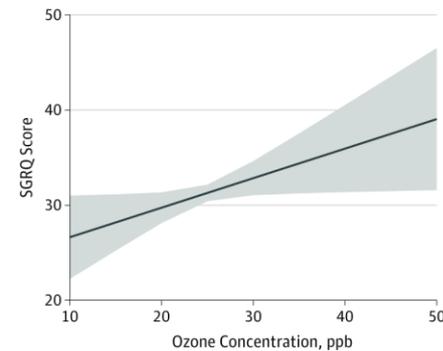
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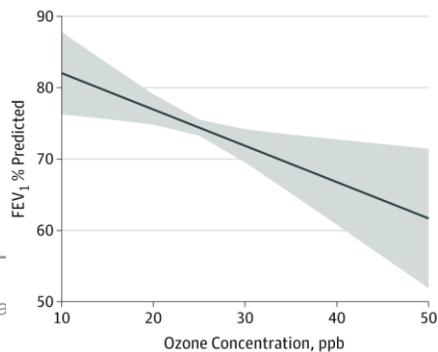
A Percentage of emphysema



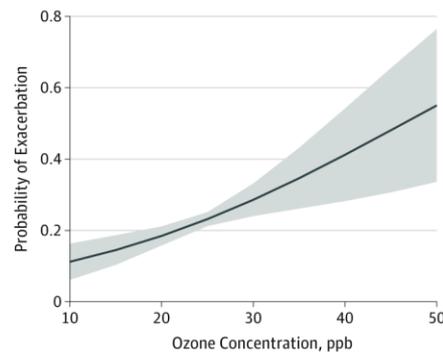
B Total SGRQ score



C FEV₁ % predicted



D Probability of exacerbation 12 mo before enrollment

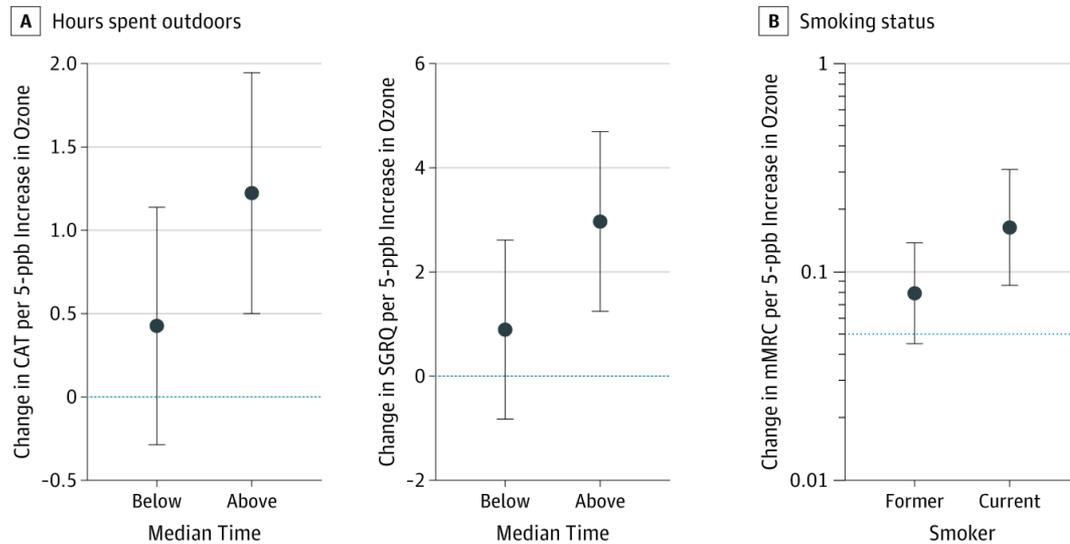


Adjusted Associations of 10-Year Historical Ozone Concentration With Selected Outcomes of Interest: FEV₁ indicates forced expiratory volume in the first second of expiration; ppb, parts per billion; SGRQ, St. George Respiratory Questionnaire. Shaded areas represent 95% CIs.



From: **Association of Long-term Ambient Ozone Exposure With Respiratory Morbidity in Smokers**

JAMA Intern Med. 2020;180(1):106-115. doi:10.1001/jamainternmed.2019.5498



Effect Estimate of Ozone on Outcomes by Time Spent Outdoors and Smoking Status. CAT indicates COPD (chronic obstructive pulmonary disease) Assessment Test; mMRC, modified Medical Research Council Dyspnea Scale; ppb, parts per billion; SGRQ, St. George Respiratory Questionnaire. Error bars represent 95% CIs. P for interaction <.05 for analyses.

Date of download: 10/20/2021

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Air Pollution and COVID-19

- The lockdown in response to the pandemic has led to improved air quality, best seen in China and Europe
- Air pollution exposure is associated with increased risk of, and poor outcomes from, pneumonia
 - Dysregulated inflammation
 - Oxidative stress
- Prior to COVID, air pollution exposure is associated with increased risk of acute respiratory distress syndrome from viral infections, trauma



| Study | Country | Exposure | COVID-19 Outcome | Principal Findings |
|---|-----------------|--|--|--|
| Wu <i>et al.</i> (April 2020) | United States | PM _{2.5} Average between 2000 and 2016 | Mortality rate | Increase of 1 µg/m ³ in PM _{2.5} associated with an 8% increase in mortality rate |
| Cole <i>et al.</i> (June 2020) | The Netherlands | PM _{2.5} , NO ₂ , and SO ₂ Average between 2015 and 2019 | Cases Hospital admissions Deaths | 1 µg/m ³ increase in PM _{2.5} associated with 9.4 more cases, 3.0 more hospital admissions, and 2.3 more deaths NO ₂ (but not SO ₂) associated with cases and deaths |
| Hendryx <i>et al.</i> (June 2020) | United States | PM _{2.5} [*] , diesel PM [†] and O ₃ [*] | Cases Deaths | PM _{2.5} and diesel PM associated with higher prevalence and mortality Diesel PM appeared to be the primary driver for associations with PM _{2.5} |
| Travaglio <i>et al.</i> (June 2020) | England | PM _{2.5} , PM ₁₀ , SO ₂ , NO ₂ , NO, and O ₃ Average between 2018 and 2019 | Cases Deaths Infectivity rate | NO and NO ₂ significant predictors of cases independent of population density 1 µg/m ³ increase in SO ₂ and NO associated with 17% and 2% higher mortality, respectively O ₃ negatively associated with cases and deaths PM _{2.5} , PM ₁₀ , and SO ₂ associated with SARS-CoV-2 infectivity (OR, 1.12, 1.07, and 1.32, respectively) but not cases or deaths |
| Konstantinoudis <i>et al.</i> (August 2020) | England | PM _{2.5} and NO ₂ Average between 2014 and 2018 | Mortality rate | 1 µg/m ³ increase in PM _{2.5} and NO ₂ associated with 1.4% and 0.5% higher mortality rate, respectively |
| Liang <i>et al.</i> (October 2020) | United States | PM _{2.5} , NO ₂ , and O ₃ Annual mean between 2010 and 2016 | CFR Mortality rate | NO ₂ associated with 11.3% and 16.2% higher CFR and mortality rate, respectively, per IQR PM _{2.5} associated with 14.9% higher mortality rate per IQR, but not with CFR No association between O ₃ and CFR or mortality rate |



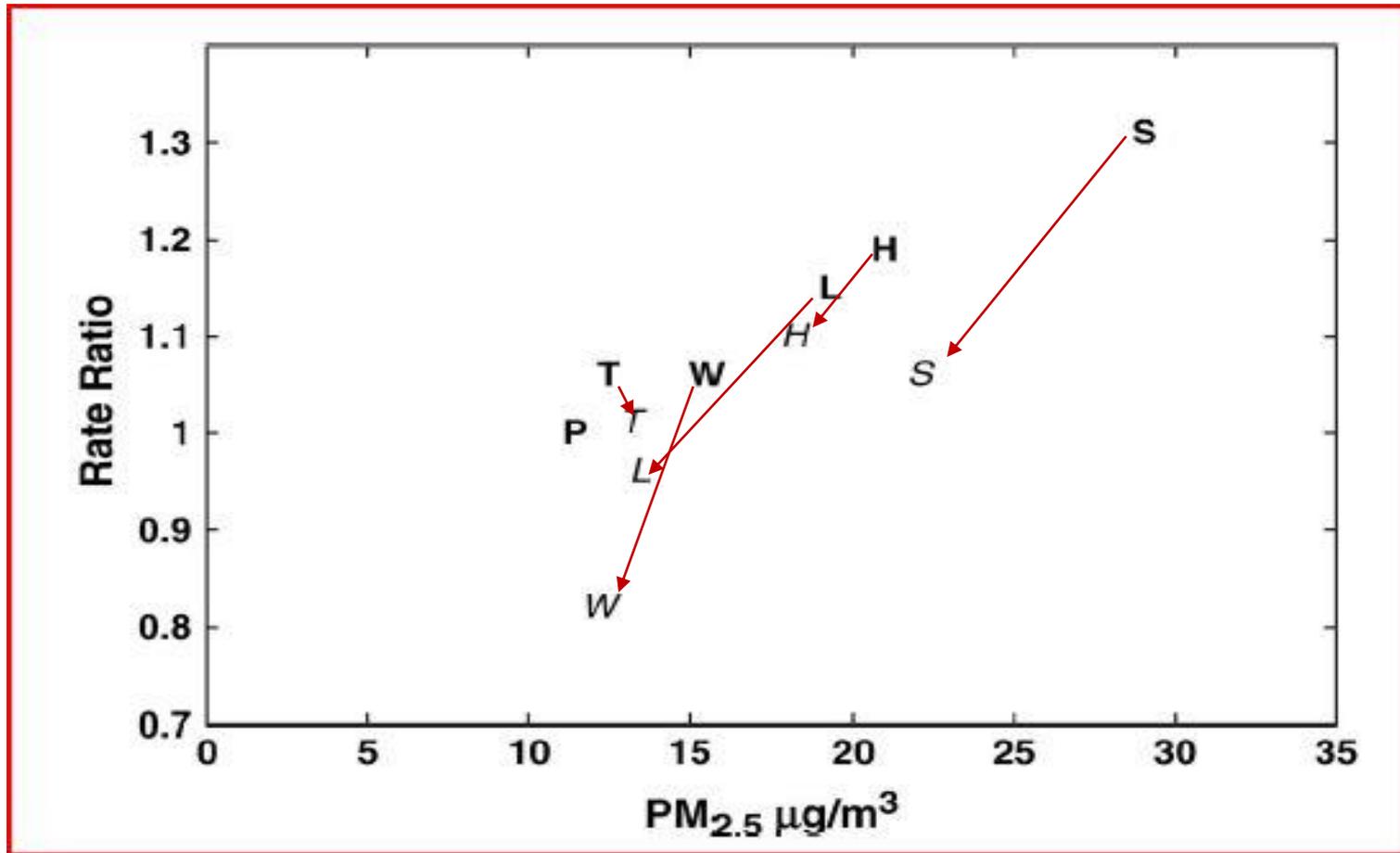
Do efforts to improve air quality make a difference for health?



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Reduction in Fine Particulate Air Pollution and Mortality: Extended Follow-up of the Harvard Six Cities Study



Fine particulate air pollution and life expectancy in the US

- Detailed monitoring information at the county level
- Looking at late 1970's and early 1980's and at late 1990's and early 2000's
- Cross-sectional comparisons within each period
- Proxies for smoking, access to health care, etc.
- Temporal comparisons between periods each county as its own control

Pope et al. NEJM 2009; 360: 376



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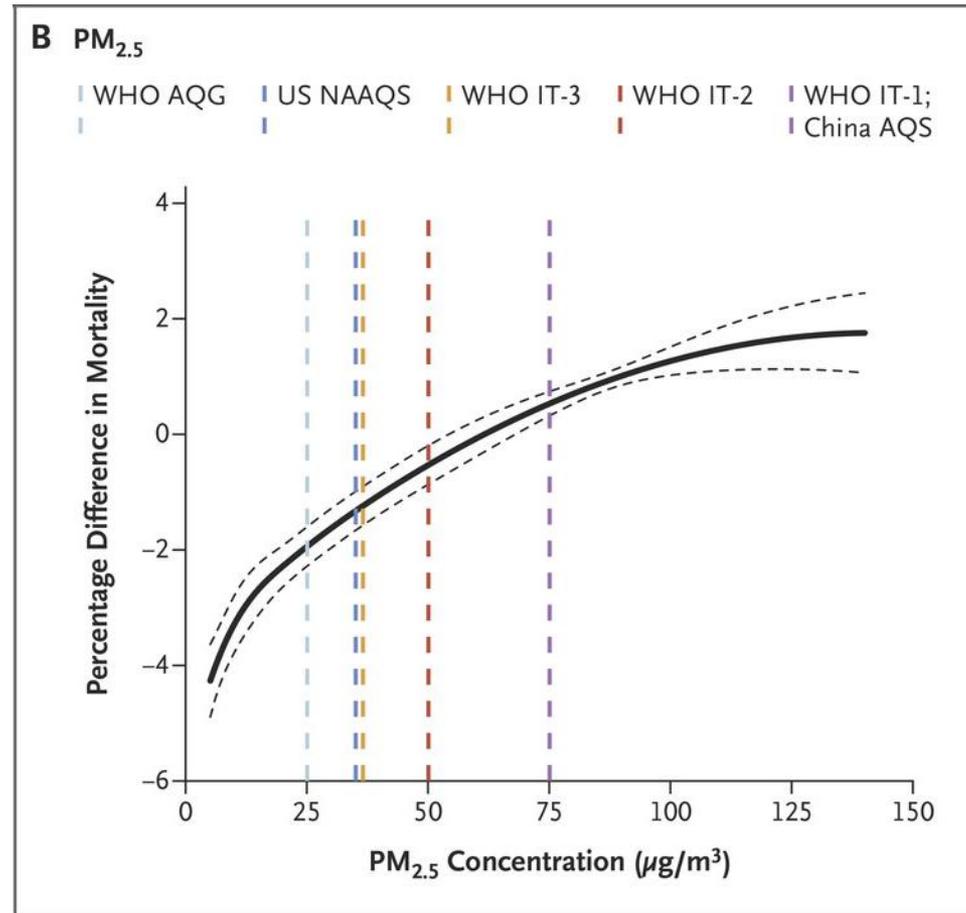
Pope et al. 2009: Results

- Air quality improved over time across the country
- Life expectancy increased (2.72 years)
- During each period, cross-sectional data showed negative association between life expectancy and pollution levels
- Reduction of $10 \mu\text{m}/\text{m}^3$ resulted in increase of 0.61 ± 0.2 years life
- Benefit regardless of baseline level; in fact greater increase for those areas with less pollution initially
- From 1980 through 2000, improved air quality accounted for approximately 15% of change in life expectancy



Increased mortality associated with short term air pollution exposure

- Data from over 600 cities in 24 countries
- Rolling 2 day average exposure
- All cause mortality
- Effects even at levels considered “safe”



C Liu et al. N Engl J Med 2019;381:705-715



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Geneva Steel Strike

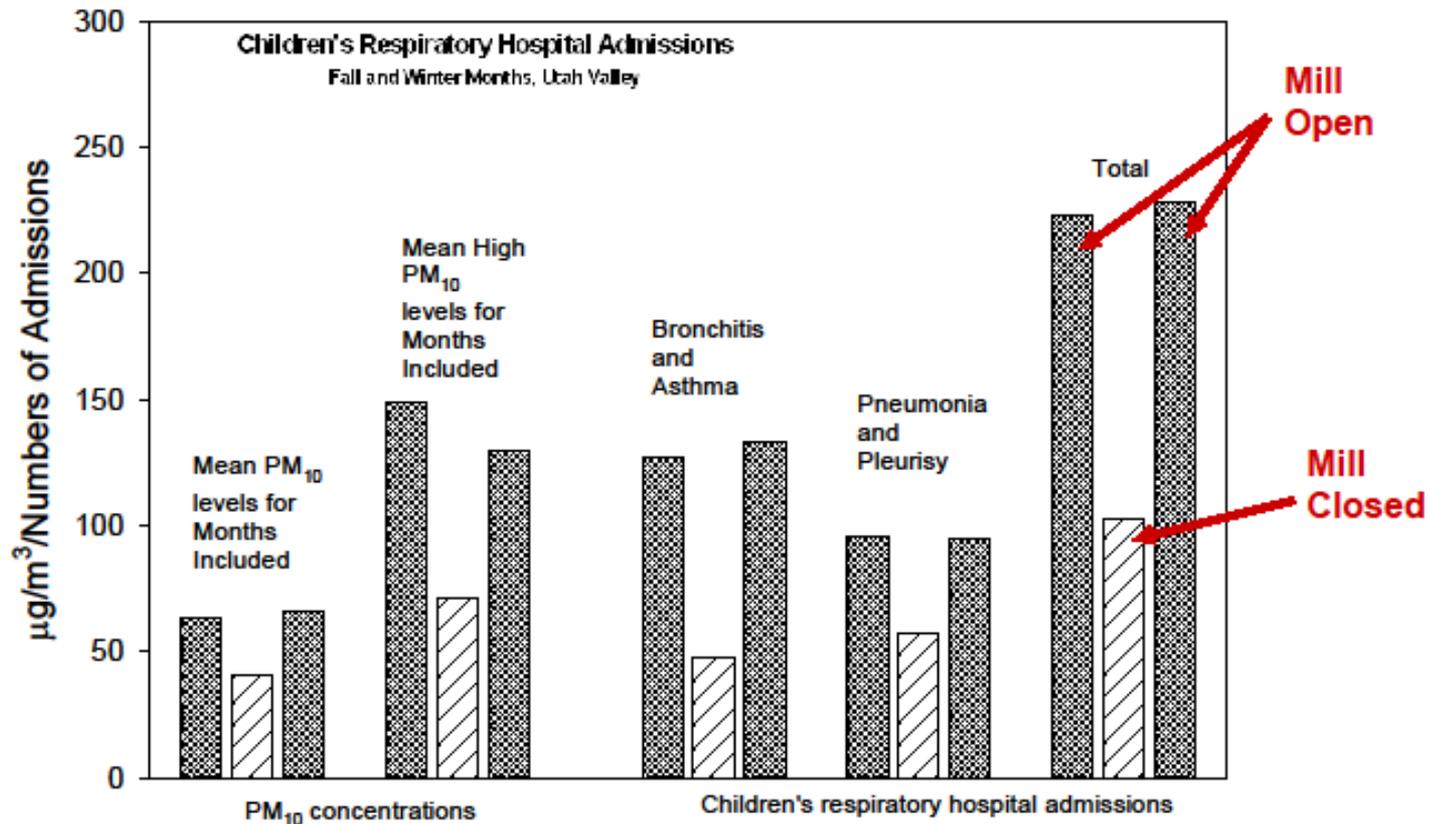


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When the steel mill was open, total children's hospital admissions for respiratory conditions **approx. doubled.**



Sources: Pope. Am J Pub Health.1989; Pope. Arch Environ Health. 1991



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Wildfires and Health



How bad is this year's Utah wildfire season? Actually, not that bad so far — unless you're trying to breathe.

Salt Lake Tribune August 17, 2018



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Constituents of wood smoke

- Regulated pollutants
 - Fine particles, carbon monoxide, nitrogen oxides
- Respiratory irritants impairing cellular function
 - Phenols, cresols, acrolein, acetaldehyde
- Organic carcinogens
 - Benzene, formaldehyde, 1,3 butadiene
- Carcinogenic cyclic compounds
 - Polycyclic aromatic hydrocarbons
 - Dioxins



Wildfires are major health events

- Wildfire activity around the world is associated with
 - Increased emergency room visits
 - Increased acute respiratory distress hospital visits
 - Increased hospital admissions for respiratory illness
 - Increased use of asthma medications
 - Decreased lung function
 - Increased evidence of inflammation in the blood
 - Increased mortality (over age 65)



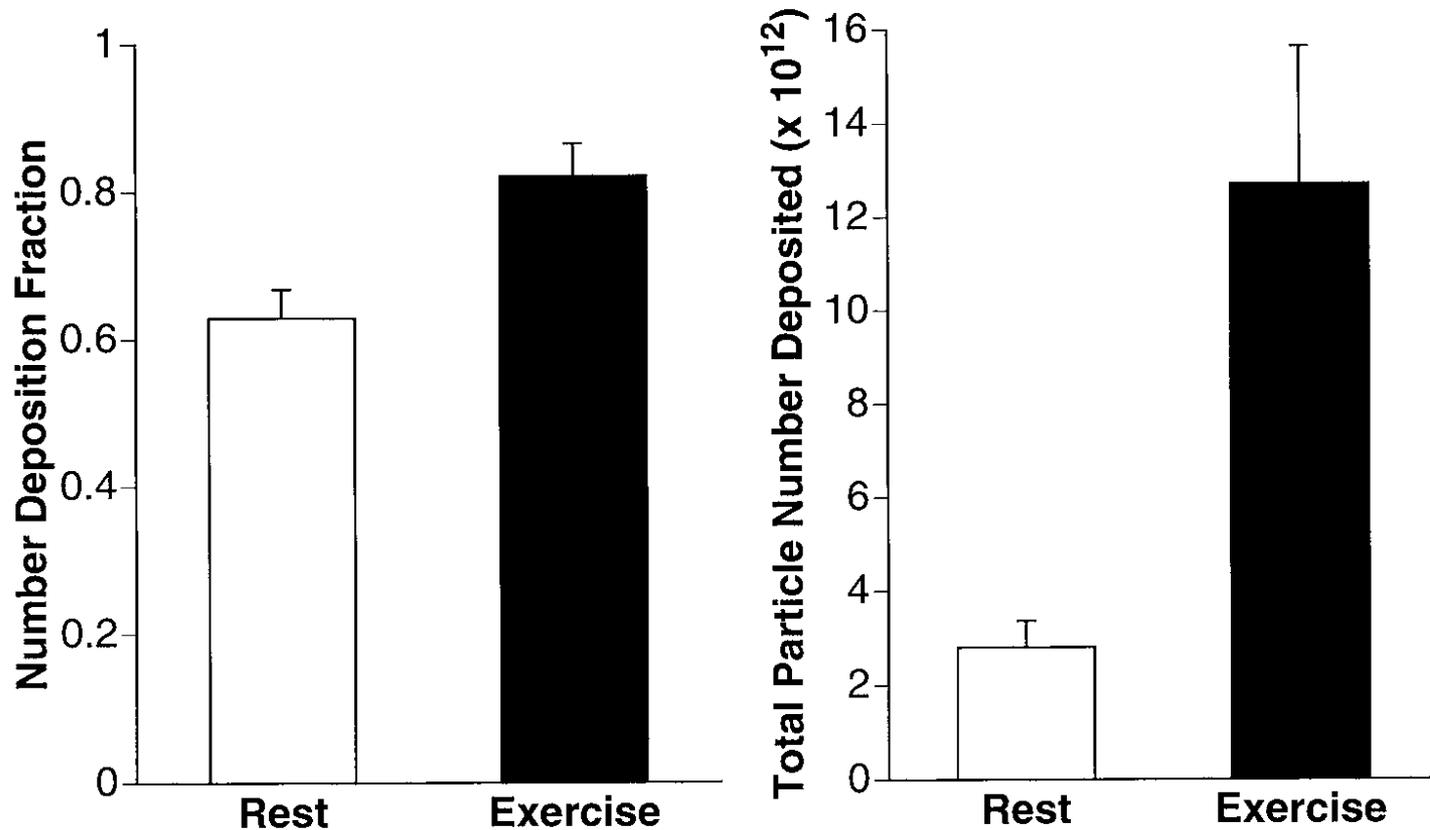
What should we tell our patients about exercise?



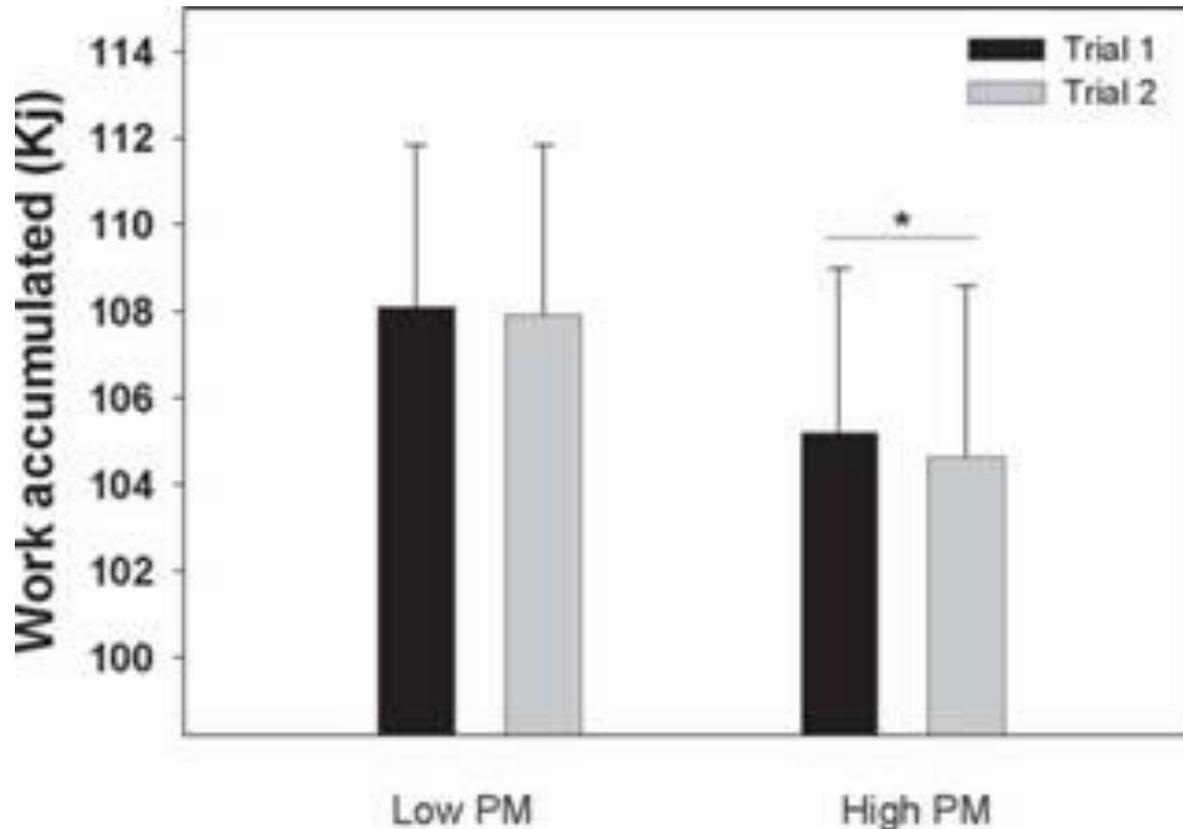
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Increased lung particle deposition during exercise



Impact of particulate air pollution on cycle work capacity



Rundell K W Br J Sports Med 2012;46:407-412



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What do I tell my patients?

- Pay attention to air quality; note if your symptoms change.
- When air quality is poor, it is particularly important to use your medications properly.
- During “bad air” days:
 - Exercise away from roadways
 - Exercise indoors
 - Stay indoors
- Starting to think about masks for high-risk individuals when outdoors



Air Quality Index

Air Quality Index

Action Forecast

Unrestricted Action



Voluntary Action



Mandatory Action



Health Forecast

Air Quality Index
(AQI)

PM 2.5

Ozone

| Air Quality Index (AQI) | PM 2.5 | Ozone |
|--------------------------------|--|-------------------|
| Good | 0 - 12.0 $\mu\text{g}/\text{m}^3$ | 0 - 0.054 ppm |
| Moderate | 12.1 - 35.4 $\mu\text{g}/\text{m}^3$ | 0.055 - 0.070 ppm |
| Unhealthy for Sensitive Groups | 35.5 - 55.4 $\mu\text{g}/\text{m}^3$ | 0.071 - 0.085 ppm |
| Unhealthy | 55.5 - 150.4 $\mu\text{g}/\text{m}^3$ | 0.086 - 0.105 ppm |
| Very Unhealthy | 150.5 - 250.4 $\mu\text{g}/\text{m}^3$ | 0.106 - 0.200 ppm |
| Hazardous | Above 250.5 $\mu\text{g}/\text{m}^3$ | Above 0.201 ppm |

Based on a 24-hour average. Based on an 8-hour average.

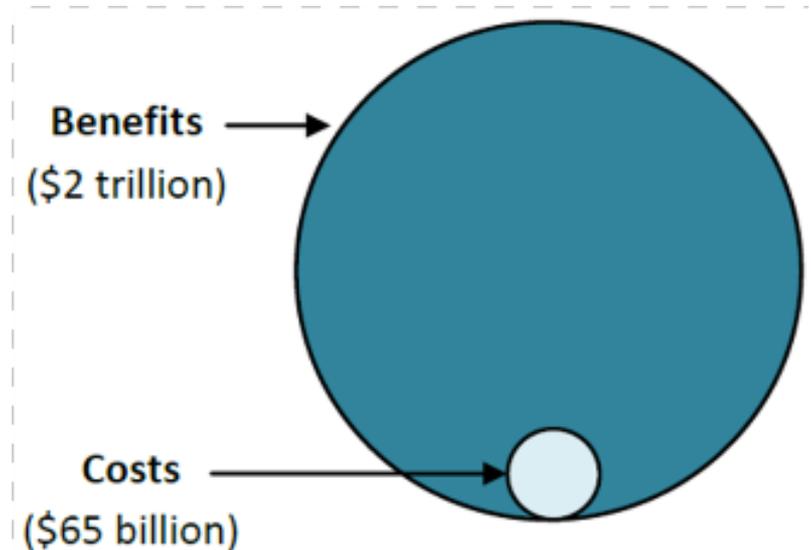


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Cost-benefit of the Clean Air Act 1990-2020

| | Cases prevented (2020) |
|------------------------------|------------------------|
| Adult mortality – particles | 230,000 |
| Infant mortality – particles | 280 |
| Mortality – ozone | 7100 |
| Chronic bronchitis | 75,000 |
| Acute myocardial infarction | 200,000 |
| Asthma exacerbation | 2,400,000 |
| Emergency room visits | 120,000 |
| Lost school days | 5,400,000 |
| Lost work days | 17,000,000 |



Clean Air Act 40th anniversary celebration





Thanks- happy to take questions



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