

### Million Hearts 2022:

# Understanding the Links between Environmental Pollutant Exposure and Cardiovascular and Cerebrovascular Events

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Justus Warren Heart Disease and Stroke Prevention Task Force

Research Triangle Park, NC May 8, 2018



### Wayne Cascio, MD

- No conflicts of interest
- The presentation represents the opinions of the speaker and does not necessarily represent the policies of the US EPA



## Air Pollution: A Leading Cause of the Global Burden of Disease

### In 2015 ambient PM<sub>2.5</sub> ranked <u>fifth</u> as a risk factor for global mortality -

#### Exposure to PM<sub>2.5</sub> caused:

- 4.2 million deaths
- 103.1 million disability-adjusted life-years (DALYs)

#### Between 1990-2015 deaths increased in association with PM<sub>2.5</sub> from:

• 3.5 million to 4.2 million

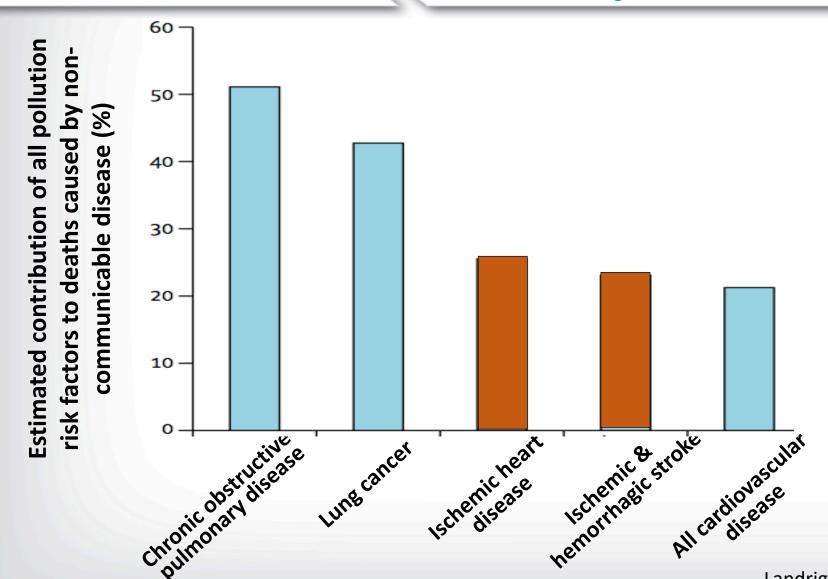
### Ozone exposure contributed to morbidity and mortality -

#### In 2015 ozone exposure is estimated to have accounted for:

- 254,000 deaths
- 4.1 million DALYs from chronic obstructive pulmonary disease



## Estimated Contribution of All Pollution to Deaths Caused by Non-Communicable Diseases, 2015



## All pollution contributes to:

- over 20% of ischemic heart disease
- over 20% of ischemic and hemorrhagic stroke
- 20% of all cardiovascular disease



## Despite Progress in the U.S. Air Pollution Continues to Impact Population Health

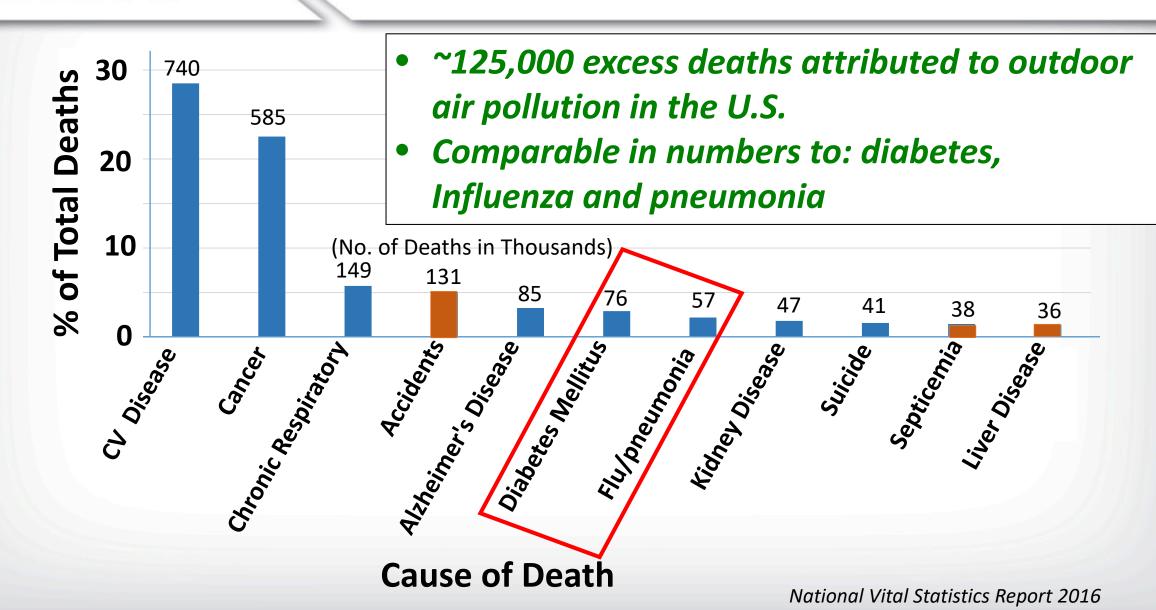
### Air Pollution Remains a Significant U.S. Public Health Concern

- Estimated excess mortality 125,000 deaths/year
- Over 20 million school days and work days lost
- Over 1 million life-years lost
- 122.5 million people living in counties with one or more pollutants exceeding the NAAQS in 2016



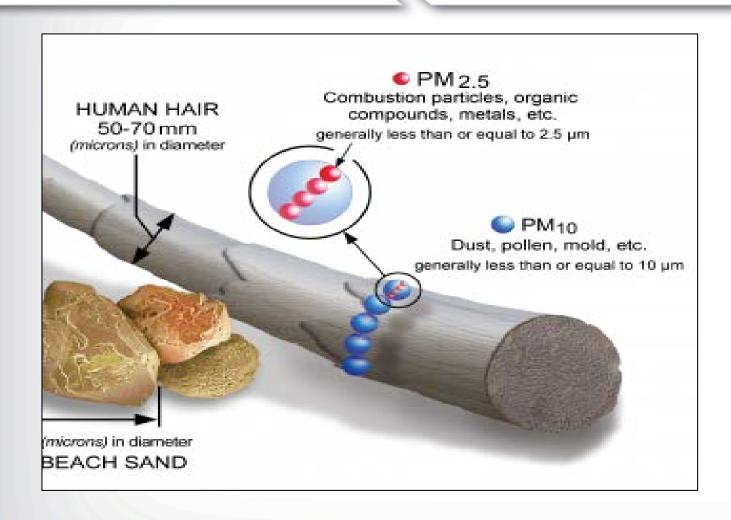


### Causes of Deaths in U.S.





### What is Airborne Particulate Matter?

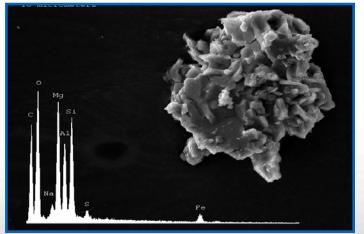


#### Particulate matter (PM) – "soot"

- from combustion sources
- mixture of solid particles and liquid droplets found in the air

## National Ambient Air Quality Standards (NAAQS)

- 35 ug/m<sup>3</sup> 24 hours
- 12 ug/m<sup>3</sup> annual average



Scanning
Electron
Micrograph
of
air particle



## Foundation of PM's Health Effect Exposure Linked to Morbidity and Mortality

#### **AHA Scientific Statement**

#### Air Pollution and Cardiovascular Disease

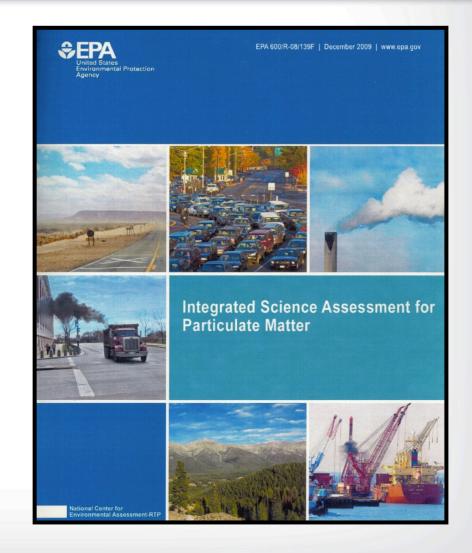
A Statement for Healthcare Professionals From the Expert Panel on Population and Prevention Science of the American Heart Association

Robert D. Brook, MD; Barry Franklin, PhD, Chair; Wayne Cascio, MD; Yuling Hong, MD, PhD; George Howard, PhD; Michael Lipsett, MD; Russell Luepker, MD; Murray Mittleman, MD, ScD; Jonathan Samet, MD; Sidney C. Smith, Jr, MD; Ira Tager, MD

Abstract—Air pollution is a heterogeneous, complex mixture of gases, liquids, and particulate matter. Epidemiological studies have demonstrated a consistent increased risk for cardiovascular events in relation to both short- and long-term exposure to present-day concentrations of ambient particulate matter. Several plausible mechanistic pathways have been described, including enhanced coagulation/thrombosis, a propensity for arrhythmias, acute arterial vasoconstriction, systemic inflammatory responses, and the chronic promotion of atherosclerosis. The purpose of this statement is to provide healthcare professionals and regulatory agencies with a comprehensive review of the literature on air pollution and cardiovascular disease. In addition, the implications of these findings in relation to public health and regulatory policies are addressed. Practical recommendations for healthcare providers and their patients are outlined. In the final section, suggestions for future research are made to address a number of remaining scientific questions. (Circulation. 2004;109:2655-2671.)

Brook RD et al. Circulation 2004

Short-term and long-term exposure to ambient air particulate matter is causally associated with cardiovascular morbidity and mortality (EPA ISA 2009)





## Short-term Air Pollutant Exposure Contribution to Cardiovascular Events

#### AHA Scientific Statement

## Particulate Matter Air Pollution and Cardiovascular Disease An Update to the Scientific Statement From the American Heart Association

Robert D. Brook, MD, Chair; Sanjay Rajagopalan, MD; C. Arden Pope III, PhD;

Jeffrey R. Brook, PhD; Aruni Bhatnagar, PhD, FAHA; Ana V. Diez-Roux, MD, PhD, MPH;

Formando Holquin, MD; Vuling Hong, MD, PhD, FAHA; Puscell V, Luceker, MD, MS, FAHA

#### Fine particulate matter (PM) or particle pollution can:

- Trigger heart attacks
- Trigger arrhythmia

- Trigger stroke
- Worsen heart failure

Heart disease patients should reduce their exposure to air pollution when levels are high



## Call for Public Health & Healthcare Action

 "Air pollution should" be viewed as one of several major modifiable risk **factors** in the prevention and management of cardiovascular disease."



European Heart Journal (2015) **36**, 83–93 doi:10.1093/eurheartj/ehu458

**CURRENT OPINION** 

## Expert position paper on air pollution and cardiovascular disease

David E. Newby<sup>1</sup>, Pier M. Mannucci<sup>2</sup>, Grethe S. Tell<sup>3</sup>, Andrea A. Baccarelli<sup>4</sup>, Robert D. Brook<sup>5</sup>, Ken Donaldson<sup>6</sup>, Francesco Forastiere<sup>7</sup>, Massimo Franchini<sup>8</sup>, Oscar H. Franco<sup>9</sup>, Ian Graham<sup>10</sup>, Gerard Hoek<sup>11</sup>, Barbara Hoffmann<sup>12</sup>,

 "Health professionals, including cardiologists, have an important role to play in supporting educational and policy initiatives as well as counseling their patients."



## Long-term Air Pollutant Exposure Contributes to Cardiovascular Morbidity & Mortality

December 2012 EPA/600/R-12/056F



### Provisional Assessment of Recent Studies on Health Effects of Particulate Matter Exposure

National Center for Environmental Assessment RTP Division Office of Research and Development U.S. Environmental Protection Agency Research Triangle Park, NC 27711

#### Ambient air particle pollution is associated with:

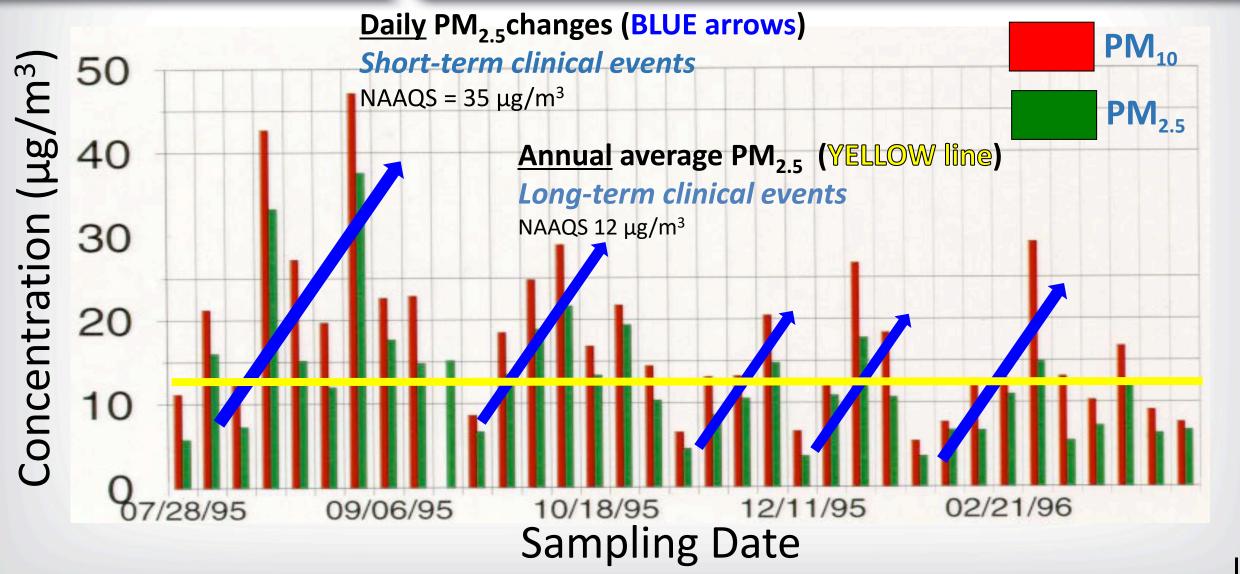
- Hypertension
- Ischemic heart disease

- Stroke
- Cardiovascular Mortality

Communities should be able to improve their cardiovascular health by complying with ambient air quality standards



### Daily Variability of PM<sub>10</sub> & PM<sub>2.5</sub> Chapel Hill, NC 1995-96





### Populations At-Risk Are Known

### Populations At-Risk from PM<sub>2.5</sub>

### Susceptible populations include -

- those with pre-existing cardiovascular disease
- those with pre-existing respiratory disease
- older adults
- those with lower socio-economic status
- children & the developing fetus

### Populations suspected to be at greater risk -

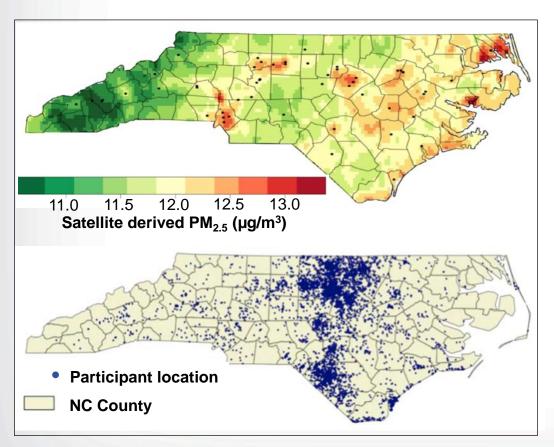
- those with chronic inflammatory diseases (e.g., diabetes, obesity)
- those with specific genetic polymorphisms (e.g., GSTM1)



## Health & Long-term Air Pollution Exposure

**Association between PM and Coronary Artery Disease** 

## 5,679 patients who underwent coronary angiography at Duke University between 2002–2009 and resided in North Carolina\*



## 1 $\mu$ g/m³ increase in annual average PM<sub>2.5</sub> was associated with an:

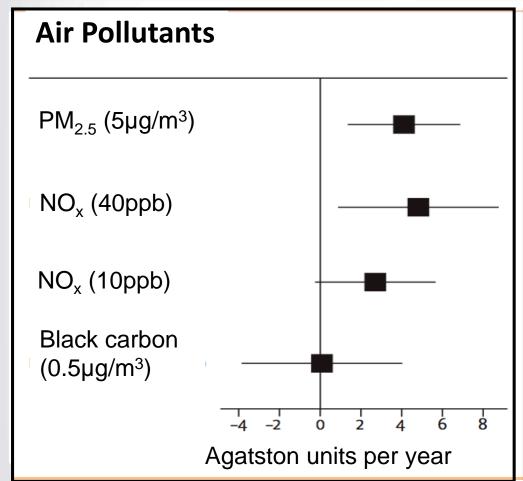
- 11.1% relative increase in odds of significant CAD
- 14.2% increase in the odds of having had a heart attack during the previous year

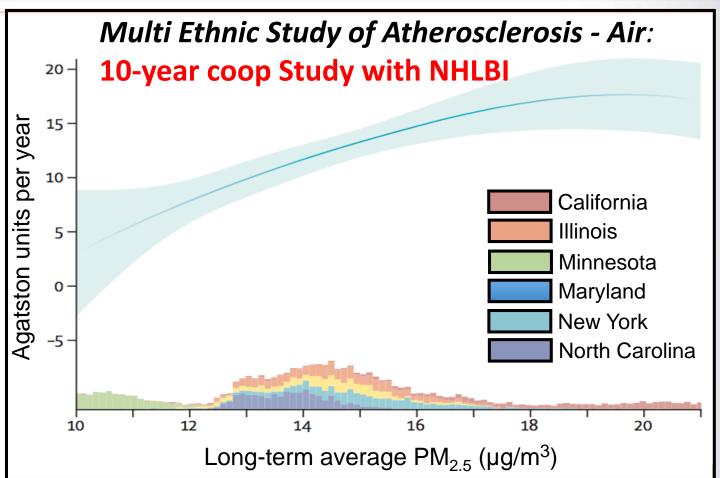
6,575 Ohio residents undergoing elective diagnostic coronary angiography found the same

relationship\*\*



## Long-Term PM<sub>2.5</sub> & NO<sub>2</sub> Exposure Increases Coronary Artery Calcium





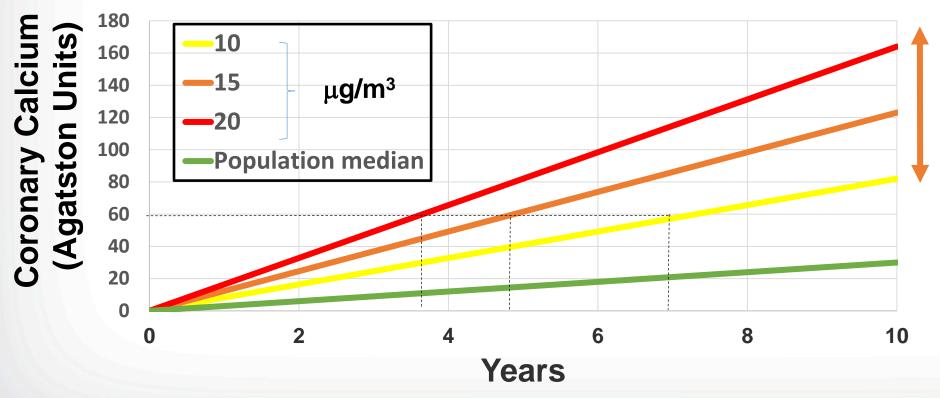
Long-term PM<sub>2.5</sub> and NO<sub>2</sub> increased coronary calcium, an indictor of atherosclerosis



## Long-term PM<sub>2.5</sub> & Nox Exposure Associated Atherosclerosis Progression

### MESA Air Study – Led by University of Washington

PM<sub>2.5</sub> and Coronary Calcium

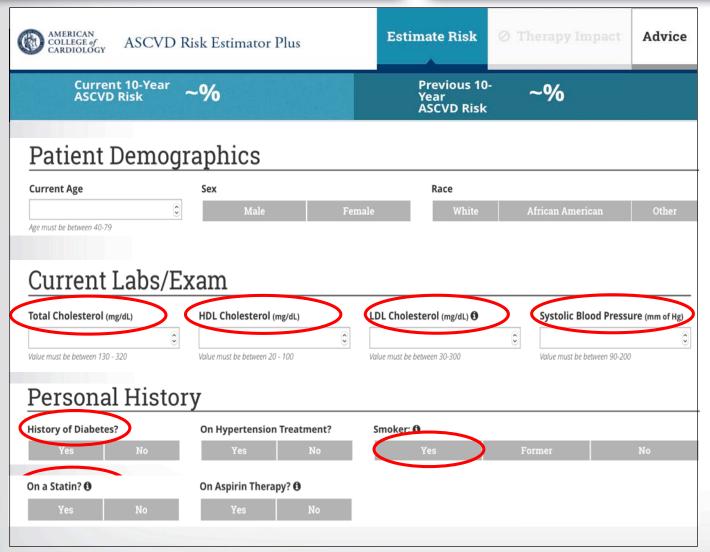


After 10 years 80 Agatston Unit difference between annual PM<sub>2.5</sub> of 10 and 20 µg/m<sup>3</sup>



### Air Pollution Worsens Vascular Risk Factors

Risk Factors for Atherosclerosis and Air Quality



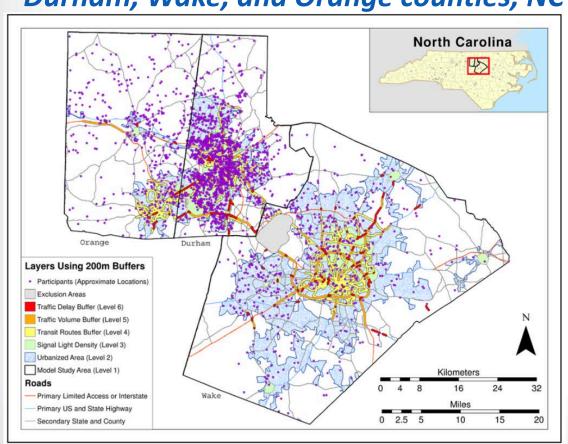
### **Poor Air Quality:**

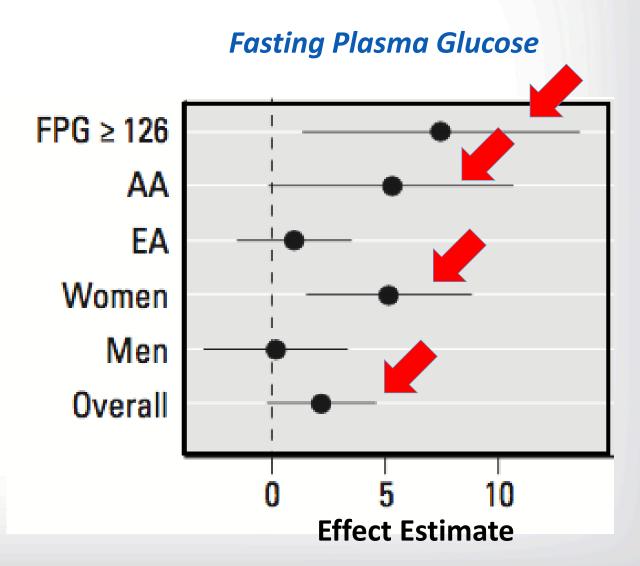
- Age might accelerate aging
- Total Cholesterol increases cholesterol
- HDL decreases HDL particle number
- LDL oxidizes LDL and ox-LDL receptor
- Systolic BP increases blood pressure
- Diabetes associated with type II diabetes
- Statin Therapy might be protective



## Roadway Proximity and Fasting Plasma Glucose

### Distribution of CATHGEN participants within the study area of Durham, Wake, and Orange counties, NC



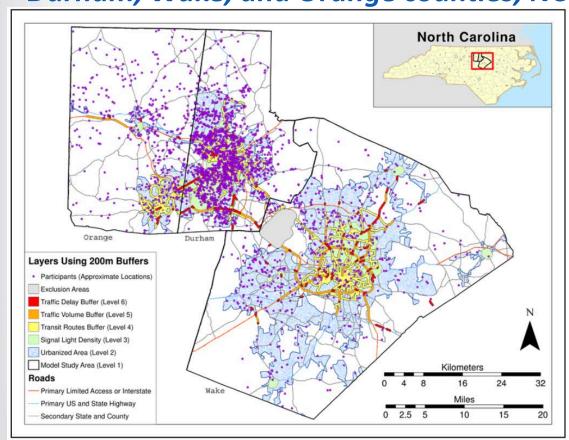


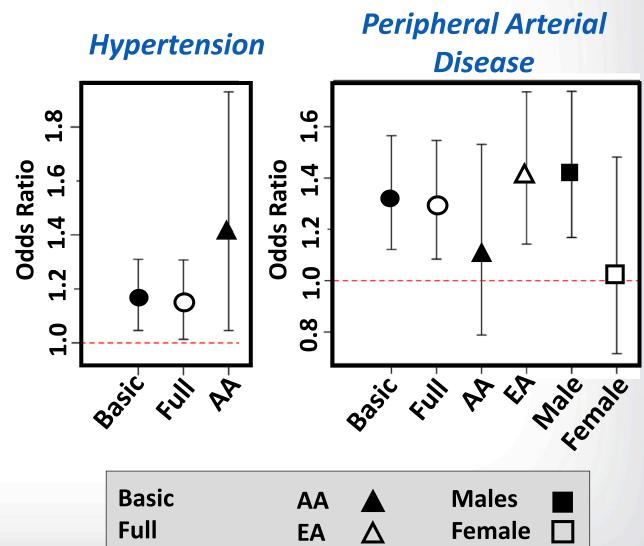
Ward-Caviness C et al. Environ Health Perspect 123:1007-1014



## Residential Proximity to Traffic Hypertension, Vascular Disease and

Distribution of CATHGEN participants within the study area of Durham, Wake, and Orange counties, NC

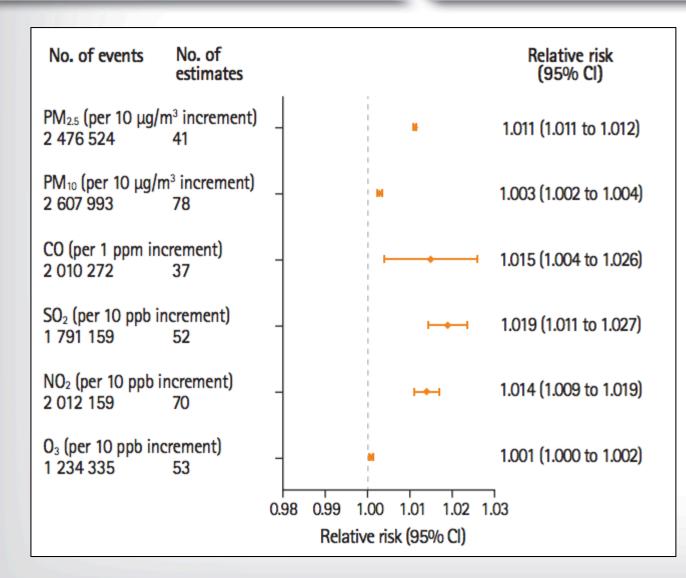




Ward-Caviness W et al. Arterioscler Thromb Vasc Biol. 2018;38:275-282



## Association Between Air Pollutants and Hospitalization or Mortality from Stroke



Systematic review and metaanalysis reported stroke to be associated with PM and air pollutants gases:

- PM<sub>2.5</sub> and PM<sub>10</sub> associated with admission to hospital for stroke and mortality
- NO<sub>2</sub> associated with hospital admission or mortality
- SO<sub>2</sub> and CO associated with admission or mortality

Lee KK, et al. Journal of Stroke 2018;20(1):2-11



### Air Pollution and Mortality

### Effect of PM on Survival and Subsequent Clinical Events

Zanobetti A & Schwartz J.

**Environ Health Perspect 2007** 

Koton et al.

Prev Med 2013

Tonne et al.

Eur Heart J 2013

Tonne et al.

*Int J Hyg Envir Health* 2016

Chen et al. EHP 2016

Environ Health Perspect 2016

**Mortality** 

CHF hospitalization

**MI** hospitalizations

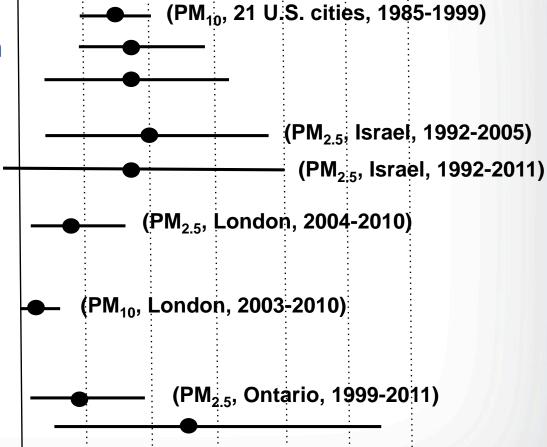
MI, CHF, Stroke

**Mortality** 

**Mortality** 

**Mortality** 

Mortality
MI Mortality



1.0 1.25 1.50 1.75 2.00 2.25 2.50 Hazard Ratio (per 10µg/m³)



## Personal Health Care Spending in U.S. for Chronic Disease is High

COPD	\$ 53.8 billion
Asthma	\$ 32.5 billion
Pneumonia	\$ 37.1 billion
Lung cancer	<b>\$ 13.1 billion</b>
Ischemic heart disease	\$ 88.1 billion
High blood pressure	\$ 83.9 billion
Stroke	<b>\$ 43.8 billion</b>
Heart failure	<b>\$ 28.5 billion</b>
Atrial fibrillation	<b>\$ 27.7 billion</b>
Peripheral vascular disease	\$ 2.7 billion
Diabetes	\$101.4 billion
Preterm birth	\$ 4.9 billion

1/2 Trillion

Dollars

in

2013

Dieleman JL et al. JAMA 2016





- Air pollution adversely affects:
  - Health, Longevity, Healthcare Resource Utilization and Public Welfare (e.g. effects on visibility, vegetation, and ecosystems)
- Most healthcare professionals & patients at-risk know of air pollution's adverse health effects

Despite Knowledge of the Risks the Healthcare System is Not Engaged

- Few healthcare professionals discuss the risks with their patients
- At-risk patients don't take action to reduce exposure



## EPA's Healthy Heart Program

Increasing Environmental Health Literacy

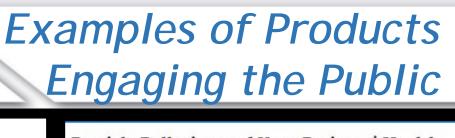


EPA's Healthy Heart program aims to prevent heart attacks and strokes by:

- Raising public awareness about the role outdoor air pollution plays in cardiovascular health, and
- Steps individuals can take to reduce their pollution exposure



## Engaging the Public





#### Healthy Heart Toolkit and Research: Steps You Can Take

Steps You Can Take to Reduce Health Effects

from Air Pollution

Studies show that air pollution can trigger are at risk for these conditions. If you have a exposure to high levels of air pollution.

#### When are air pollution levels h

- · Any time of year
- · When weather is calm
- Near busy roads In urban areas
- In industrial areas







**Daily Air Ouality** 

#### Heart Disease, Stroke, and Outdoor Air Pollution

 Did you know that air pollution can trigger heart attacks, stroke, and other health effects?

Medical studies show that air pollution can trigger heart attacks, stroke, and irregular heart rhythms—especially in people who are already at risk for these conditions. Also, for people with a medical condition called heart failure, air pollution can further reduce the ability of the heart to pump blood the way that it should. Very small particles are the pollutants of greatest concern for triggering these effects. Particle pollution is found in haze, smoke, and dust—and sometimes in air that looks clean. This fact sheet tells you how you can:

· Get up-to-date information about your



3 How can you protect your health?

#### Particle Pollution and Your Patients' Health

Helps health care providers advise their patients about particle pollution exposure.

Air Sensor Guidebook



EnviroFlash

27707

Current

2/6/2013

8:00 PM EST

Pollutant:

PM2.5

40

Good

Current

Location

Go

The Air Quality Index (AQI) for Raleigh-Durham-Chapel Hill

This course is designed for family medicine physicians, internists, pediatricians, occupational and rehabilitation physicians, nurse practitioners, nurses, asthma educators, pulmonary specialists,

AirNow

Today's AQI Forecast

Current AQI AQI Loop More Maps

cardiologists, and other medical professionals. Start the Course





recitions repaid for reducts made again del teste factors Consignite Publish





Good

#### Efectos de los Contaminantes Comunes del Aire

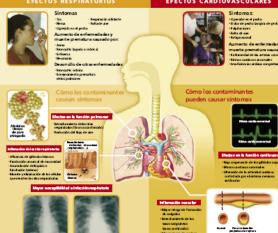
Click to see map

U.S. Embassies and Consulates
Data from air quality monitors at select U.S. embassies and consulates

6/15/17: National Air Quality Conference, September 11 13, 2017. Registration Now Open and Call for

6/7/17: 2017 Air Quality Flag Program Spring Challenge

Air Quality Index | Ozone | Particle Pollution | Smoke



Reduzca su riesgo, usando el Índice de Calidad del Aire (AQI por sus siglas en inglés; al planear actividades al aire libre – www.aimow.gov



## Public Education Air Quality Index Available at AirNow.gov

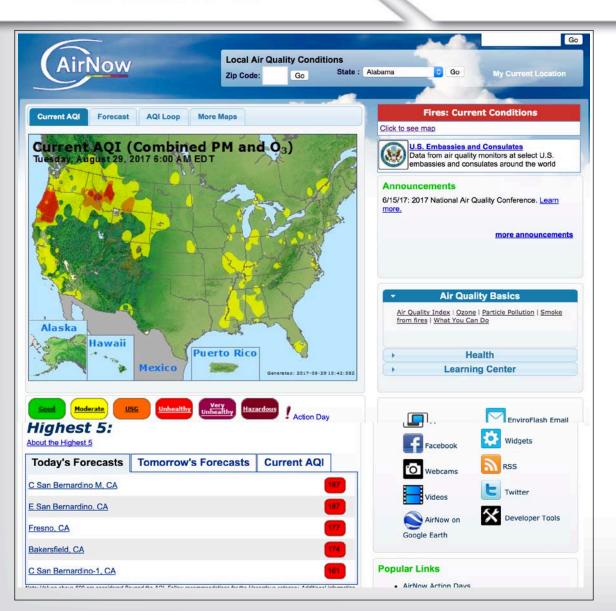


- Color scale detailing how clean or polluted the air is
- Local air quality conditions also often part of local weather reports
- Where can it be found?
  - Local TV, radio or newspapers
  - AirNow app
  - Email alerts at www.enviroflash.info

Descriptors	Cautionary Statement
Good 0 – 50	No message
Moderate 51 – 100	Unusually sensitive individuals
Unhealthy for Sensitive Groups 101 - 150	Identifiable groups at risk - different groups for different pollutants
Unhealthy 151 - 200	General public at risk; sensitive groups at greater risk
Very Unhealthy 201 - 300	General public at greater risk; sensitive groups at greatest risk

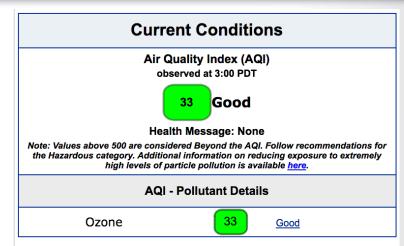


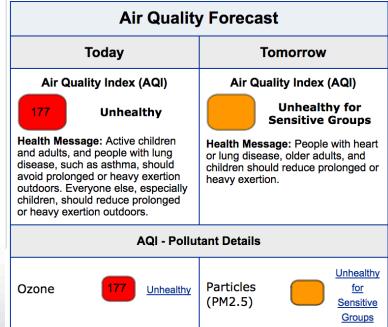
## Informing the Public www.airnow.gov via the Internet



AirNow
National AQIs
& forecast
August 29,
2017

Fresno, CA
AQI & forecast
August 29,
2017







### Healthy Heart Toolkit

www.epa.gov/air-research/healthy-heart-toolkit-and-research



### Healthy Heart Toolkit and Research: Steps You Can Take

## Steps You Can Take to Reduce Health Effects from Air Pollution

Studies show that air pollution can trigger heart attacks, strokes and worsen heart failure in people who are at risk for these conditions. If you have a heart condition, you could benefit by reducing your exposure to high levels of air pollution.

#### When are air pollution levels high?

- Any time of year
- When weather is calm
- Near busy roads
- In urban areas
- In industrial areas
- When there is smake

#### Daily Air Quality

- Check Pollution Forecasts
- Get Free Email Alerts
   EXIT

#### Resources

- Be Smart, Protect Your Heart video EXIT
- Heart Disease, Stroke and Outdoor Air Pollution
- Million Hearts Initiative:

- When are air pollution levels high?
- Are you at risk?
- Steps to Protect Your Heart
- How to Reduce your Risk?
- Warning Signs of a Heart Attack
- Warning Signs of a Stroke





## CME for Health Care Professionals Particle Pollution and Your Patients' Health



www.epa.gov/particle-pollution-and-your-patients-health

**Environmental Topics** 

**Laws & Regulations** 

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Patient Exposure and the Air Quality Index

Cardiovascular Effects

**Respiratory Effects** 

Patient Exposure and High Particle Pollution Events

#### **Clinical Scenarios**

Course Home

About this course

What is Particle Pollution?

Particle Pollution Exposure

**Frequent Questions** 

Course Outline/Key Points

**Review Questions** 

**Patient Education Tools** 

**Course Evaluation** 

References

Glossary

#### Particle Pollution and Your Patients' Health

An evidence-based training for healthcare professionals that:

- Describes the biological mechanisms responsible for the cardiovascular and respiratory health effects associated with particle pollution exposure.
- Provides educational tools to help patients understand how particle pollution exposure can affect their health and how they can use the Air Quality Index to protect their health.



This course is designed for family medicine physicians, internists, pediatricians, occupational and rehabilitation physicians, nurse practitioners, nurses, asthma educators, pulmonary specialists, cardiologists, and other medical professionals.

Start the Course

Course developers



## Partnering with HHS' Million Hearts® Recommendations Supported by EPA Science





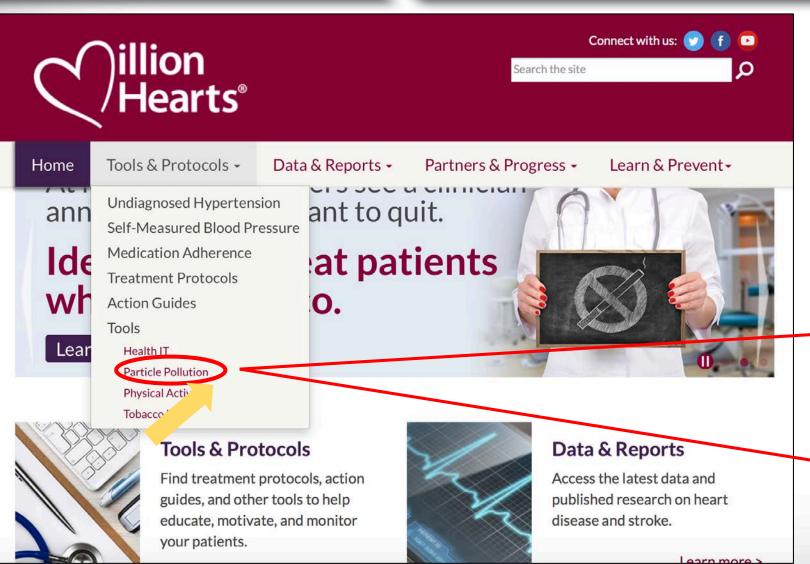
## EPA contributes the *Healthy Heart* program to lower air pollutant exposures in at-risk populations in an effort to:

- decrease heart attacks and strokes
- improve vascular disease outcomes
- decrease disability and healthcare expenditures
- decrease the societal burden of vascular diseases



### Million Hearts®

### **Provides Educational Tools on Particle Pollution**







## The Environmental "Buckets" of Prevention Framework

## **Total Population Community-Wide Prevention**

NAAQS
Built-Environment
Health Literacy

- Attain & maintain NAAQS Stds
- Improve built-environment:
  - Places for physical activity
  - Create healthier nearroad environments
- Improve overall CV health status

## Innovative Clinical Prevention



- Optimize clinical care of the at-risk priority population
- Increase awareness of health effects of PM among physicians, health care professionals, and the at-risk population
- Provide guidance to lower exposure & associated risk

## **Traditional Clinical Prevention**

"evidence-based" clinical prevention management strategies

 Long-term indoor air filtration lowered markers of oxidative stress and inflammation

> (Chuang H-C, et al. Environ International 2017)

#### **Public Health**

#### **Health Care**



## High Resolution Air Pollution Mapping Small Scale Variability due to Local Sources

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Article

pubs.acs.org/est

#### Oakland, CA

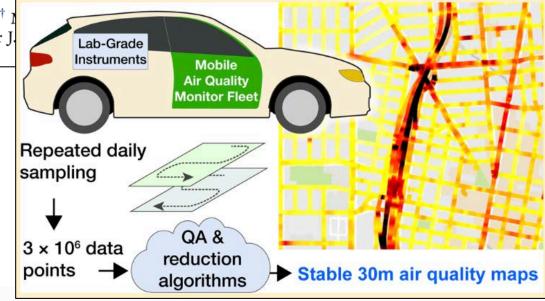
Spatial distribution of Black Carbon (BC)

High-Resolution Air Pollution Mapping with Google Street View Cars: Exploiting Big Data

Joshua S. Apte,\*,†© Kyle P. Messier,†,‡ Shahzad Gani,† Melissa M. Lunden, Julian D. Marshall,‡ Christopher J. and Steven P. Hamburg‡

Apte JS et al. Environ Sci Technol 2017

Hankey S et al. Population-Level Exposure to Particulate Air Pollution during Active Travel: Planning for Low-Exposure, Health-Promoting Cities. Environmental Health Perspectives 125:527–534, 2017







### Moving to the Future

## **EDVIRONMENTAL** Science & lechnology

Policy Analysis

pubs.acs.org/est

Forecast-Based Interventions Can Reduce the Health and Economic Burden of Wildfires

Ana G. Rappold,\*',† Neal L. Fann,‡ James Crooks,† Jin Huang,§ Wayne E. Cascio,† Robert B. Devlin,† and David Diaz-Sanchez†

Forecast-based interventions
predicted to reduce the health
and economic burden of
wildfires

Rappold AG, et al.

Environ Sci Technol 2014

Cost effectiveness is improved by intervening only in the homes of those at highest risk, e.g. older persons ORIGINAL ARTICLE

WILEY

Effectiveness and cost of reducing particle-related mortality with particle filtration

Indoor Air. 2017;1-12.

W. J. Fisk D | W. R. Chan

Fisk WJ, Chan WR Indoor Air 2017



The Global Threat of Outdoor Ambient Air Pollution to Cardiovascular Health
Time for Intervention

Robert D. Brook, MD

Anthropogenic ambient fine particulate matter less

etal air pollution problem appears to be

"... we believe that the time is ripe to definitively test the efficacy of personal-level interventions..."

Brook RD, et al. JAMA Cardiol. 2017



### Summary and Opportunities

- High attributable health burden
- Particle pollution increases short- and long-term cardiovascular morbidity & mortality
- Improvements in air pollution levels reduce health impacts and increase life expectancy
- Many regions of US fail to meet EPA standards >100 million exposed
- EPA is working with the States to help communities meet NAAQS
- Older-people, those with pre-existing heart & lung disease, & diabetes are at higher risk from air pollution



### Summary and Opportunities

- High-risk patients should be educated about risks of air pollution and educated about measures to reduce exposure
- Decreased short-term exposure in high patients is predicted to mitigate adverse health effects in high-risk individuals
- Randomized controlled trials are needed to proved effectiveness of interventions to reduce exposure
- Health risks need to be addressed through integrated efforts of public health and health care at the community and individual level
- More effective health communication strategies are needed to encourage health protective behaviors





## Thank you

Wayne E. Cascio, MD, FACC Director, National Health and Environmental Effects Research Laboratory Office of Research and Development U.S. Environmental Protection Agency

Email: cascio.wayne@epa.gov

- No conflicts of interest
- The presentation represents the opinions of the speaker and does not necessarily represent the policies of the US EPA



### Fall in Air Pollution Related Deaths Over Time

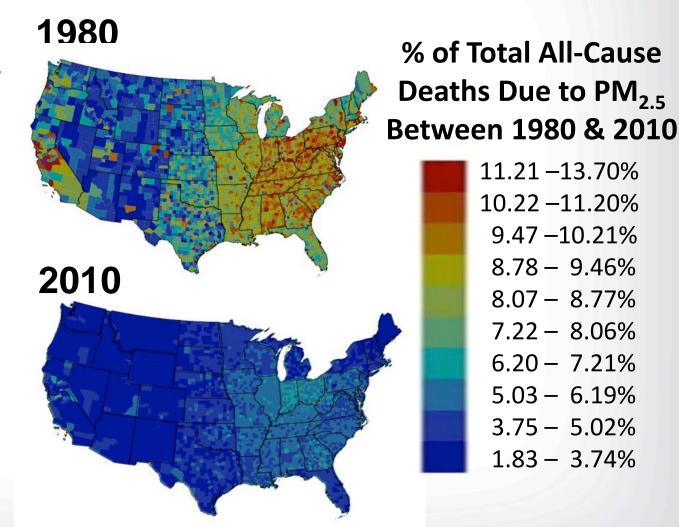
Fraction of Total All-Cause Deaths Attributed to PM<sub>2.5</sub>

## After the implementation of local, state, and federal air quality policies

 PM<sub>2.5</sub> precursor emissions declined over the course of several decades

Between 1980 - 2010, PM<sub>2:5</sub> exposures fell by about half, and estimated excess deaths decreased by about a third

 California, Virginia, New Jersey, and Georgia had some of the largest estimated reductions in PM<sub>2.5</sub>-attributable deaths





## Long-term Health Effects of Air Pollution PM<sub>2.5</sub> Exposure and Post-MI Survival in Ontario, Canada

### Post-Myocardial Infarction Survival Ontario, Canada 1999-2011

- 8,873 patients with 4,016 non-accidental deaths
- Mortality follow-up through 2011
- Cumulative time-weighted exposures to PM<sub>2.5</sub>
   were derived from satellite observations
- For each 10-μg/m³ increase in PM<sub>2.5</sub> non-accidental mortality increased by 22%

#### **Conclusions:**

 Long-term air pollution exposure adversely affects the survival of Heart Attack patients

### Post-Stroke Survival London, England 1995-2006

- 3,320 patients with 1,856 deaths
- Stroke follow-up to mid-2006
- Outdoor NO<sub>2</sub> and PM<sub>10</sub> modeled for 2002
- HRs were adjusted for relevant factors
- 10-μg/m³ increase in NO₂ and PM₁₀ was associated with a 28% and 52% increase in risk of death, respectively

#### **Conclusions:**

- Improvements in outdoor air quality might contribute to better survival after stroke
- A 10  $\mu$ g/m³ reduction in NO<sub>2</sub> exposure is expected to reduce mortality comparable to that for stroke units



## Air Pollution

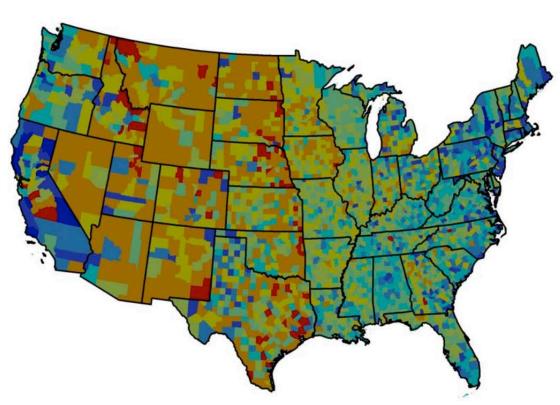
### Public Health Benefits of Decreasing PM<sub>2.5</sub> 1980-2010

## Change in the % of Death Due to PM<sub>2.5</sub> Between 1980 - 2010



6.08 - 8.11%

8.12 - 11.7%



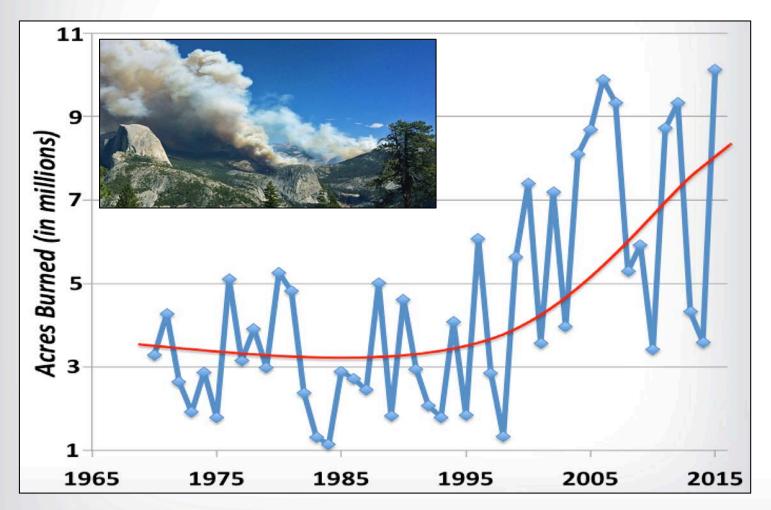
# Relative to a hypothetical population with exposures held constant at 1980 levels

- people born in 2050 would live about 1 year longer
- there would be a cumulative gain of 4.4 million life years among adults ≥30 years of age



## Wildfire: An Issue of Concern for the States

Impacts of Local and National Importance



Adapted from https://www.nifc.gov/fireInfo/fireInfo\_stats\_totalFires.html

- Wildland fires accounted for 38% of PM<sub>2.5</sub> emissions in 2014
- 2017 was substantially worse than the average of the last 10 years
- Between 2001 2010 over 40% of the country's large wildfires occurred in the Southeast
- U.S. spends more than
   \$2 billion each year to fight wildfires



## Wildfire-PM<sub>2.5</sub> Increases Heart Attack & Stroke

California 2015 Wildfire Study

**Results:** Wildfire-PM<sub>2.5</sub> associated with heart attacks and strokes for all adults, particularly for those over 65 years old

- Increase in risk the day after exposure:
  - All cardiovascular, 12%
  - Heart attack, 42%
  - Abnormal heart rhythm, 24% (same day)
  - Heart failure 16%
  - Stroke 22%
  - All respiratory causes 18%

*Impact:* Highlights the importance of decreasing exposure in at-risk populations

#### All Cardiovascular Causes

