US ERA ARCHIVE DOCUMENT





CLEAN **AIR** RESEARCH PROGRAM

BUILDING A SCIENTIFIC FOUNDATION FOR SOUND ENVIRONMENTAL DECISIONS

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The EPA Air, Climate, and Energy Intramural Research Program (ACE)

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Office of Research and Development

CLARCs Kick-Off Meeting April 7-8, 2011 RTP, NC



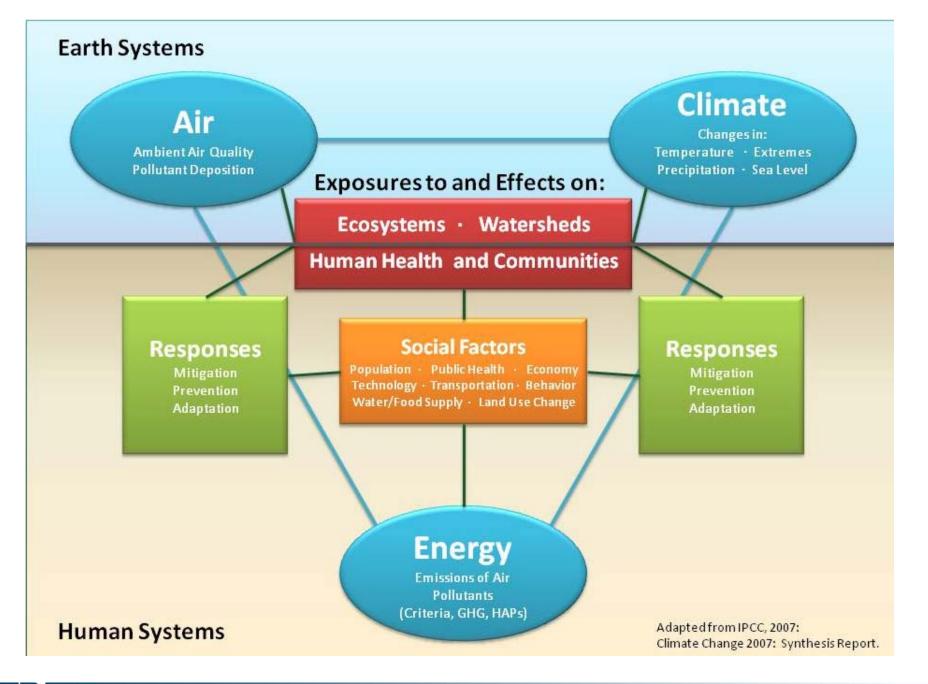
Problem Statement:

Protecting health and the environment from the impacts of climate change and air quality are central 21st century challenges. These challenges are complicated by the interplay between air quality, the changing climate, and emerging energy options.

Vision:

The ACE program will provide cutting-edge scientific information and tools to support EPA's strategic goals to protect and improve air quality and take action on climate change.







The ACE Approach to Research



Epidemiology

Clinical and Animal Toxicology



Intramural Research Program

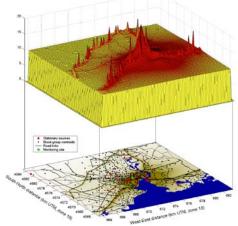


Emission source characterization

Federal / State /
Academic
Partners

EPA Ambient
Monitoring Network
(NCore)





Atmos. measurements & models for exposure



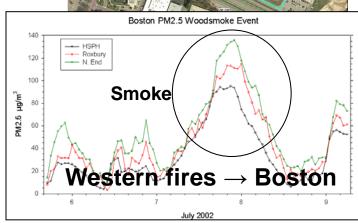
Research Theme 1: Assess Impacts

Assess human and ecosystem exposures and effects associated with air pollutants and climate change at individual, community, regional, and global scales.



Objectives:

- Assess multipollutant exposures and effects
- Develop innovative approaches to assess exposures and effects of pollutants in the atmosphere



- Identify characteristics of populations and ecosystems susceptible to exposure to air pollutants and climate change
- Inform review of NAAQS



Research Theme 2: Prevent and Reduce Emissions

Provide data & tools to develop and evaluate approaches to prevent and reduce emissions of pollutants to the atmosphere, particularly environmentally sustainable, cost effective, and innovative multipollutant and sectorbased approaches



Objectives:

- Support management air pollution problems at different scales of time and space
- Develop methods and data for life-cycle analyses of alternative pollution reduction and energy options
- Innovative technologies to support implementation of management strategies
- Evaluate pollution reduction and prevention solutions
- Support implementation of NAAQS



Research Theme 3: Respond to Challenges in Climate and Air Quality HEALTH EFFECTS OF AIR POLLUTION

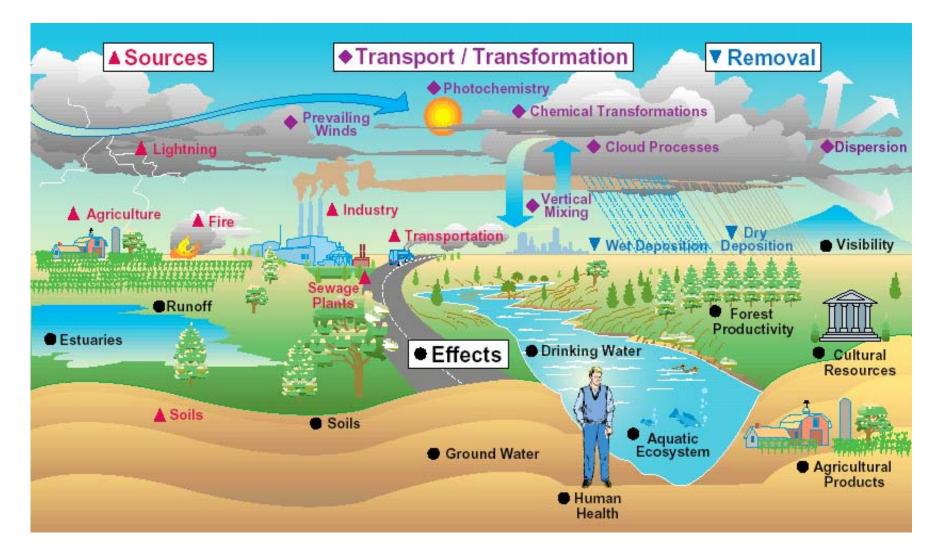
Provide human exposure and environmental modeling, monitoring, metrics and information needed by individuals, communities, and governmental agencies to adapt to the impacts of climate change and make informed public health decisions regarding air quality

Objectives:

- Evaluate alternative adaptation strategies
- Devise innovative methods to inform individual- and community-level adaptation to climate change and decision making for air quality
- Evaluate social, behavioral, and economic factors that influence adaptation strategies for climate change and decision making for air quality



Moving Towards a Multi-pollutant Research Program





EPA – the 5th CLARC?

- The EPA intramural air pollution research program is often referred to as the "5th Center."
- However, in terms of scientific personnel and the amount of dollars invested in air pollution research, this program is larger than all 4 Centers combined.
- Approximately 225 people are engaged in air pollutant research in EPA
- The EPA intramural program spent nearly 80 million dollars on air pollution research last year.

It is not feasible to discuss the entire program in 30 minutes. This presentation will focus on a small number of projects which are similar to projects being done by the CLARCs, to facilitate potential collaborations between ORD and the CLARCs.



Project 1. Near Road Studies

 Near-road Exposures and effects from Urban air pollutants Study (NEXUS)

Cooperative agreement with U. Michigan

 Research Triangle Area Mobile Source Emissions Study (RAMSES)





NEXUS Looks at Effect of Traffic on Asthmatic Children in Detroit

- Which measures of traffic-related air pollution are most closely associated with aggravated asthma?
- Do children with asthma who live near major highways show more inflammation and other biological responses?
- Does traffic exposure influence the likelihood of respiratory viral infections in children with asthma?





NEXUS Study Design

- 105 children ages 6-16 with persistent asthma
- 3 groups of 35 each who live within 150 m of:
 - High traffic/high diesel truck roadways
 - High traffic/low diesel truck roadways
 - Low traffic roadways
- Respiratory health evaluated seasonally and when children report respiratory viruses
- Leveraged with on-going studies of asthma in Detroit funded by NIEHS
- Concurrent measures of traffic-related air pollutants:

Home indoor/outdoor
Select schools outdoor





RAMSES Looks at Effect of Traffic Components on People with Cardiovascular Disease in RTP

- Although near road exposures are known to be particularly potent in causing adverse health effects, there is no consensus as to which components are responsible for these effects.
- What are the short and long term impacts of living near roads in those with CV disease.





RAMSES Has Three Phases

- Phase 1. A cohort of 7000 people who visited a Duke Clinic for angioplasty and have been followed over time will be used to determine if long term residence near roads with different traffic patterns is associated with severity or progression of CV disease.
- Phase 2. A panel of 50 volunteers from the cohort will be used in a follow-up study to associate acute effects with ambient PM, PM components, and gases.
 - Exposure assessment by community, home, and personal monitoring as well as modeling and proximity to roads
- Phase 3. The phase 2 cohort will be brought to roads that different in traffic patterns (e.g. smooth flow vs stop and go) and acute health effects associated with specific components of near road pollution.



Near Road Toxicology Studies

 Size-fractionated particles collected near (20m) and far (275m) from road

• Extracted particles:

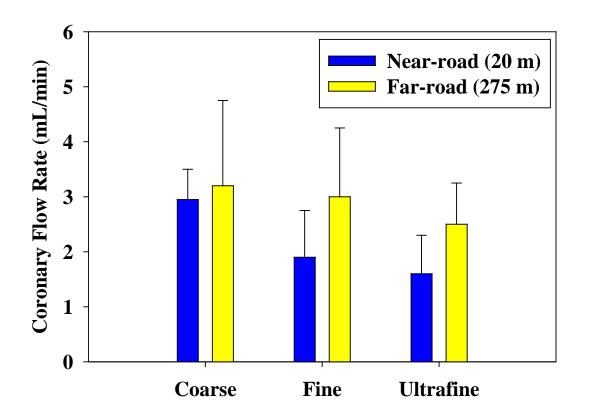
Chemically analyzed

Instilled in mice and allergy & cardiopulmonary responses

evaluated

Used for in vitro toxicology studies





Near-road fine and ultrafine PM reduced blood flow to the heart, suggesting traffic related particles may adversely affect the heart



Project 2. Are there combined effects of multiple pollutants (e.g. synergistic, additive, antagonistic)?

 It is difficult for epidemiology studies to disentangle the effects of pollutants which are co-located in time and space.

Does NO₂ cause health effects or is it a surrogate for near road exposures?

 Animals or humans can be exposed to simple mixtures (binary, trinary) of air pollutants or agents of climate change.

CAPS + O3, or diesel + NO2

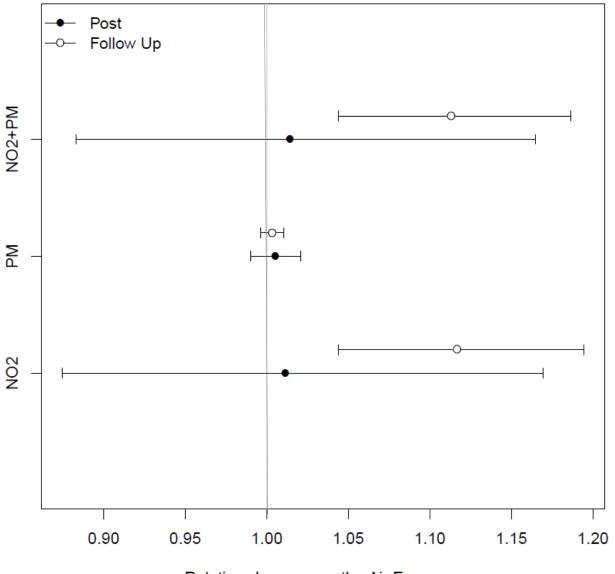


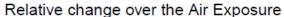


Exposure	PM Mass Concentration (ug/m³)	Pulmonary Function	BAL Fluid Cells	BAL Fluid Markers	Cardiac Endpoints	Plasma Markers
Coarse	89.0 ± 49.5	No Effect	↑PMNs	↓Protein	↓SDNN	↓tPA
Fine	120.4 ± 14.1	↓ DLCO	↑ Cells ↑ PMNs ↑ Monocytes	↓ IL-8 Fibrinogen	↑Norm. QT Var	↑Fibrinogen ↓ WBCs ↓ VLDL
Ultrafine	47.0 ± 20.2	No Effect	No Effect	↑ IL-8	↑ HF, LF ↑ Norm. QT Var ↓ QTc	↑ D-Dimer ↑ PAI-1 ↓ HDL, VLDL



Change in Low Density Lipoprotein







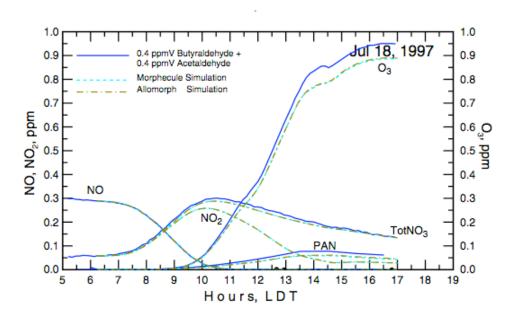
Project 3. Simulated Atmosphere Research

- Goal is to create artificial atmospheres that are reflective of real world MP mixtures.
- Health effects can be associated with different mixes.
- Various precursors, including those associated with climate change, can be used.
- Goal is to compare biological effects in animals exposed using two different types of chambers.



UNC Chapel Hill Chamber





Physicochemical Analyses

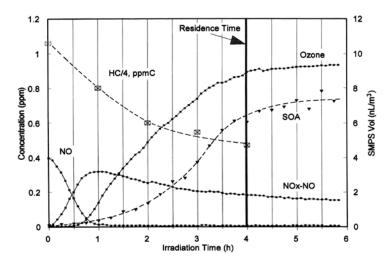
- PM size, mass, and morphology
- Inorganic elements
 - : XRF, ICP-OES, ICP-MS
- Organic components
 - : GC-MS
- Carbonaceous components
 - : Elemental and organic carbon, black carbon
- Reactive Oxygen Species
 - : EPR, reduction-oxidation potential
- Real-time PM analysis
 - : Aerosol Time-Of-Flight Mass Spectrometer (ATOFMS)
- Continuous emission monitoring for gaseous components



EPA Chamber







Biological Models

- Pulmonary inflammation
- Allergic potential
- Viral infectivity
- Atherosclerosis
- Hypertension
- Heart failure
- Mutagenicity

Project 4. Multi-City Studies

- Enhance results of epidemiology studies through the use of improved exposure metrics
- Multi-city Epidemiology Studies
- Multi-city Toxicology Studies



EPA Cooperative Agreement Research to Enhance Results of Epidemiology Studies Through the Use of Improved Exposure Metrics

- Numerous epidemiologic studies have used measurements from central-site ambient monitors to estimate exposures to air pollution
- Central-site monitors may not account for: spatial and temporal heterogeneity of urban air ambient pollution human activity patterns infiltration of ambient pollutants indoors contributions of indoor sources that may be effect modifiers
- Central-site are especially problematic for certain PM components and species (e.g., EC, OC, coarse, ultrafine) that exhibit significant spatial heterogeneity
- A number of enhanced exposure assessment approaches have recently been developed and applied in the investigation of air pollution health effects



Emory Coop

Develop and evaluate six exposure metrics for ambient traffic-related (CO, NO_x and $PM_{2.5}$ EC) and regional (O_3 and SO_4^{2-}) pollutants by applying them to two of ongoing epidemiologic studies on ambient air pollution and acute morbidity in Atlanta, GA

Rutgers Coop

Examine associations between PM_{2.5} mass and species and adverse health using two established epidemiology studies in New Jersey by testing four different tiers of exposure specification

University of Washington Coop

Improve air pollution cohort study health effect estimates by (i) incorporating dispersion model output into an existing spatiotemporal concentration model and (ii) quantifying the impact of exposure misspecification and methods for minimizing this source of error



Overall Goal: Enhance results of epi studies through the use of improved exposure metrics									
Collaborators	Pollutants	Exposure Approach	Health Study	EPA Role					
Turpin, Rich, Lunden (Rutgers/UMDNJ /LBNL	(1) PM2.5 (2) PM2.5 species: EC,OC,SO4, NO3,NH4	(1) Ambient Monitoring(2) SHEDS(3) Infiltration Model (LBNL)(4) SHEDS-Infiltration Hybrid	(1) NJ MI (2) NJ Birth Outcomes	(1) SHEDS (2) CMAQ					
S. Sarnat, J. Sarnat (Emory/Ga Tech)	(1) PM2.5 (2) PM2.5(EC) (3) CO (4) NOx (5) Ozone (6) SO4	 (1) Ambient Monitoring (2) Interpolation (3) CMAQ (4) CMAQ/Baseline+AERMO D (5) Air Exchange/Infiltration (6) SHEDS & APEX Exposure modeling 	(1) Atlanta ED (2) Atlanta ICD	(1) CMAQ(2) AERMOD(3) Data Fusion(4) SHEDS and APEX					
Sheppard, Sampson, Vedal, Larson (University of Washington)	(1) PM2.5 (2) NOx*	Assimilated Data from	(1) MESA-Air (2) WHI-OS*	(1) CMAQ** (2) AERMOD** (3) Data Fusion					



Multi-city Epidemiology Studies

Research Questions

- What metropolitan areas should have daily monitoring?
- What are the cause-specific concentration-response curves for specific PM-components / co-pollutants?
- Can built-environment and human behavior patterns explain heterogeneity of city-specific associations?
- Do concentration-response slopes for criteria pollutants and components vary over time? (criteria pollutants as indicators)
- What is the modifying effect of climate indices on pollutant mixtures and cause-specific mortality?
- Is SES a vulnerability factor?



Research in Progress

- Provide OAQPS with candidate list of metropolitan areas for enhanced daily speciation monitoring
- Standard time series methods associations of causespecific mortality with PM components / co-pollutant
- Innovative Bayesian methods of temporal-spatial models NCSU cooperative agreement
- Bivariate models for temperature / pollutants



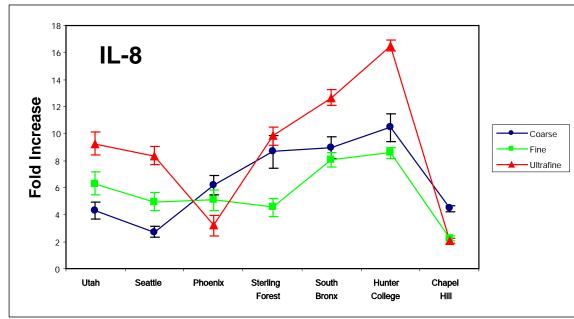
Map of selected STN monitor sites: 49 STN monitors, 2001-2005.



Multi-city Toxicology Studies

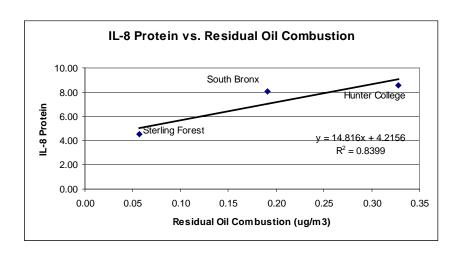
- UF, F, C particles were collected in 7 US (and 2 German) cities
- Cities were chosen because they have different source profiles
- Each size fraction was analyzed for multiple PM components
- Animal instillation and in vitro toxicology studies were used to link biological effects with particle size, components, and sources

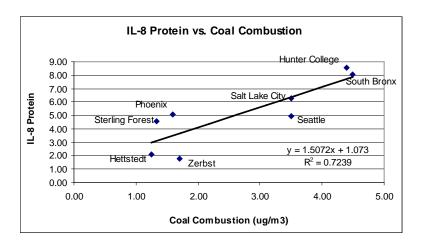




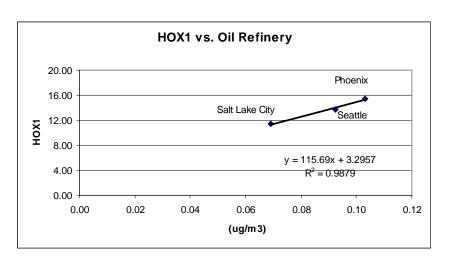


Association of Biological Changes to Sources in MAPS Cities





IL-8 protein is strongly correlated with coal combustion (r2=0.72) and residual oil combustion (r2=0.84)



HOX1 is strongly correlated with Oil refinery (r2=0.98)



Size fractionated samples have been collected in additional cities.





Project 5. Factors that Modify Health Effects to Multipollutants

- Retrospective panel study of subjects used in controlled exposure.
- Animal CV responses to pollutants.



Human Retrospective Study

 Hundreds of volunteers have been exposed to air pollutants in Chapel Hill over the past six years

Each person exposed to air and pollutant

A large number of clinical measurements were taken before and after each exposure

 Air pollutant levels have been retrieved for several days prior to each person's clinic visits

Regional monitors and rooftop monitors in Chapel Hill facility (for PM components)

 Associations are being made between ambient levels of pollutants and changes in clinical tests performed prior to air or pollutant exposure.

Can stratify the population on things such as age, gender, race, genetics, disease, medications, smoking, near road, etc.



Clinical Measurements for Retro Study

Pulmonary Function

Bronchoalveolar Lavage

- **FEV1**
- FVC
- DLCO

- Cell Differentials
- Soluble Mediators of cell injury and inflammation
- Epithelial Cells
- Macrophages
- Bronchial Biopsies

Vascular

- BAU
- CBC and differential
- Clotting/coagulation factors
- Cytokines

Cardiac

Holter monitoring
time domain
Frequency domain
Repolarization
Ectopic beats

Clinical

Medical History
Genotyping
Age, smoking, medications
Residence

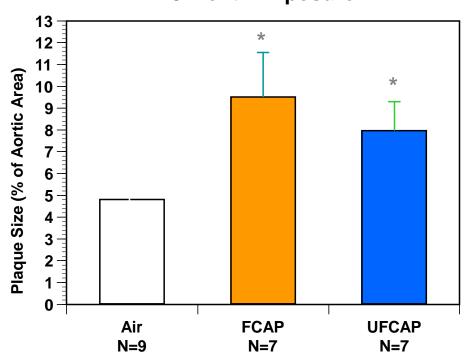


Vascular Effects of a Subchronic Inhalation Exposure to Concentrated Fine and Ultrafine Ambient Air Particles in Atherosclerosis Susceptible Mice

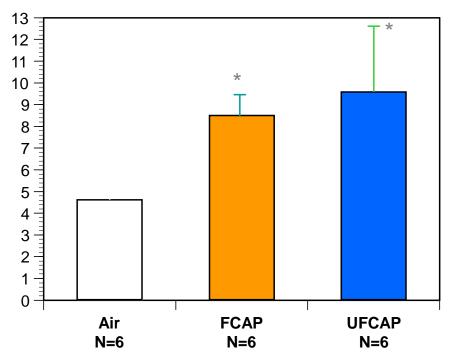




3 Month Exposure



3 Month Exposure/1 Month Recovery



* p<0.05 vs. Air



Acute Effects of Diesel Exhaust on Cardiovascular Function in Aged Heart Failure Rats

Pre-exposure
Ultrasound
(Echocardiogram)



4 h whole-body inhalation of diesel exhaust (500 μg/m³)



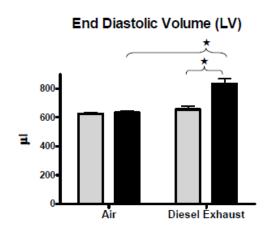
Post-exposure
Ultrasound
(Echocardiogram)

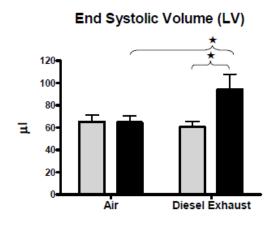


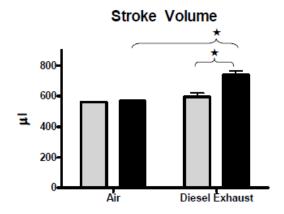




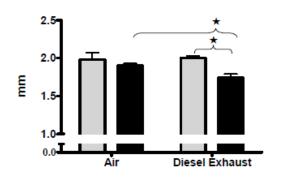
A Single Diesel Exhaust Exposure Increases Stroke Volume and Decreases Left Ventricular Wall Thickness



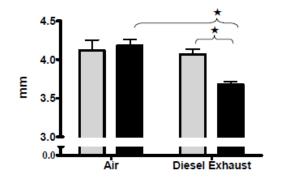


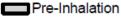


Posterior Wall Thickness - Diastole (LV)





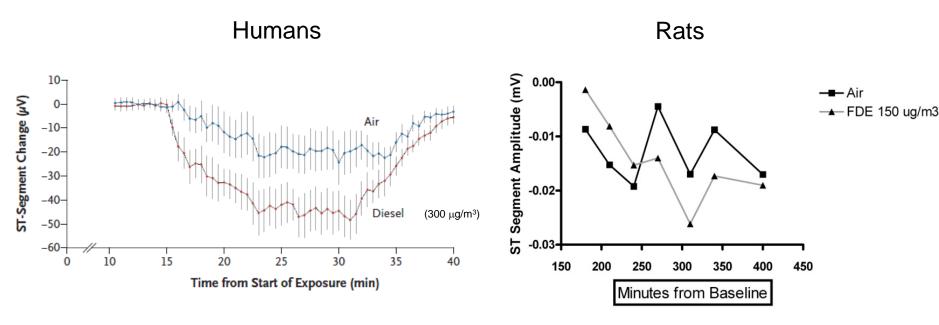




Post-Inhalation



Cardiac Ischemia during Diesel Exhaust Exposure in Humans and Rats at Comparable Concentrations



Mills et al, 2007

ST segment depression: an electrocardiographic indicator of cardiac ischemia



EPA's Air Pollution Program Provides A Science Foundation for Protecting the Air We Breathe

Air Pollution health and exposure research has significant benefits and impacts:

- Indicates that the current standards are necessary to protect public health
- Strengthens confidence in the scientific basis for these standards
- Provides information to inform people, particularly at risk populations, of the harmful effects of air pollutants and to educate them to minimize exposure to air pollutants

