

# OZONE

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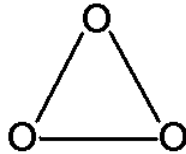
School of Public Health

PubH 242 - Toxicology

# Ozone Chemistry

- Formula –  $O_3$

- Structure –



- Synonyms – Triatomic Oxygen, Trioxygen
- Molecular Weight – 48.0 grams

(NTP, 2001)

# Ozone Chemistry (cont.)

- Gas (25° Celsius, Standard pressure)
  - Melting Point - -193° Celsius
  - Boiling Point - -112° Celsius
  - Vapor pressure: 1 mm Hg @ -180.4° C
- Slightly water soluble
- Unstable, highly reactive, potentially explosive  
(NTP, 2001)

# Sources of Ozone

- Naturally occurring stratospheric molecule
  - Created during thunder and lightning storms
  - Creation in the atmosphere by a catalyzed reaction of  $O_2$  and  $O$
  - Provides barrier to ultraviolet radiation
- Photochemical air pollution in the troposphere
  - Produced as an indirect by-product of combustion
- Commercially produced material
  - Produced by passing air through charged electrodes

(NTP, 2001)

# Common Routes of Exposure

- Exposure occurs by respiration of ozone rich air pollution.
  - Exposure to Ozone due to occupational exposure.
  - Route of exposure is via the respiratory tract.
    - Accumulation of  $O_3$  within the alveolar region of the lung
    - Volume reduced moderately by nasopharyngeal region
- (Costa, 2000; Vavra, 2000)

# Toxicokinetics

- Absorption
  - Some removal of volume due to nasopharyngeal scrubbing
    - Mucous absorption
- Distribution
  - Target organ: lung

(Santiago, 2000; Medinsky, 2000)

# Toxicokinetics (cont.)

- Metabolism
  - Ozone interferes with epithelial lining of the lung
    - $O_3$  seeks free electrons
    - Reacts with lipids in dissolved in lung fluids
    - Breaks down lipids imbedded in the cell membrane
    - Metabolites absorbed across the air-blood membrane
- Excretion
  - Exhaled during respiration

(Costa, 2000; Vavra, 2000)

# Ozone Interactions: Copollutants

- Multiplicative effect of ozone with other air contaminants
  - Reduced dose of air contaminants may suggest additive effect.
- $O_3$  and  $NO_2$
- $O_3$  and  $SO_2$
- $O_3$  and atmospheric acids

(Costa, 2000; Zeghnoun, 2001)

# Toxicity Thresholds

- NATIONAL AMBIENT AIR QUALITY STANDARDS
  - 1 hour (average): 0.12 parts per million (235  $\mu\text{g}/\text{m}^3$ )
  - 8 hour (average): 0.08 parts per million
- OSHA
  - 8 hour (average): PEL - 0.1ppm
  - 15 minute (average): STEL – 0.3 ppm

# Toxicity Thresholds (cont.)

- Acute Effect

(animal models and/or humans studies)

- > 0.01 ppm - odor threshold
  - < .2 ppm – reduced respiratory volume
  - > .25 ppm – coughing, breathing difficulty
  - > 1 ppm – general irritation of respiratory tract, vomiting, lethargy, decreased lung function.
  - > 2 ppm - pulmonary edema (mice)
- Increased mast cell proliferation

# Toxicity Thresholds (cont.)

- Chronic Effects
  - Generally all effects are considered reversible
  - Exposures have a cumulative effect
  - Formation of tolerance
    - Occurs after 2 to 4 days continual exposure

(Costa, 2000; Gong, 1998; Kleeberger, 2001)

# Potential Health Outcomes

- Induces immune response
  - Activation of cytokine-mediated response
  - Increased permeability of the lung epithelium
    - Greater probability of infection while in ozone rich environment
  - Increased likelihood of an asthmatic response
    - Potential allergen
  - Increases quantity of Polymorphonuclear cells
    - May reduce level of antigen required for allergic response

# Potential Health Outcomes (cont.)

- Reduction in Forced Air Volumes
- Cardiac stress
- Formation of precursor emphysema tissue
- Exacerbated lung dysfunction by hyperthyroidism

(Costa, 2000; Gong, 1998; Huffman, 2001)

# Appropriate Treatment

- Removal from area of exposure
- Anti-inflammatory drugs to reduce effects of immune response
- Antioxidants to reduce decrease in lung function

(Costa, 2000; Trenga, 2001)

# Biomarkers

- Lavage Fluids
  - Increased concentration of immune related cells
- Exhaled gases
  - NO
  - Ethane, Pentane

(Costa, 2000; Graham, 2000)

# Regulatory Issues

- Multiple cities are in violation of mandated air quality standard.
  - Increased number of hospital visits during periods of high Ozone concentrations.
    - Economic burden on society.
    - Health burden on society.
- Benefits of reduction of air pollutants.

(Costa, 2000; Cifuentes, 2001, Burnett, 2000)

# References

- Burnett, Richard T., et al.** Association between Ozone and Hospitalization for Acute Respiratory Diseases in Children Less than 2 Years of Age. *Am J Epidemiol* 153(5): 444-452, 2001.
- Cifuentes, L., et al.** Hidden Health Benefits of Greenhouse Gas Mitigation. *Science* 293: 1257-1259, 2001.
- Cohen, Mitchell D., et al.** Ozone-Induced Modulation of Cell-Mediated Immune Responses in the Lungs. *Toxicol Appl Pharmacol* 171: 71-84, 2001.
- Costa, Daniel L.** “Air Pollution”, Toxicology: The Study of Poisons. 6<sup>th</sup> ed. New York: McGraw-Hill, 2000.
- Delfino, Ralph J., et al.** Symptoms in Pediatric Asthmatics and Air Pollution: Differences in Effects by Symptom Severity, Anti-inflammatory Medication Use and Particulate Averaging Time. *Environ Health Perspect* 106(11): 751-761, 1998.
- Friser, T., et al.** Ambient ozone exposure is associated with eosinophil activation in healthy children. *Clin Exper Allergy* 31:1213-1219, 2001.
- Gong, Henry JR., et al.** Relationship between Acute Ozone Responsiveness and Chronic Loss of Lung Function in Residents of a High-Ozone Community. *Arch of Env Health* 53(5): 313-319, 1998.
- Gong, Henry Jr., et al.** Cardiovascular Effects of Ozone Exposure in Human Volunteers. *Am J Respir Crit Care Med* 158: 538-546, 1998.

# References (cont.)

- Graham, Regina M., et al.** Sensory Nerves Promote Ozone-Induced Lung Inflammation in Mice. *Am J Respir Crit Care Med* 164: 307-313, 2001.
- Holz, O., et al.** Flow cytometric analysis of lymphocyte subpopulations in bronchoalveolar lavage fluid after repeated ozone exposure. *Int Arch Occup Environ Health* 74:242-248, 2001.
- Huffman, L.J., et al.** Hyperthyroidism Increases the Risk of Ozone-Induced Lung Toxicity in Rats. *Toxicol Appl Pharmacol* 173: 18-26, 2001.
- Kleeberger, Steven R., et al.** Airway responses to chronic ozone exposure are partially mediated through mast cells. *J Appl Physiol* 90:713-723, 2001.
- Medinsky, Michele A. and Bond, James A.** Sites and mechanisms for uptake of gases and vapors in the respiratory tract. *Toxicology* 160:165-172, 2001.
- Olin, A.-C., et al.** Nitric oxide (NO) in exhaled air after experimental ozone exposure in humans. *Respiratory Med* 95:491-495, 2001.
- Petroeschovsky, Anna.** Associations between outdoor air pollution and hospital admissions in Brisbane, Australia. *Arch of Env Health* 56(1): 37-52, 2001.
- Santiago, Lizzie Y, et al.** Ozone absorption in the human nose during unidirectional airflow. *J Appl Physiol* 91:725-732, 2001.
- Shore, Stephanie A., et al.** Tumor Necrosis Factor Receptor 2 Contributes to Ozone-induced Airway Hyperresponsiveness in Mice. *Am J Respir Crit Care Med* 164:602-607, 2001.

# References (cont.)

- Trenga, C.A., et al.** Dietary Antioxidants and Ozone-Induced Bronchial Hyperresponsiveness in Adults with Asthma. *Arch of Env Health* 56(3): 242-249, 2001.
- Zeghnoun, A, et al.** Short-Term Effects of Air Pollution on Mortality in the Cities of Rouen and Le Havre, France, 1990-1995. *Arch of Env Health* 56(4): 327-336, 2001..