Chemical and Pathology Studies of Particulate Matter
A Medical Geology Perspective

Jose A. Centeno, PhD, FRSC
Email: jose.a.centeno@us.army.mil

Division of Biophysical Toxicology
The Joint Pathology Center – JTF CapMED
Washington, D.C.

International Symposium on Advances in Geospatial Technologies for Health
September 12-13, 2011
Santa Fe, NM
Disclaimers:

The opinions presented are the private views of the speaker, and should not be construed as official or representing the views of the Department of Army, the Department of Defense, the Joint Pathology Center, or other federal agencies.

Certain equipment and instruments or materials are identified in this presentation to adequately specify the experimental details. Such identification neither implies recommendation by the Department of the Army, nor that the materials are necessarily the best available for the purpose.
OVERVIEW

- Introduction

- From dust to disease: a review of toxicological pathways

- Health outcomes of bio-reactive dust-borne elements:
  ** Respiratory effects**
  ** Carcinogenic effects**
  ** Indirect health effects**

- Case Studies in Military Medical Geology
## Key Variables in Relation to Health Impacts of Mineral Dusts

<table>
<thead>
<tr>
<th>Environmental Mechanisms</th>
<th>Exposure Pathways</th>
<th>Progression to Health End-Points</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Anthropogenic</strong></td>
<td>Discrete / Defined</td>
<td>Well-established disease processes/ readily measurable health end-points</td>
</tr>
<tr>
<td><strong>Geogenic / Geochemical</strong></td>
<td>Diffuse / Mixed</td>
<td>Poorly-defined disease processes/ health end-points difficult to identify or measure</td>
</tr>
</tbody>
</table>
Including:

- Volcanoes
- Dust storms (regional storms)
- Long-range transport episodes of desert dust (intercontinental dust)
- Displacement through natural processes such as landslides and earthquakes
- Mine tailings spills
- Chemical or industrial spills
- Terrorist attacks

Dominant dust source regions around the world (in brown).

Courtesy of Prof. Dr. Edward Dervishire, UK and Dr. Geoffrey Plumlee, USGS
Annual volumes at over 1 billion tons from the main African deserts alone

Anthropogenic influences (such as from farming and irrigation) have contributed to dust formation by enhancing the process of desertification, with losses of over 10 million hectares (~29 million acres) of farmland per year.
Dispersal phenomena occur on all the major continents:

- mobilization of Saharan dust to southern Europe and the Americas
- inundation of Chinese cities and Japan with dust from central Asia
- regular dispersal in Australia and south-western parts of the US
Absorption of particulate materials from soil, food and water has been established as an additional avenue of exposure via the Lymphatic system (i.e. 1,2,3). In the example above microprobe mapping has enabled the analysis of microscopic Al-Si-Fe particles concentrated within human gut associated lymphoid tissue (i.e. Peyer’s patches). The presence of such particles have been associated with tissue damage, immune suppression and parallels have been drawn with lung pneumoconiosis.

1 Shepherd et al., 1987, Hum Pathol, 18, 50-54.
3 Ziegler, 1993, The Lancet, 342, 1348-1351

Sample courtesy of J.Powell, Rayne Institute, St Thomas’s Hospital, London
The process of inhalation of mineral aerosol particles leads to deposition in the pulmonary alveoli. *The process varies* with several factors, notably:

- mineral type (composition) and inclusions;
- dust particles size and shape:
  - $< 10\text{–}20\ \mu\text{m}$ (inhaled)
  - $< 2\ \mu\text{m}$ (respired)**
- Length of exposure;
- Certain lung and immune system functions.

**Atmospheric dust finer than 2.5 µm is of particular importance with respect to community health, as in the PM standard.**

*Pulmonary alveoli*: out-pouchings on the fine lining passages in which oxygen exchange with the blood stream occurs.

*The PM (particulate matter) standard* of the U.S. Environmental Protection Agency is based on the total mass of particles measuring 2.5 microns or less observed in a 24-hour period.
Particles <4 µm frequently penetrate more deeply into the lungs.

Prolonged exposure can lead to the pneumoconioses (including silicosis*, asbestosis+, and other lung conditions).

Many symptoms are non-specific in the absence of radiography.

[Crystalline silica is WHO listed as a human carcinogen]

*Silicosis: inflammation of the lung, leading to fibrosis, caused by foreign bodies, notably inhaled silica particles.

+Asbestosis: degenerative fibrosis of the lung resulting from chronic inhalation of asbestos fibers.
Airborne dust may contain asbestos, silica, chromium, nickel, arsenic, cadmium, well known carcinogenic for the respiratory system.

**Silica** is the most significant cause of dust-associated morbidity and mortality.

Airborne dust may absorb harmful gases, disease-generating bacteria and even carcinogenic hydrocarbon compounds (eg, benzo(a)pyrene).

Respiratory disease may exacerbate cardiac problems.

Another major group of toxins which may be contained within, or adsorbed to, dust particles are the **metals and metalloids**. Among the best characterized are aluminum (Al), lead (Pb), zinc (Zn), mercury (Hg), arsenic (As), chromium (Cr), cadmium (Cd), copper (Cu), and iron (Fe).
Exposure to Metals and Metalloids may produce a wide spectrum of respiratory effects, including:

- upper airway injury or sensitivity, such as rhinitis and sinusitis from arsenic and mercury;

- lower airway inflammation, including tracheitis, bronchitis and asthma from mercury, zinc and numerous other inhaled metals;

- acute inflammation, edema, and fibrosis of the lung parenchyma with inhalation of such elements as cadmium
Patterns of toxicity vary considerably with the forms, valences and formation ("species") of compounds:

- Although the process of lung injury has often been attributed to silicates, the action of divalent iron (Fe\(^{2+}\)) trapped on dust surfaces may play a role.

- In this reactive form, iron may result in catalyzing reactions and the production of free radicals (reactive oxygen species), leading the tissue injury.
A number of metals and metalloids, including cadmium, aluminum, and nickel, are classified as confirmed or suspected carcinogens using the IARC classification.

Arsenic, for example, is a group I carcinogen and has been linked to an extensive list of cancers, including those of the skin and most abdominal viscera, as well as angiosarcomas and hemato-lymphatic malignancies.
Force Health Protection: Integrating Environmental Earth Sciences and Human Health

Needs, Benefits and Military Relevance

Problems Related to “Place”:

Air: Sand & Dust Clouds
- Kuwait Oil Fires
- Desert Dust Storms

Water
- Arsenic, Mercury, Radon
- Lead, Uranium

Soil
- Leishmaniasis
- Valley Fever
Pulmonary Sarcoidosis

Definition: multisystem granulomatous disease of unknown etiology.

Possible association with exposure to silicates and toxic metals (Al, Pb, Hg).

Health Effects of Middle East Sand (Dust)
A Medical Geology Research Case


- Obstructive bronchitis and bronchiolitis in 86 autopsied casualties from Kuwait, with observation of sand particle in lung parenchyma. (NS Irey, Kuwait Casualties: Morphologic and Toxicologic Findings, NIH Technical Assessment Statement, April 27-29, 1994).

- From March through August 2003, 19 US military personnel developed pneumonia severe enough to warrant medical evacuation and mechanical ventilation; two died. (AF Shorr, et al., Acute eosinophilic pneumonia among US military personnel deployed in or near Iraq. JAMA. 2004 Dec 22;292(24):2997-3005.)

Health Effects of Dusts

Military Medical Geology

Operation Desert Storm (April 1991)
Health Impact of the Kuwait Oil Fires

AFIP Studies on:
1. Animal Tissues (Kuwait City vs Ahmadi)
   a. Histologically – anthracotic pigment
   b. Analysis of Toxic Metals-
      Ni, V, Cr, Cd, Pb, Cu, Zn, Mn

2. Biological Surveillance
   a. Epidemiology, clinical assessment, chemical toxicology

Results: Inhalation of air contaminated with smoke from the oil fires had little or no long-term effects.

Chemical and Microscopy (Particle Size) Analysis of Middle East sand (dust)

<table>
<thead>
<tr>
<th># particles</th>
<th>123</th>
</tr>
</thead>
<tbody>
<tr>
<td>average (µm)</td>
<td>6.4</td>
</tr>
<tr>
<td>median (µm)</td>
<td>4.3</td>
</tr>
<tr>
<td>max (µm)</td>
<td>46.1</td>
</tr>
<tr>
<td>min (µm)</td>
<td>1.1</td>
</tr>
</tbody>
</table>

Other metals detected: Al, As, Cd, Co, Ni, Cr, Mn, Pb

Lyles M, Centeno JA et al. unpublished results
### Links between selected elements and some known lung function conditions and diseases

<table>
<thead>
<tr>
<th>Element</th>
<th>Desert Dust (&lt;10 µm)</th>
<th>Desert Dust (20-40 µm)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mn (ppm)</td>
<td>450</td>
<td>331.98</td>
</tr>
<tr>
<td>Fe (ppm)</td>
<td>25500</td>
<td>18111.61</td>
</tr>
<tr>
<td>Co (ppm)</td>
<td>11.72</td>
<td>8.24</td>
</tr>
<tr>
<td>Pb (ppm)</td>
<td>17.22</td>
<td>9.45</td>
</tr>
<tr>
<td>Cu (ppm)</td>
<td>220</td>
<td>152.64</td>
</tr>
<tr>
<td>Cd (ppm)</td>
<td>1.24</td>
<td>0.70</td>
</tr>
<tr>
<td>Mg (ppm)</td>
<td>13230.49</td>
<td>10572.70</td>
</tr>
<tr>
<td>Al (ppm)</td>
<td>15912.39</td>
<td>13154.60</td>
</tr>
<tr>
<td>Ca (ppm)</td>
<td>139577.64</td>
<td>140250.15</td>
</tr>
<tr>
<td>Na (ppm)</td>
<td>1098.28</td>
<td>1476.86</td>
</tr>
<tr>
<td>Cr (ppm) [but species critical]</td>
<td>181.32</td>
<td>187.36</td>
</tr>
<tr>
<td>Zn (ppm)</td>
<td>105.18</td>
<td>72.30</td>
</tr>
<tr>
<td>Ni (ppm)</td>
<td>93.28</td>
<td>60.44</td>
</tr>
<tr>
<td>Ti (ppm)</td>
<td>1095.52</td>
<td>539.81</td>
</tr>
</tbody>
</table>

- **Cancer**
- **Cancer suspected**
- **Cancer & asthma**
- **Emphysema**
- **Asthma**
SUMMARY

• Dust and other particulates ("Dust" is used here to cover dust and other finer particulates) from both natural and anthropogenic sources have major implications for human health, agriculture and livestock and the natural environment.

• Several studies have demonstrated that fine particles may contain relatively high concentrations of transition metals implicated to catalyzed the formation of oxygen radical species, increasing the oxidative stress burden and tissue damage.

• Substantial research has been undertaken on aspects of this subject, but an integrated understanding of these materials from an inter-disciplinary point of view is still lacking. It is important for risk assessment studies, quantifying the public health impact of fine-particulate exposure using a multi-disciplinary approach.

Photo by David Fitzpatrick, NYPD