#### EE-527: MicroFabrication

Toxicity of Materials

R. B. Darling / EE-527

# Toxicity

- A toxic substance or poison is a material that will produce serious disease of death upon introduction into a normal healthy person's bloodstream.
- Unique susceptibilities or allergies are not included.
- Acids and bases are not strictly poisons, even though they can cause local destructive effects on tissue severe enough to be fatal.

# Types of Poisons

- <u>Cytotoxins</u>: injure all cells that they contact, usually by destroying the ion permeability of the cell membrane
- <u>Neurotoxins</u>: injure the central nervous system, usually the ability of neurotransmitters to regulate the Na<sup>+</sup> and K<sup>+</sup> membrane channels
- <u>Hemotoxins</u>: injure the circulatory system, usually the ability of hemoglobin to carry oxygen

### Factors Affecting Toxicity

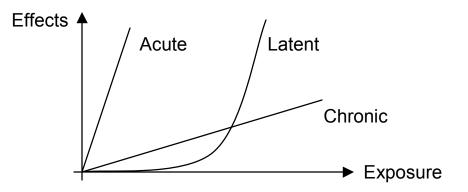
- Quantity of material: generally less than 1/8 ounce
- Rate and extent of absorption into the bloodstream
  - routes of exposure:
    - (1) inhalation-- most common for adults
    - (2) oral-ingestion-- most common for children
    - (3) cutaneous
    - (4) subcutaneous
    - (5) intravenous
    - (6) intramuscular
    - (7) intraperitonial-- puncture wounds to the gut sac
- Rate and extent the material is broken down by the body
- Rate and extent the material is excreted by the body

## Exposure by Inhalation

- Respiratory system is most potentially hazardous route of intake of poisoning
  - asphyxiants- reduce or eliminate oxygen uptake
  - irritants- mucus, nasal, skin
  - anesthetics- loss of consciousness
  - lacrymators- cause tearing, gagging
- Small volumes within the aveoli and bronchi tend to trap aerosol particles in the micron size range
- Tidal volume of lungs is small compared to stagnant volume- a long time is required to fully exchange
- Strong irritants can cause swelling which closes passageways and produces asphyxiation

### Types of Poisoning Effects

- All materials are toxic in sufficient quantity:
  - 5g of caffiene in a single dose is usually fatal to an adult
  - many people drown in water
- Effects versus exposure:
  - acute: short term; immediately obvious
  - chronic: long term; requires repeated exposures
  - latent: requires an incubation period
    - Example: benzene poisoning: delayed onset of aplastic anemia



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- Lethal Dose, 50 % Kill  $(LD_{50})$ 
  - amount of material which kills 50 % of laboratory animals
  - expressed as mg of poison per kg of animal weight
  - Example: asprin:  $LD_{50} = 1750 \text{ mg/kg}$ 
    - 100 kg person will have 50 % chance of death consuming 175 grams
- Lethal Concentration, 50 % Kill (LC<sub>50</sub>)
  - concentration of material in ppm by volume that kills 50 % of laboratory animals during exposure period
- Lethal Dose, Low (LD<sub>LO</sub>)
  - smallest dose ever reported to have caused a fatality
- Lethal Concentration, Low (LC<sub>LO</sub>)
  - smallest concentration ever reported to have caused a fatality

- Threshold Limit Value, (TLV)
  - upper limit of material concentration that an average healthy person can be exposed to on a continual every day basis without adverse effects
  - expressed as ppm for gases in air
  - expressed as  $mg/m^3$  or  $\mu g/m^3$  for fumes or mists in air
  - recommendations set by American Conference of Governmental Industrial Hygienists (ACGIH)

- Example: carbon monoxide, CO
  - hemotoxin; combines with hemoglobin 300X more readily than  $O_2$
  - TLV = 100 ppm; body can tolerate 0.01 % in air
  - @ 1000 ppm (0.1 %) causes headache and nausea
  - @ 10,000 ppm (1.0 %) fatal to adults in 1 min.
- Example: carbon dioxide, CO<sub>2</sub>
  - TLV = 5,000 ppm (0.5 %); present atmosphere is 320 ppm
- Example: hydrogen cyanide, HCN
  - TLV = 10 ppm; 0.2-5.0 ppm is the odor threshold (almond smell)
  - @ 100 ppm, causes death in 1 hr.
  - @ 180 ppm, causes death in 10 mins.
  - @ 280 ppm, immediately fatal

- Time-Weighted Average TLV, (TWA-TLV)
  - This is a TLV for an 8 hour day, 40 hour work week exposure
- Immediately Dangerous to Life and Health, (IDLH)
  - Defined by OSHA and NIOSH
  - Maximum concentration from which one could escape within 30 minutes without adverse health effects

#### Notorious Poisons

– carbon monoxide CO TLV = 100 ppmTLV = 5000 ppm– carbon dioxide  $CO_2$ TLV = 10 ppm– hydrogen cyanide HCN hydrogen sulfide  $H_2S$ TLV = 10 ppmTLV = 5 ppm $SO_2$ – sulfur dioxide  $N_2O$ used for anesthetic purposes – nitrous oxide - nitric oxide TLV = 25 ppmNO  $NO_2$ – nitrogen dioxide TLV = 5 ppmTLV = 50 ppmNH<sub>3</sub> ammonia  $As_2O_3$  0.1 g usually fatal – arsenic trioxide  $Pb(C_2H_5)_4$  $0.075 \text{ mg/m}^3$  via skin – tetraethyl lead – liquid mercury Hg TLV = 0.1 ppm via skin contact

#### Changes in Attitudes

- 1970s 1980s: "Solution to polution is dilution"
- 1990s: "Reduce, reuse, recycle"
- Minimata Bay, Japan: methyl mercury poisoning
  - thousands of people affected: poor health, shortened lifespans, serious birth defects
  - dilution rendered ineffective because marine life re-concentrates the waste materials