# 4th year Emergency Medicine Didactic Series: Common Toxicologic Emergencies

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### **OBJECTIVES**

- DISCUSS TOXIDROMES
- DISCUSS DECONTAMINATION METHODS
- DISCUSS THE WORKUP OF AN OVERDOSED PATIENT
- DISCUSS THE COMMONLY USED ANTIDOTES
- REVIEW SOME OF THE COMMONLY SEEN DRUG OVERDOSES IN A COMMUNITY HOSPITAL

## **OVERDOSES**

Approximately 5 million annually.

Less than 1% mortality.

Mean time to presentation is 3-4 hours.

SUPPORTIVE CARE

Educational opportunities for patients

## **TOXIDROMES**

 constellation of signs and symptoms that suggest a specific class of poisoning

## **OPIODS**

- Morphine
- Meperidine
- Propoxyphene
- Mixed agonist/antagonist
- Synthetic opiates (fentanyl)
- Drug combinations: Lortab, Percocet, Vicodin
- Heroin

## **OPIOIDS**

- CNS Depression
- Respiratory Depression
- Miosis
- Seizures
- Nausea & Vomiting
- Hypotension and Bradycardia

## Sympathomimetic Agents

- Drugs of abuse
- Adrenergic agonists
- Amphetamines
- Cocaine
- PCP
- TCA
- Early MAOI
- Sedative/hypnotic withdrawal

## SYMPATHOMIMETIC

- **Hypertension**
- Tachycardia
- Mydriasis
- **CNS Excitement**
- Hyperthermia
- Diaphoresis

## CHOLINERGIC AGENTS

- Organophosphates
- Carbamates
- Insecticides
- Nerve agents

## CHOLINERGIC

#### **DUMBELLS/SLUDGE/SLUG BAM**

- Salivation/Sweating
- Lacrimation
- Urination
- GI complaints (N/V/D)
- Bradycardia/Bronchoconstriction
- Abdominal cramping
- Miosis/Muscle fasciculations

## Anticholinergic Agents

- TCA
- Scopolamine
- Neuroleptics
- Parkinson meds
- Benztropine mesylate (Cogentin)/Trihexyphenidyl (Artane, Apo-Trihex),
- Antihistamines: Benadryl
- Gyromitra
- Jimson weed

## ANTICHOLINERGIC

- Hot as Haiti (hyperthermia)
- Red as a beet (flushed skin)
- Dry as a bone (dry skin, urinary retention)
- Blind as a bat (mydriasis)
- Mad as a hatter (delirium, hallucinations, myoclonic jerking)

## SALICYLATES

#### **ASPIRIN**

- Altered mental status
- Sweating
- Pulmonary edema
- Irritable
- Ringing in ears (tinnitus)
- Increased temp., breathing, and heart rate
- Nausea and vomiting

## PATIENT WORK UPS

## Do the **DONT**

- Dextrose
- Oxygen
- Naloxone
- Thiamine

## What type of work up needs to be done?

## **ESSENTIAL WORK UP**

- ELECTROLYTES
- EKG
- ACETAMINOPHEN LEVEL
- CARDIAC MONITORING

## ADDITIONAL WORK UP

- ALCOHOL LEVEL
- SALICYLATE LEVEL
- PREGNANCY TEST
- TOXIC ALCOHOL LEVELS
- URINE DRUG SCREEN

## URINE TOX SCREEN

- TCA FALSE POSITIVES
  - -- Cyclobenzaprine (Flexeril)
  - -- Benadryl
  - -- Quetiapine fumarate (Seroquel)
  - -- Carbamazepine (Tegretol)
  - -- Cyprohepatidine
  - -- Hydroxyzine (Vistaril)

## ANION GAP METABOLIC ACIDOSIS

#### A MUD PILE CAT

- ◆ Aspirin
- Methanol
- Uremia
- ◆ Diabetic Ketoacidosis
- ◆ Paraldehyde, Phenformin
- ◆ INH, Iron, Ibuprofen
- ◆ Lactic acidosis
- Ethylene glycol

- ◆ CO, CN, Caffeine
- ◆ Alcoholic ketoacidosis
- ◆ Theophylline, Toluene

## DECONTAMINATION TECHNIQUES

## **IPECAC**

Only for very LIMITED home use.

Not for use in the emergency department.

Do not use on caustic substances

 Less effective than activated charcoal and equivalent to gastric lavage.

## Gastric Lavage

Frequently used in the past, now very infrequently used.

Only to be used within 30-60 minutes of ingestion

Only to be used on LIFE-THREATENING ingestions.

## Gastric Lavage

- Decreases absorption of toxin ~26% @ 30 minutes
   vs. 12 % @ 60 minutes
- ? Airway protection
- Time consuming
- ? Promotes absorption of toxins

## ACTIVATED CHARCOAL

- Dose is 1 gram/kg
- Aqueous charcoal vs. charcoal with sorbitol
- Recommendation to use within 1 hour of ingestion
- May use later if the patient has ingested a substance that slows gut motility.
- Mean drug bioavailability is reduced by 69% at 30 minutes vs. 34% at 60 minutes

### **ACTIVATED CHARCOAL**

#### DOES NOT WORK FOR THESE INGESTIONS:

Heavy Metals	<u>Inorganic Ions</u>	<u>Hydrocarbons</u>
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Arsenic Lithium Alkanes

Lead Na/K/Mg Alkenes

Mercury Fluoride Alkyl halides

Iron Iodide Aromatic hydrocarbons

Zinc

Cadmium <u>Corrosives</u> <u>Essential Oils</u>

Acids/alkali

<u>Alcohols</u>

## ACTIVATED CHARCOAL

- Be aware of the contraindications.
  - Doesn't work for some meds
  - Needs to be given early
  - Do NOT give to someone who is at risk for aspiration

## MULTI-DOSE CHARCOAL

- INDICATIONS:
  - -- Theophylline
  - -- Phenobarbital
  - -- Carbamazepine (Tegretol)
  - -- Digitalis
  - -- TCA

## WHOLE BOWEL IRRIGATION

- Polyethylene glycol (GoLytely, MiraLax) 2 L/hr until clear rectal effluent
- Nasogastric tube
- Drugs (Packers vs. Stuffers)
- Sustained released or enteric coated drugs
- Toxins not well adsorbed by AC (e.g., lithium or other heavy metals)

## Acetaminophen

## 

- Most commonly used drug
- Combined in over 100 different drugs
- Majority absorbed within 2 hours
- Peak plasma levels around 4 hours

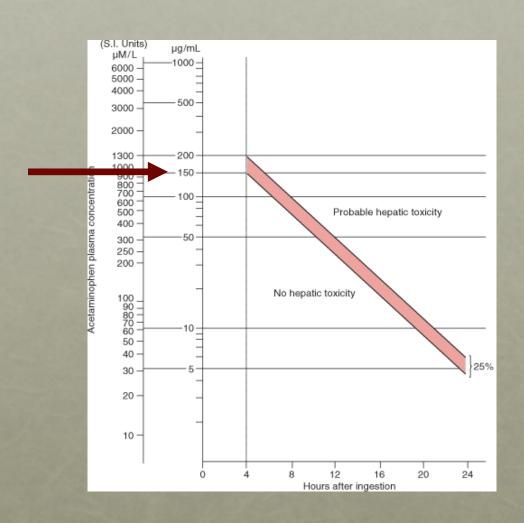
#### Knowledge of APAP Containing Products

- 71.2% recognized brand Tylenol as having acetaminophen
- 50% recognized Tylenol PM
- Fewer than 15% recognized Vicodin, Darvocet, Tylox,
   Percocet, and Lorcet as containing APAP
- 6.7 to 19.2% indicated that Motrin, ibuprofen, Sudafed, Aleve, and Benadryl contained APAP

Estimates of Acetaminophen Associated Overdoses in the United States

- 56,000 emergency room visits
- 26,000 hospitalizations
- 127,000 exposures reported
- 458 deaths of which 100 were unintentional
- Doubling of fatalities from 1997 to 2001
- Depending on the data source, 8 to 26% of overdoses were unintentional

## RUMACK-MATTHEW NOMOGRAM



• Therapeutic dose: 10-15 mg/kg

Toxic dose: <u>150</u> mg/kg

## STAGES OF TOXICITY

#### **STAGE ONE**

- -- 0.5-24 HOURS
- -- Malaise
- -- N/V
- -- Pallor
- -- Lethargy
- -- Diaphoresis
- -- Lab tests are normal

#### **STAGE TWO**

- -- 24 to 72 hours
- -- RUQ pain
- -- 1 LFT's
- -- 🋧 PT, Tbili
- -- Oliguria and renal dysfunction

\*\* AST is the first sign of hepatotoxicity \*\*

### STAGES OF TOXICITY

#### **STAGE THREE**

- -- 72-96 hours
- -- LFT's peak
- -- Stage one symptoms reappear.
- -- Severe hepatotoxicity =
  ALT/AST > 10,000, elevated
  PT/INR, low glucose, lactic
  acidosis, Tbili > 4.0
- -- DEATH is most common in this stage.

#### **STAGE FOUR**

- -- 4 days to 2 weeks
- -- If survival, liver function usually returns to normal

# Acetaminophen Toxicity

#### • TREATMENT:

- -- N-Acetylcysteine (NAC)
  - Oral or IV preparations
- -- Activated Charcoal

\*\* Treatment is 100% effective if given in <8 hours \*\*

# Acetaminophen Toxicity

#### • TREATMENT:

 Oral Regimen – 140 mg/kg load, then 70 mg/kg every 4 hours times 17 doses

IV Regimen – 150 mg/kg in 250 ml of D5W over 30-45 minutes, then 50 mg/kg in 500 ml of D5W over 4 hours, then 100 mg/kg in 1000 ml of D5W over 16 hours

\* Be cautious of fluid volume in pediatrics.

# Chronic Ingestions

- Start N-Acetylcysteine (NAC)
- Check Acetaminophen level, LFTs, Coags
- Repeat levels in 4 hours to determine half life.
- Plot two or three levels on the nomogram to determine toxicity
- Check acetaminophen level and LFTs
  - -- If not detectable and no abnormalities = OK

# **ASPIRIN**

#### **ASPIRIN**

- Altered mental status
- Sweating
- Pulmonary edema
- Irritable
- Ringing in ears (tinnitus)
- Increased temp., breathing, and heart rate
- Nausea and vomiting

## SALICYLATES/ASPIRIN

- Alka-Seltzer
- Anacin
- Excedrin
- Pepto-Bismol
- Oil of Wintergreen

- Therapeutic levels 10-30 mg/dL
- Mild toxicity 40-50
- Severe toxicity > 100

- Peak blood levels occur within one hour of therapeutic doses. Greater than 6 hours in overdoses.
- May result in pylorospasm, bezoars. Caution with enteric coated and extended release meds.

- Tachypnea stimulates medullary resp. centers
- Hyperthermia uncouples oxidative phosphorylation
- Tachycardia hypovolemia, agitation, distress
- Tinnitus seen at levels around 20mg/dL
- Acid-base abnormalities respiratory alkalosis or mixed respiratory alkalosis/metabolic acidosis

- AMS 1. Direct toxicity of salicylate in CNS
  - 2. Cerebral edema
  - 3. Low CNS glucose levels

#### TREATMENT

DONE Nomogram = Don't Use!!

- ABCs Do not intubate!
- Activated Charcoal
- Alkalinize the urine (NaHCO3 gtt)
- Hemodialysis Call Nephrology early!
- Check labs (K<sup>+</sup> and ABG) and levels every 2 hours until declining.

# Indications for Dialysis

- Altered Mental Status
- Pulmonary or Cerebral Edema
- Renal insufficiency
- Fluid overload that prevent NaHCO<sub>3</sub>- administration
- Level > 100 mg/dL
- Clinical deterioration despite treatment.

# Opioids

## **OPIOIDS**

- CNS Depression
- Respiratory Depression
- Miosis
- Seizures
- Nausea & Vomiting
- Hypotension and Bradycardia

## **OPIODS**

- Morphine
- Meperidine
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- Drug combinations
- Heroin

#### **OPIOIDS**

- Naloxone (Narcan)
  - -- Repeat doses every 20-60 minutes prn
  - -- May need drip. Start at 2/3 initial successful bolus
  - -- Need high doses for propoxyphene, methadone and fentanyl ~ 10-20 mg
  - -- If no re-dosing after 4 hours, then may disposition patient

# Benzodiazepines

# Benzodiazepines

- Intubation and supportive care
- Flumazenil
  - -- Use with extreme caution
  - -- May precipitate life threatening withdrawal and seizures

## Calcium Channel Blockers

## Calcium Channel Blockers

- Dihydropyridines
  - Nifedipine, amlodipine, felodipine, nicardipine
  - Potent vasodilators and minor effect on contractility and conduction

- Non-dihydropyridines
- -Verapamil and diltiazem
- -Depressive effect on contractility and conduction, but minor effects of vasodilation

## Calcium Channel Blockers

- Hypotension is seen with both types
- Bradycardia is usually only seen with verapamil and diltiazem

#### Treatment

- Gastric Lavage
- Activated Charcoal
- Whole Bowel Irrigation
- IV Calcium
- Glucagon
- Vasopressor = Levophed
- \*\* INSULIN \*\*

# BETA BLOCKERS

#### BETA BLOCKERS

- Most are symptomatic within 2 hours, all by 6 hours
- Use caution with sustained release products and sotalol
- EKG Changes
  - -- Decreased AV node conduction = PR prolongation
  - --Slowed automaticity in SA node = bradycardia
  - -- QRS prolongation
  - -- QTc prolongation with sotalol toxicity

## BETA BLOCKERS

- HYPOTENSION and BRADYCARDIA
- Hypoglycemia
- Seizures
- Arrhythmias

## TREATMENT

- Activated charcoal
- Whole bowel irrigation
- Gastric lavage

### **TREATMENT**

- Fluids for hypotension
- NaHCO<sub>3</sub><sup>-</sup> and Magnesium for arrhythmias
- Glucagon is considered first line treatment
  - -- 5 mg bolus, if no effect repeat in 10 minutes, then start drip at 2-5 mg/hour
- **◆**Calcium
- ◆Epinephrine drip poor outcomes, last resort
- **♦**HIGH DOSE INSULIN THERAPY

# HIGH DOSE INSULIN THERAPY

- One amp of D50
- Regular insulin bolus of 1-2 units/kg over 5 minutes
- Regular insulin infusion of 0.5 U/kg/hr, with a goal rate of 2 units/kg/hr

Frequent monitoring of glucose and potassium

#### Alcohol

- There are different kinds of alcohol so find out exactly what was taken
  - don't be fooled: Joose, mouthwash, colognes and perfumes, and OTC meds
- Do a good history and a great physical exam
- Be cautious in assuming altered mental status is secondary to alcohol
  - consider the possibility of a head injury and other causes

#### Alcohol

- Pathophysiology
  - metabolized at about 20 mg/dL per hour
  - decreases gluconeogenesis so be wary of hypoglycemia (do the DON'T)
  - patient's may be malnourished and have a low thiamine level (do the DON'T)
- Blood alcohol levels are rarely useful and correlate poorly with the clinical examination

#### Alcohol

#### Disposition

- Offer counseling or referral resources for alcohol abuse.
- If alert, oriented x 3, appropriately conversant and able to walk with a steady gait and if considered clinically sober => may be d/c'd from the ED under the care of a responsible adult
- If intoxicated with an abnormal mental status or neurologic exam, the patient should not be allowed to leave the ED, even if they want to.
  - place them on a hold as they are danger to themselves

#### Alcohol withdrawal

- Clinically
  - can occur at any level of blood alcohol
  - the hallmark is CNS excitation
  - usually begins 24 hours after reduction in EtOH intake
  - lasts 2-7 days

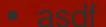
#### **Alcohol Withdrawal**

#### Mild

- occurs 6-24 hours after decrease in consumption
- peaks at 24-36 hours
- mild autonomic hyperactivity: nausea, anorexia, tremor, tachycardia, HTN, anxiety, sleep disturbance

#### Major

- occurs more than 24 hours after decrease
- peaks at 50 hours
- same symptoms as above, but more severe. Also decreased seizure threshold, auditory and visual hallucinations, and delirium



#### **Delirium Tremens**

- the extreme end of the EtOH withdrawal spectrum
- tremor, hallucinations, profound confusion, agitation, and hyperadrenergic syndrome (fever, tachycardia, and HTN)

# Treatment of EtOH withdrawal

- Supportive care: IVF, MVI, thiamine, folate, Mg
- Medications:
  - First line: benzos (lorazepam, diazepam)
    - Ativan 1-5 mg IV q 5-15 minutes prn agitation/tremor
  - Second line: butyrophenones (haloperidol)
    - Haldol 5-10 mg IV for agitation
  - BE AGGRESSIVE WITH MEDS
    - don't be afraid of large doses (there are reported cases of patients needing 480 mg of lorazepam and 240 mg of haloperidol in 24 hours)

## SUMMARY

- Toxidromes are clues to the mystery.
- Ipecac = NO
- Flumazenil = No
- Gastric Lavage = Probably no
- Whole Bowel Irrigation = Maybe
- Activated Charcoal = Probably
- NAC = YES
- Naloxone = Yes

# Summary

Acetaminophen = NAC (150)

Aspirin = Bicarb + Nephrologist

• Opioids = Narcan

Benzo's = Intubation

Ca<sup>++</sup> Channel and Beta Blockers = High Dose Insulin

#### Poison Control Center

• CALL THEM!!!!

•1-800-222-1222

# That's it!

Thanks.

Contact me for any questions or concerns.

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