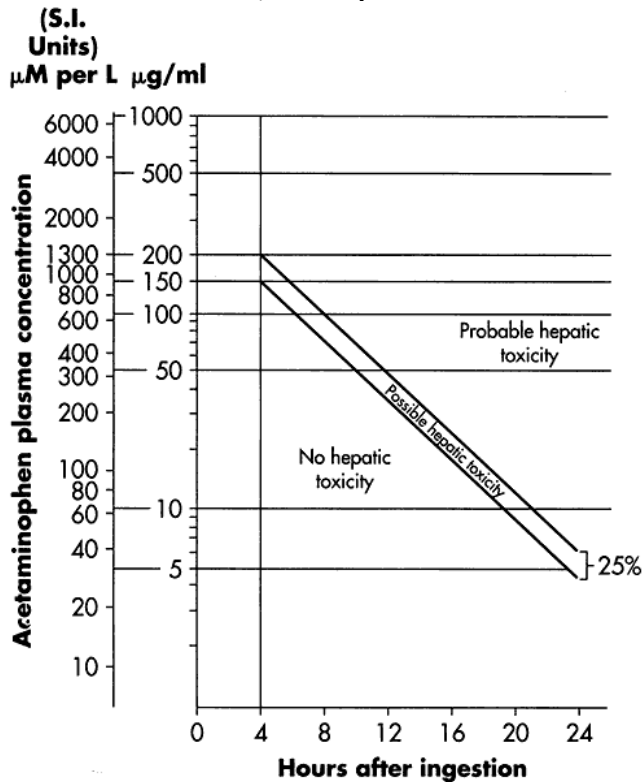


Tylenol (Acetaminophen) Ingestion

California Poison Control 1-800-876-4766

Toxicity

1. > 150 mg/kg is considered toxic and the dosage which N-acetylcysteine (mucomyst) therapy is initiated if the plasma acetaminophen level is unavailable
2. Rumack-Matthew nomogram for the single acute acetaminophen poisoning. Semilogarithmic plot of plasma acetaminophen levels versus time. Cautions for use of this chart: (1) Time coordinates refer to time of ingestion. (2) Serum levels drawn before 4 hours may not represent peak levels. (3) The graph should be used only in relation to a single acute ingestion. (4) The lower solid line 25% below the standard nomogram is included to allow for possible errors in acetaminophen plasma assays and estimated time from ingestion of an overdose. (Adapted from Rumack BH, Matthew H: Pediatrics 55:871-876, 1975.)



Pathogenesis

1. Hepatocellular Damage
 - a. Acetaminophen is metabolized by cytochrome P450 and glutathione in the liver to a mercapturic acid conjugate
 - b. With an overdose, the hepatic stores of glutathione are depleted to <70% of normal resulting in toxic damage by a highly reactive intermediate from the acetaminophen metabolic pathway

Clinical Features

1. Day 1: anorexia, diaphoresis, lethargy, malaise, nausea & vomiting, pallor
2. Day 2: (Day 1 symptoms disappear) Hepatic necrosis begins:
 - a. Abdominal pain and tenderness, hepatomegaly
 - b. Elevated AST/SGOT, ALT/SGPT, bilirubin, PT
3. Day 3 – 4: (Day 1 symptoms reappear) Hepatic necrosis peaks:
 - a. Jaundice, encephalopathy, acute renal failure, bleeding, hypoglycemia
4. After Day 4: resolution of symptoms and hepatic dysfunction
 - a. Fatalities from: ARDS, cerebral edema, coagulopathy, infection, multiorgan failure

Investigations

1. Serum
 - a. Liver function tests (Aspartate Aminotransferase (AST)/Serum Glutamic Oxalacetic Transaminase (SGOT), Alanine Aminotransferase(ALT)/Serum Glutamic Pyruvic Transaminase(SGPT), bilirubin), glucose, BUN, Creatinine, PT daily if acetaminophen levels are in the toxic range
 - b. Drug screen (for other toxins)
2. Urine
 - a. Drug screen (for other toxins)

Management

1. California Poison Control 1-800-876-4766
2. Initial management
 - a. Airway, Breathing, Circulation
 - i. If hemodynamically unstable Normal Saline or Lactated Ringers at 10-20 cc/kg IV
 - b. Draw Blood: liver function tests, PT, glucose, BUN, Creatinine
 - c. Acetaminophen level (if >4 hours post ingestion)
3. Gastric Lavage
 - a. Insert a large bore NG tube and check position
 - b. Suction out stomach contents and save for analysis
 - c. Place patient on side
 - d. Inject 15 cc/kg of saline per lavage
 - e. Contraindications: unprotected airway, coma, convulsions
4. Activated Charcoal
 - a. 1 g/kg
 - b. Not recommended if using oral N-Acetylcysteine

Maintenance Therapy

1. N-Acetylcysteine (Mucomyst) used for Oral Administration
 - b. Begin within 10 hours of ingestion if possible but may be used as late as 24 hours post ingestion
 - c. Indicated if plasma acetaminophen level is in the toxic range or if the level is not available, the ingested dose is >150 mg/kg
 - d. Administer orally or via NG tube

- e. Loading dose: 140 mg/kg/dose PO diluted in 3 volumes of soft drink
 - f. Maintenance dose: 70 mg/kg/dose PO q4h x 17 doses (for 3 days)
2. N-Acetylcysteine (Acetadote) used for Intravenous Administration
- a. Loading Dose
 - i. Dose 1: 150mg/kg over 30 minutes
 - b. Maintenance Dose
 - i. Dose 2: 50mg/kg (12.5 mg/kg/hr) over 4 hours
 - ii. Dose 3: 100mg/kg (6.25 mg/kg/hr over 16 hours)
 - c. Patients <40 kg and those requiring fluid restriction
 - i. The final concentration should be 40 mg/ml
 - d. Patients >40 kg
 - i. Loading Dose = Dose 1: in 200 ml 5% Dextrose (800ml/hr)
 - ii. Maintenance Dose = Dose2: in 500 ml 5% Dextrose (125ml/hr)
 - iii. Maintenance Dose = Dose 3: in 1000 ml 5% Dextrose (62.5ml/hr)
 - e. Compatible in D5W

Hepatic Toxicity

- 1. Consult Gastroenterologist
- 2. Decrease protein intake
- 3. Bowel decontamination with neomycin
- 4. Antacids to prevent bleeding

Prognosis

- 1. Mortality rate is <0.5%
- 2. There is no long term sequelae after acute toxicity