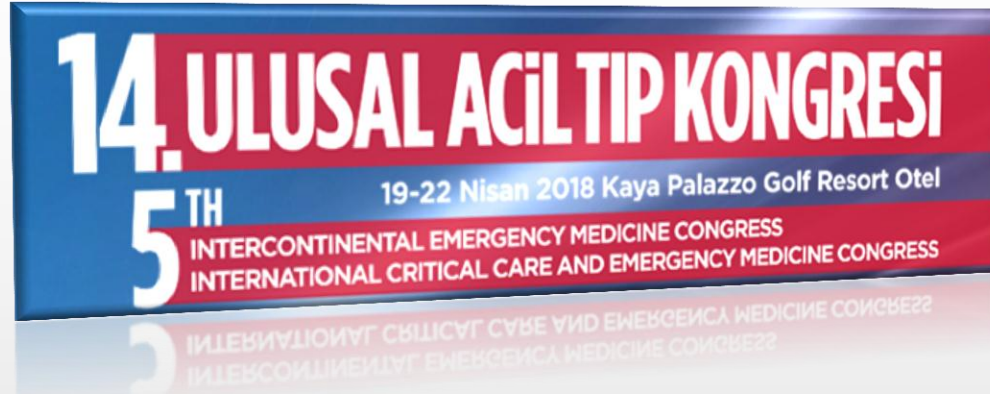




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**Which antidotes are in our
pocket during daily practices?**

Afşin Emre Kayıpmaz, MD



Initial stabilization and therapy

- Endotracheal intubation as needed for airway protection, oxygenation, ventilation, and orogastric lavage
- Supplemental oxygen for hypoxia
- Pulse oximetry
- Cardiac monitor
- IV access



- **Hypotension:**
 - Administer 0.9 % normal saline IV fluid bolus
 - Trendelenburg position
 - Vasopressors for persistent hypotension
- **Bradycardia:**
 - Atropine
 - Cardiac pacing
- **If altered mental status:**
 - Thiamine,
 - D50W
 - Naloxone



- **Decontamination:**

- Gastric Decontamination.
- Prevents systemic absorption of ingested toxin

- **Orogastric lavage:**

- Consider in potentially lethal ingestions without known antidote within 1 hour of ingestion
- Protected airway essential prior to lavage



- **Activated charcoal:**
 - Most effective within a few hours of most toxic ingestions
 - Contraindicated if caustic ingestion, unprotected airway, or bowel obstruction



- Drugs not effectively bound to charcoal:
 - Metals (borates, bromide, iron, lithium),
 - Alcohols,
 - Potassium



- **Activated charcoal:**
 - Multiple-dose activated charcoal:
 - Theophylline
 - Carbamazepine
 - Phenobarbital
 - Dosing: 1–2 g/kg PO



- **Whole-bowel irrigation:**

- Polyethylene glycol evacuates bowel without causing electrolyte disturbances
- In toxins not well adsorbed by charcoal (e.g., iron and lithium), body packers/stuffers, sustained-release ingestions.
- Contraindicated if bowel obstruction, perforation, or hypotension



- **Urinary alkalization:**
 - Salicylates,
 - Phenobarbital
- **Hemodialysis/hemoperfusion:**
 - Salicylates,
 - Lithium
 - Theophylline
 - Toxic alcohols
 - Valproate



Acetaminophen: N-acetylcysteine

- **Oral dosing** is suitable for non-pregnant patients with a functional GI tract (no hepatotoxicity)
- **140 mg/kg** loading dose,
 - **followed by 17 doses of 70 mg/kg every 4 hours**
- If vomiting occurs within 1 hour of NAC dosing, a full dose should be repeated



N-acetylcysteine: IV dosing

- In patients with no biochemical evidence of hepatic failure (ie, INR <2):
- 21 hour IV protocol:
 - **150 mg/kg** loading dose over 60 minutes,
 - followed by **50 mg/kg** infused over 4 hours,
 - with the final **100 mg/kg** infused over the remaining 16 hours



N-acetylcysteine: IV dosing

- In patients with biochemical evidence of hepatic failure (ie, INR >2),
- 21 hour IV protocol:
 - **150 mg/kg** loading dose over 60 minutes,
 - followed by **50 mg/kg** infused over 4 hours,
 - followed by **100 mg/kg** infused over the next 16 hours)
 - followed by a continuous IV NAC **infusion at 6.25 mg/kg** per hour until INR is <2



Anticholinergic: Physostigmine

- Patients who manifest both peripheral & moderate central anticholinergic toxicity should be treated with:
 - **0.5 to 2 mg** (0.02 mg/kg IV, up to a maximum of 0.5 mg per dose in pediatric patients);
(should be given by slow IV push, over five minutes)



Benzodiazepines: Flumazenil

- Nonspecific competitive antagonist of the BZD receptor
- It can be used to reverse BZD-induced sedation following general anesthesia, procedural sedation, or overdose



Benzodiazepines: Flumazenil

- The use of flumazenil in the setting of overdose remains highly controversial
- It can precipitate withdrawal seizures in patients who have developed a tolerance to BZDs through chronic use or abuse



Benzodiazepines: Flumazenil

- It does not consistently reverse respiratory depression caused by BZD overdose
- It appears to be safe and effective when used to reverse the sedating effects of a BZD in patients who do not use BZDs chronically.



Benzodiazepines: Flumazenil

- Initial dose is **0.2 mg** IV over 30 seconds
- Repeated doses of **0.2 mg**, to a **maximum dose of 1 mg** can be given until the desired effect is achieved (no more than 3 mg within any one hour)



β -blockers: Glucagon et al.

- For severe poisoning (eg, profound hypotension), give following treatments simultaneously:
 - IV glucagon, **5 mg** IV bolus, may be repeated if the initial bolus is ineffective
 - IV calcium salts,
 - Calcium chloride (10% solution) **10 to 20** mL
 - Calcium gluconate (10% solution) **30 to 60** mL
 - Vasopressor (eg, epinefrin),
 - IV high-dose insulin (with glucose),
 - IV lipid emulsion therapy.



- IV high-dose insulin (with glucose): Bolus of **1 U/kg** IV of regular insulin, followed by infusion of **0.5 U/kg/hour**; (titrate infusion upwards until hypotension corrected or dose reaches 2 U/kg/hour)
- Relative hypoglycemia and hypokalemia must be corrected prior to therapy
- IV lipid emulsion therapy: **1.5 mL/kg** over 2 minutes, followed by **1.5 mL/kg** infusion over 60 minutes



Carbon monoxide: Oxygen, hyperbaric oxygen

- Intubate as clinically indicated
- Apply high-flow oxygen to **all** poisoned patients regardless of pulse oximetry or arterial pO₂



Carbon monoxide: Oxygen, hyperbaric oxygen

- Hyperbaric oxygen (HBO) for:
 - CO level >25 percent (>20 percent if pregnant)
 - Loss of consciousness
 - Severe metabolic acidosis (pH <7.1)
 - Concern for end-organ ischemia (chest pain, ECG changes, altered mental status)



Vitamin K antagonist (eg, warfarin)

- If 4-factor prothrombin complex concentrate is available (preferred approach):
 - 4F PCC **1500 to 2000 units** IV over 10 minutes.
 - Check INR 15 minutes after completion of the infusion
 - If INR is not ≤ 1.5 , give additional 4F PCC
 - Give vitamin K 10 mg IV over 10 to 20 minutes.



Vitamin K antagonist (eg, warfarin)

- If 3-factor prothrombin complex concentrate is available
 - 3F PCC **1500 to 2000 units** IV over 10 minutes.
 - Check INR 15 minutes after completion of the infusion
 - If INR is not ≤ 1.5 , give additional 3F PCC
 - Give vitamin K 10 mg IV over 10 to 20 minutes.



Vitamin K antagonist (eg, warfarin)

- **Neither 3F PCC nor 4F PCC is available**
 - FFP 2 units IV by rapid infusion.
 - Check INR 15 minutes after completion of infusion. If $INR \geq 1.5$, administer 2 additional units
 - Repeat process until $INR \leq 1.5$
 - Administer loop diuretic if volume overload occurs
 - Give vitamin K 10 mg IV over 10 to 20 minutes



Vitamin K antagonist (eg, warfarin)

- These products and doses are for use in life-threatening bleeding only
- PCC will reverse anticoagulation within minutes of administration;
- FFP administration can take hours due to the volume required;
- Vitamin K effect takes 12 to 24 hours, but administration of it is needed to counteract the long half-life of warfarin.



Cyanide: Cyanide antidote kit, hydroxocobalamin

- If hydroxocobalamin is available
 - Hydroxocobalamin **70 mg/kg** up to 5 g IV (**5 g** is standard adult dose)
 - Sodium thiosulfate (25%): **1.65 mL/kg** up to 50 mL IV; may repeat once (maximum dose 12.5 g)



Cyanide: Cyanide antidote kit, hydroxocobalamin

- If hydroxocobalamin is not available (no contraindications to nitrites)
 - Sodium nitrite **10 mg/kg - up to 300 mg** - by slow IV infusion; may repeat once
 - Sodium thiosulfate (25%): **1.65 mL/kg** up to 50 mL IV; may repeat once



Cyanide: Cyanide antidote kit, hydroxocobalamin

- If hydroxocobalamin is not available and cyanide toxicity is possible but not certain, or the patient has contraindications to nitrites
- Sodium thiosulfate (25%): **1.65 mL/kg** up to 50 mL IV; may repeat once



Methanol, Ethylene glycol: 4-methylpyrazole (Fomepizol), Ethanol

- Block alcohol dehydrogenase with **fomepizole**,
 - **15 mg/kg** IV loading dose,
 - followed by **10 mg/kg** q 12 h x 4 doses.
 - If patient requires further treatment after this regimen, increase dose to **15 mg/kg** every 12 hours



Methanol, Ethylene glycol: 4-methylpyrazole (Fomepizol), Ethanol

- If fomepizole is unavailable or patient has a known allergy, block alcohol dehydrogenase with ethanol,
 - **10 mL/kg** of a ethanol solution (10%),
 - followed by **1 mL/kg** of ethanol solution (10%) infused per hour.
 - Titrate to serum ethanol concentration of **100 mg/dL**.



Methanol, Ethylene glycol: 4-methylpyrazole (Fomepizol), Ethanol

- Administer **sodium bicarbonate**,
 - 1 to 2 meq/kg bolus followed by infusion of 132 meq NaHCO_3 in 1 L D5W to run at 200 to 250 mL/hour for patients with pH below 7.3
- For patients with known or suspected methanol poisoning, administer **folic acid**, 50 mg IV every six hours



Methanol, Ethylene glycol: 4-methylpyrazole (Fomepizol), Ethanol

- For patients with known or suspected ethylene glycol poisoning, administer
 - **thiamine, 100 mg IV, and pyridoxine, 50 mg IV**



Methemoglobinemia: Methylene blue

- If the patient is symptomatic and does not have G6PD deficiency, we recommend the immediate use of intravenous MB
- Prefer MB over ascorbic acid because of its more rapid onset of action.
 - The usual dose in this setting is **1 to 2 mg/kg**, given over five minutes.



Opiates: Naloxone

- If the O₂ saturation is <90% but the patient is breathing spontaneously, administer supplemental oxygen followed by IV naloxone, **0.05 mg**
- Repeat until ventilation is adequate.
- The goal of treatment is adequate ventilation, **not** normal mental status



Opiates: Naloxone

- If the response is inadequate after 5 to 10 mg, reconsider the diagnosis
- If the patient is apneic, ventilate using a bag-valve mask attached to supplemental oxygen and administer naloxone in doses of **0.2 to 1 mg IV or IM**



Organophosphates: Atropine, pralidoxime

- **Atropine 2 to 5 mg IV/IM/IO bolus**
 - Escalate (double) dose every 3-5 minutes until bronchial secretions and wheezing stop
 - Tachycardia and mydriasis are not contraindications
 - Hundreds of milligrams may be needed over several days in severe poisonings
 - Inhaled ipratropium 0.5 mg with parenteral atropine may be helpful for bronchospasm



Organophosphates: Atropine, pralidoxime

- **Pralidoxime (2-PAM)**
 - **2 g** IV over 30 minutes; may repeat after 30 minutes or give continuous infusion if severe
 - Continuous infusion at 8 mg/kg/hour in adults
 - If no IV access, give pralidoxime 600 mg IM
 - is given with atropine
 - Diazepam **10 mg** IV, repeat as necessary if seizures occur. Do not give phenytoin.



Tricyclic antidepressants: NaHCO_3

- for QRS duration >100 msec or any ventricular arrhythmia caused by TCA poisoning
 - The initial dose of sodium bicarbonate is **1 to 2 mEq/kg**.
 - In adults, this may be given as two to three **50 mEq (50 mL)** vials or prefilled syringes of 8.4 percent sodium bicarbonate given as a rapid IV push through a large bore IV



Digoxin: Digibind

- Clinically significant manifestations of digitalis poisoning be treated with digoxin-specific antibody (Fab) fragments
- As temporizing measures or if Fab fragments are not immediately available, bradycardia can be treated with atropin 0.5 mg IV in adults and hypotension with IV boluses of isotonic crystalloid



Iron: Deferoxamine

- The usual dose of deferoxamine is **15 mg/kg** per hour intravenously.
- In cases of severe overdose, higher doses of deferoxamine (up to 35 mg/kg per hour) be administered during the first 24 hours of treatment



Iron: Deferoxamine

- Patients with a significant number of radiopaque pills on abdominal radiograph receive gastric lavage and/or whole bowel irrigation



Isoniazid: Pyridoxine (vitamin B6)

- **1 g** intravenously for every gram of INH ingested.
- When the quantity of INH ingested is unknown, **5 g IV** may be administered to an adult
- The dose can be administered at a rate of **0.5 g/min**, and may be repeated in patients with refractory seizure activity

4-metil pirazol (Fomepizol sulfat)	FOMEPIZOL	5 amp / kutu	100 mg / 20 ml	Etilen glükol ve Methanol zehirlenmeleri
Botulismus Polivalan Antiserum (A-B-E)	BOTULİSMUS ANTİTOKSİN	250 ml şişe	Tip A 750 IU/ml Tip B 500 IU/ml Tip E 50 IU/ml	Botulismus vakaları için Antitoksin
Calcium Edeate sodyum	CALCIUM EDEDATE DE SODIUM % 5	10 amp / kutu	500 mg / 10 ml	Kurşun zehirlenmeleri
Di cobalt EDTA	KELOCYANOR % 1,5	6 amp / kutu	300 mg / 20 ml	Siyanür zehirlenmeleri
Digoksin İmmün Fab	DİGİFAB	1 vial / kutu	40 mg / vial	Digoksin zehirlenmeleri
Dimercaprol	B.A.L.	12 amp / kutu	200 mg / 2 ml	Ağır metal şelatörü
DMP5	DİMAVAL	20 cap / kutu	100 mg kapsül	Ağır metal şelatörü (Hg)
D-penisilamin	METACAPTASE	100 cap / kutu	150 mg kapsül	Ağır metal şelatörü (Pb, Cu)
Etil Alkol	ETİL ALKOL % 10	500 ml şişe	500 ml şişe	Etilen glükol ve Methanol zehirlenmeleri
Hydroxocobalamin	CYANO KİT 2,5 g	2 vial / kutu	2,5 gr vial	Siyanür zehirlenmeleri
Metilen Mavisi	METİLEN MAVİSİ % 1	1 flakon	20 ml / flakon	Methemoglobinemi yapan zehirlenmeler
Physostigmine	ANTİKOLİUM	5 amp / kutu	2 mg / 5 ml	Antikolinergik zehirlenmeler
Pralidoksim	CONTRATHİON	10 flakon/ kutu	200 mg / flakon	Organik fosfor zehirlenmeleri
Silibinin	LEGALON-SİL	4 flakon / kutu	350 mg / flakon	Mantar zehirlenmeleri
Succimer (DMSA)	SUCCİCAPTAL	15 cap / kutu	200 mg / kapsül	Ağır metal şelatörü (Hg)



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