

# Poisonings and Intoxications: What a Nephrologist Needs to Know

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- Research focus: Applications of digital health technologies in alcohol use disorder/substance use disorder, medication safety



# DISCLOSURES

None



# OBJECTIVES

1. Recognize key nephrotoxins
2. Understand specific indications for dialysis in toxicity
3. Review the role of the nephrologist in key high-risk toxicities and renal-specific considerations



# Case 1

35-year-old man comes to the Emergency Department, appears intoxicated and minimally responsive.

EMS found the patient with a suicide note and a bottle of antifreeze.

VS: Afebrile, tachycardic, tachypnea, BP 120/60



# Case 1 - Labs

11.8 > 14.8/44.8 < 188

140	112	17	< 98
5.5	<5	0.95	

Anion gap **23**

Alk Phos 74

AST 31

ALT 21

CK 289

APAP <10

Salicylate <10

Ethanol <10

pH **6.9**/18/68/3.7

Lactate **8.9**

Measured serum osm –

Measured Osmolal gap – 355

Calculated – 285

**Osm gap – 70**

Urine – calcium oxalate crystals

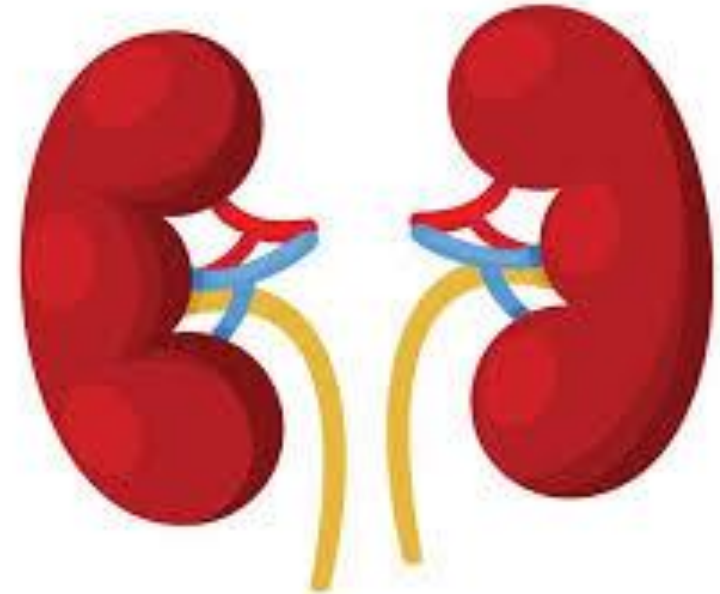


# NEPHROLOGY IS CONSULTED....



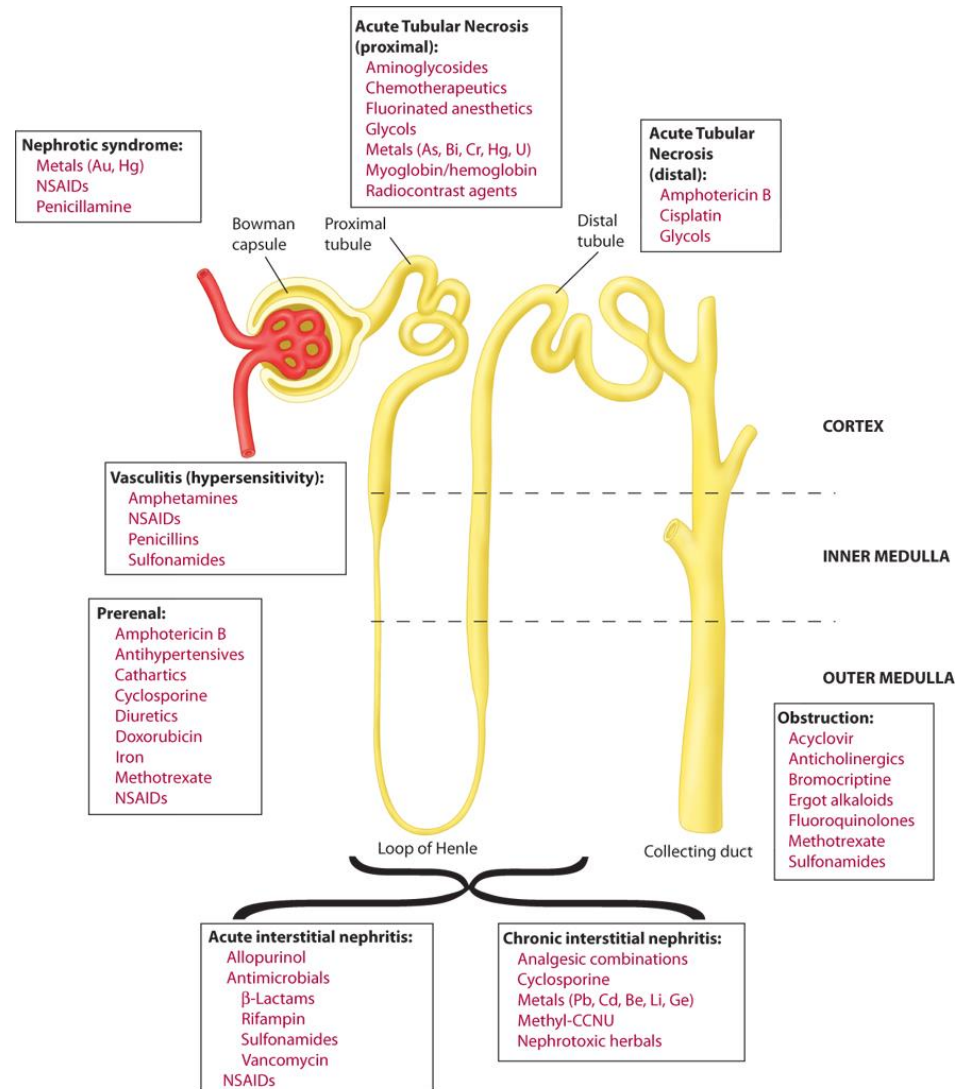
# Why This Matters for Nephrologists

- Toxins frequently cause AKI or require dialysis
- Nephrologists often consulted for RRT, fluid/electrolytes, or drug dosing
- Overlap with ED, ICU, and toxicology





# The Kidney in Toxicology - Nephrotoxins



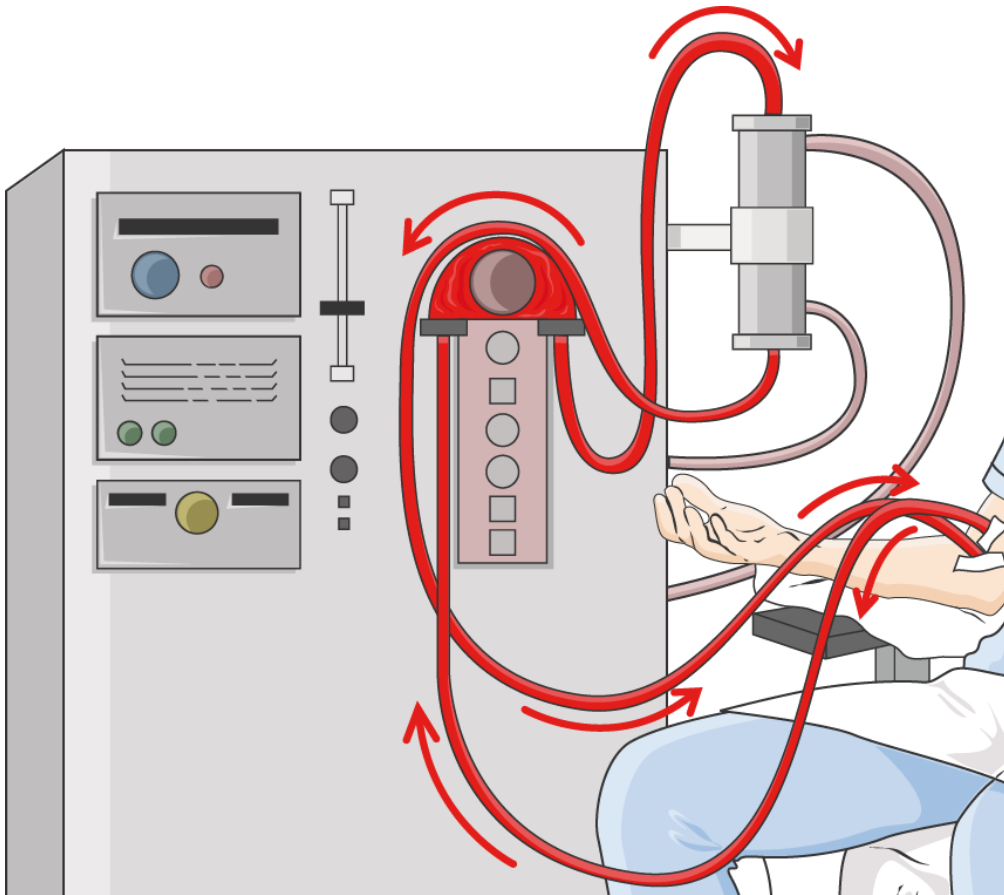
Source: L.S. Nelson, M.A. Howland, N.A. Lewin, S.W. Smith, L.R. Goldfrank, R.S. Hoffman: Goldfrank's Toxicologic Emergencies, Eleventh Edition  
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# Indications for extracorporeal removal



Methods to enhance removal:

Hemodialysis (HD)

Charcoal hemoperfusion use declining

Forced diuresis

Urinary alkalinization

Most common use of HD lithium, ethylene glycol, salicylates, acetaminophen

HD indicated for patients in cases where the responsible xenobiotics is removable AND:

- Patients fail to respond to comprehensive supportive care
- Normal route of elimination is impaired high morbidity or mortality is likely
- Correction of severe electrolyte disorders

# Characteristics of a xenobiotics that can be removed by HD

Volume of distribution below 1 L/kg

Low protein binding (or large amount of free xenobiotics in overdose)

HD enhances clearance compared to endogenous clearance by a factor of 2 or overall low endogenous clearance

Low lipophilicity

\*\*Largest factor depends on pharmacokinetics in overdose

Improved clearance prior to distribution

TABLE 6-3

Characteristics of Xenobiotics That Allow Clearance by Hemodialysis, Hemoperfusion, and Hemofiltration

For All Three Techniques	For Hemodialysis	For Hemoperfusion	For Hemofiltration
Low $V_d$ (<1 L/kg)	MW <5,000 Da	MW <50,000 Da	MW <40,000 Da
Single-compartment first-order kinetics	Low protein binding	Adsorption by activated charcoal	Low protein binding
Low endogenous clearance (<4 mL/min/kg)			

MW = molecular weight;  $V_d$  = volume of distribution.



# Intermittent Hemodialysis vs Continuous Renal Replacement Therapy (CRRT)

Intermittent HD preferred in most situations

Usually performed for 4 to 8 hours

CRRT can be used but results in slower removal rates



# EXTRIP Workgroup

Extracorporeal Treatment in Poisoning (EXTRIP) is a collaboration of experts from diverse specialties

- Clinical toxicology
- nephrology
- pharmacology
- critical care
- emergency medicine

Publish recommendations regarding the indications for dialysis for overdose

<https://www.extrip-workgroup.org/publications>



# Nephrologist role in poisonings

- Electrolyte and fluid management
- Renal dosing of antidotes
- Monitoring AKI recovery
- Preventing further renal injury
- Multi-disciplinary communication



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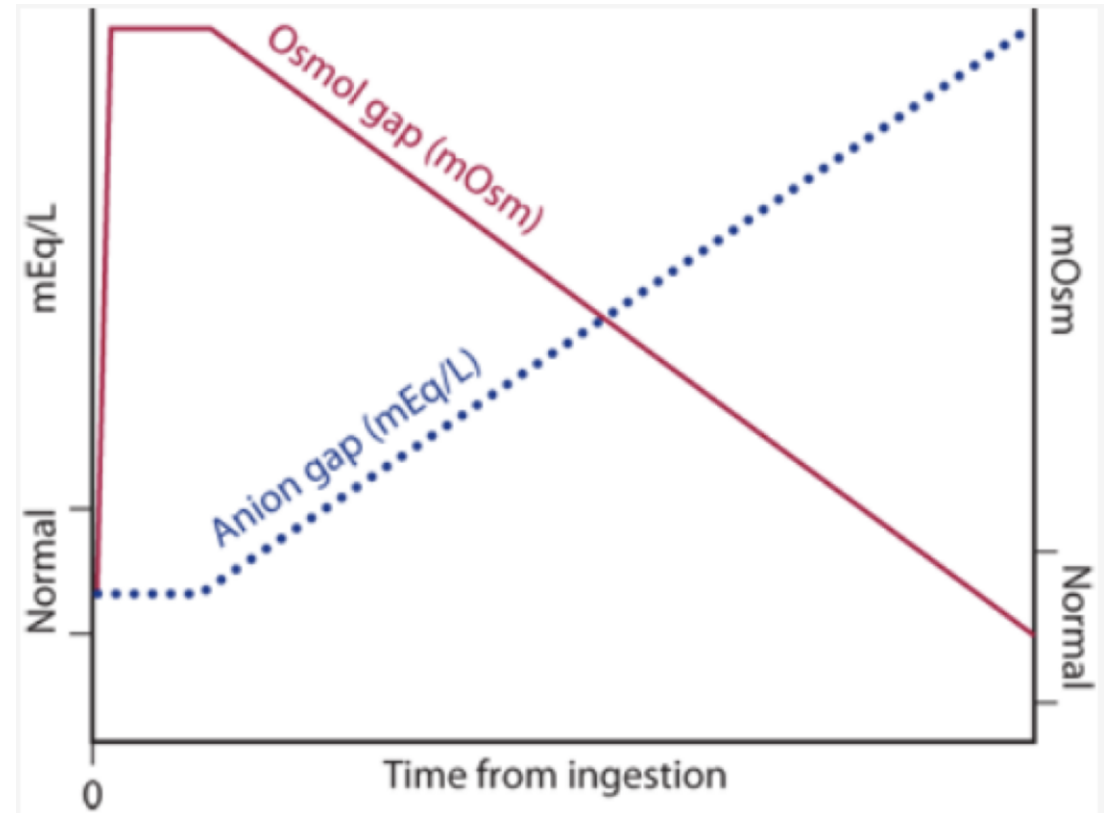
**Ethylene Glycol: 277.3 mg/dL**



# Specific Toxins – Ethylene glycol

## Ethylene glycol

- Symptoms – intoxication, nephrotoxicity, hypocalcemia, metabolic acidosis, cerebral edema
- Labs – elevated serum osm gap, large anion gap acidosis, elevated lactate,

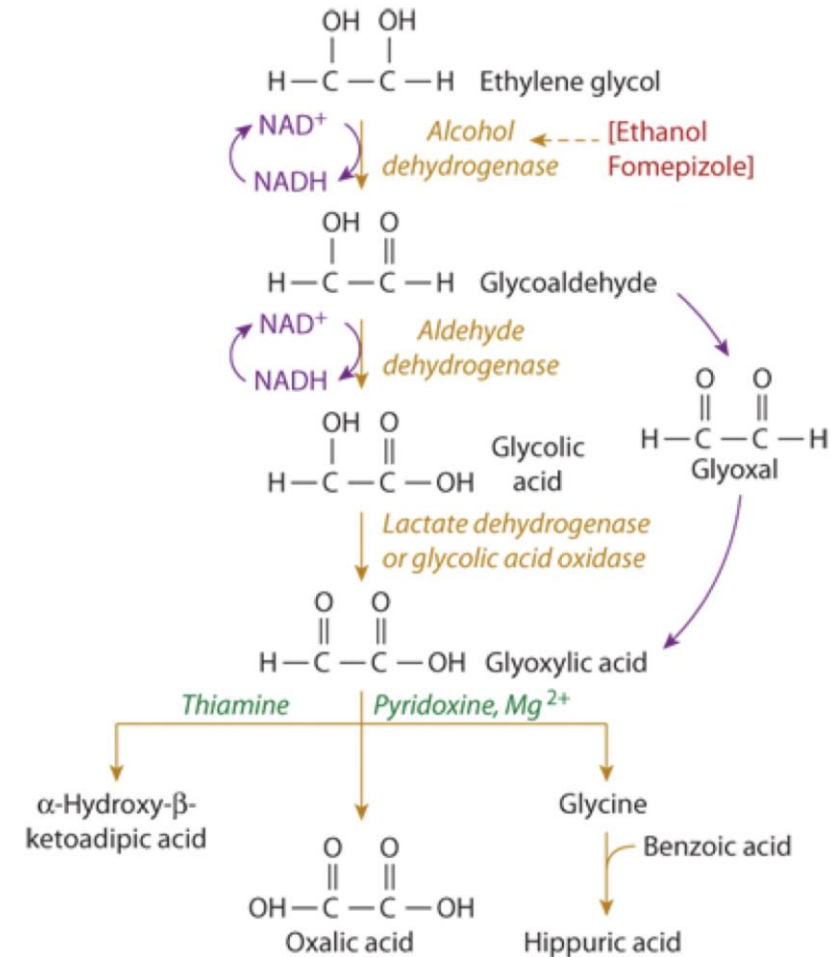




# Specific Toxins – Toxic Alcohols

## Ethylene Glycol and Methanol

- Converted to toxic metabolites by alcohol dehydrogenase
- Renal crystal injury (EG), visual symptoms (methanol)
- Treatment: Fomepizole/ethanol, dialysis



# Ethylene glycol – Indications for HD



Suspected EG ingestion with osmol gap greater than 50 mOsm/L and acidosis (pH <7.25-7.30)

Renal Failure

End organ toxicity

EG concentration greater than 50 mg/dL

Severe metabolic acidosis

## Case 2

38-year-old man with past medical history of bipolar disorder presented for increasing tremors

First noticed in his bilateral hands, which seemed to be worsening over the past two days as well as generalized “muscle spasms”

VS: BP 112/76 HR 124 RR 20 O2 96% T 98.2

Labs: Na<sup>+</sup> 129

Cr 5.9

You get one more lab....

Lithium level 4.2 mmol/L



# Specific Toxins - Lithium

Acute:

Predominately GI symptoms, nausea, vomiting

Chronic:

Predominately neuro symptoms

Acute-on-chronic:

GI and neuro effects

Other effects: chronic tubulointerstitial nephropathy, hypothyroidism, hyperparathyroidism, hypercalcemia, Ebstein anomaly, leukocytosis

SILENT - Syndrome of irreversible lithium-effectuated neurotoxicity

Nephrogenic Diabetes Insipidus



# Specific Toxins 2 – Lithium

Treatment – IV fluids and dialysis

Dialysis indications:

- Concentration > 5.0 mEq/L
- Concentration > 4.0 mEq/L with AKI or CKD
- Decreased level of consciousness, seizures, or life-threatening dysrhythmias at any lithium concentration
- Estimated time to reach lithium concentration < 1 mEq/L exceeds 36 hours



# Specific Toxins – Salicylates

Mixed acid-base disturbance

Clinical signs: confusion, tinnitus, hyperventilation

Treatment: fluid resuscitation, urine alkalinization

Dialysis Indications:

- Concentration  $> 7.2$  mmol/L (90 mg/dL)
- Concentration  $> 6.5$  mmol/L (80 mg/dL) with AKI or CKD
- Concentration  $> 6.5$  mmol/L (90 mg/dL) after IV Altered mental status
- Respiratory distress or new hypoxemia requiring supplemental oxygen
- $\text{pH} \leq 7.2$



# Specific Toxins – Acetaminophen

Frequent cause of overdoses and toxicity  
Causes hepatotoxicity

Reasons for nephrology involvement:

- Severe metabolic acidosis
- Acute kidney injury
- Hemodialysis considerations

Treatment – n-acetylcysteine

Dialysis indications:

- Acetaminophen concentration  $>1000$  mg/L and NAC is NOT administered
- Presence of altered mental status, metabolic acidosis, with an elevated lactate, and an [APAP]  $> 700$  mg/L and NAC is NOT administered
- If the patient presents with an altered mental status, metabolic acidosis, an elevated lactate, and an [APAP]  $> 900$  mg/L even if NAC is administered



# Specific Toxins – Metformin-associated lactic acidosis

Rare presentation with high mortality

Risk factors – renal injury, hepatic failure, critical illness

Presentation - Profound acidosis, shock

Treatment:

- Dialysis clears metformin and acid
- CRRT useful in unstable patients





## Case 3

28-year-old male found unresponsive at a party. Per EMS bystanders reported cocaine and alcohol use. Arrives to ED confused, agitated, diaphoretic

VS: T: 38.6°C; BP: 92/58 mmHg; HR: 136 bpm; RR: 26/min; SpO<sub>2</sub>: 96% on room air

Labs:

Creatinine: 3.2 mg/dL (baseline 0.9)

CK: 112,000 U/L

K<sup>+</sup> 6.5



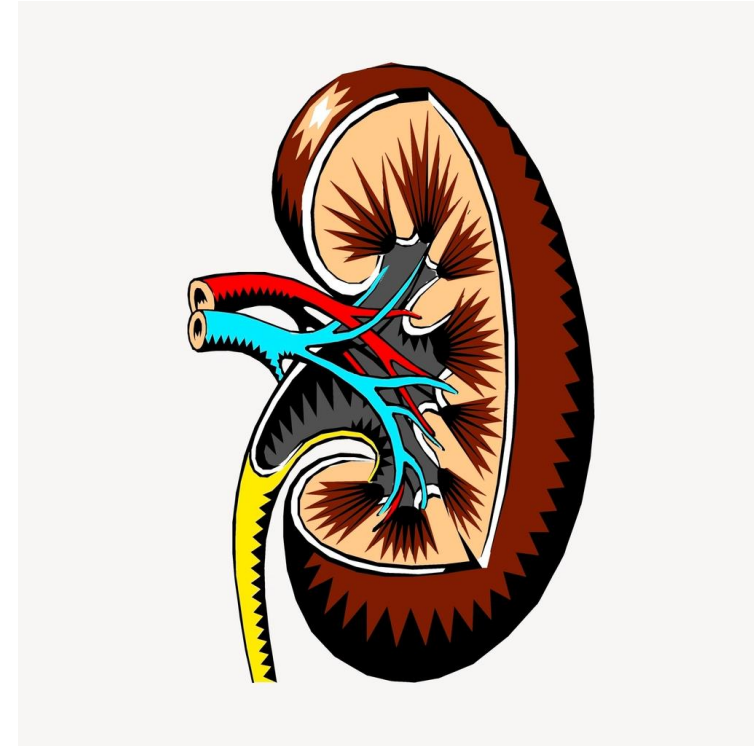
# Rhabdomyolysis-Induced AKI

Causes: drugs (cocaine, statins),  
trauma, seizures

Fluid resuscitation key

Manage electrolyte abnormalities

Dialysis if severe AKI or electrolyte  
derangements



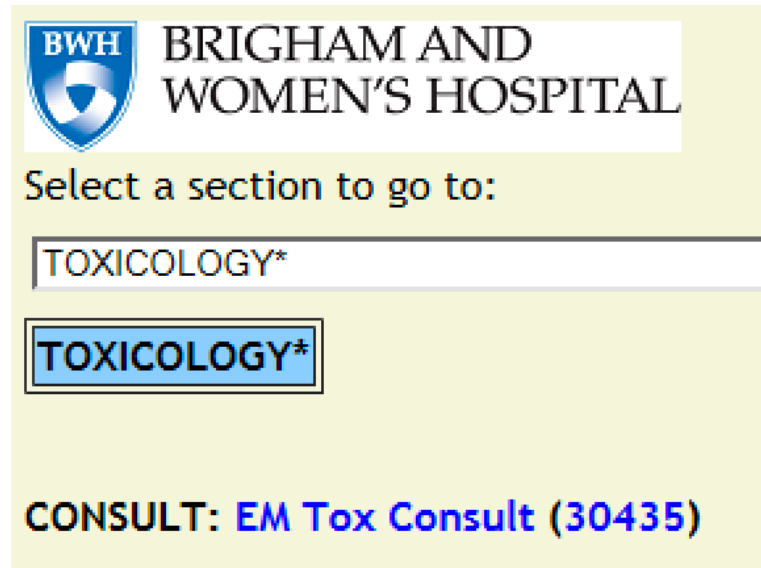
# TAKE HOME MESSAGES

- 1. Know the dialyzable toxins.** Ethylene glycol, methanol, lithium, salicylates, metformin, and in selected cases, acetaminophen may require extracorporeal removal.
- 2. Nephrologists are key in the acute phase.** Your expertise in fluid and electrolyte optimization, renal dosing of antidotes, and AKI recovery monitoring is critical to patient outcomes.
- 3. Leverage expert guidance.** EXTRIP's consensus recommendations can be useful to make timely, evidence-based decisions on initiating dialysis for toxin removal.



# Resources for Clinicians

EXTRIP.org  
Poison Control  
BWH Toxicology



The screenshot shows the BWH website's navigation menu. At the top left is the BWH logo, a blue shield with a white 'B' and 'W' and a blue 'H'. To its right, the text 'BRIGHAM AND WOMEN'S HOSPITAL' is displayed in a serif font. Below this, the text 'Select a section to go to:' is followed by a search bar containing the text 'TOXICOLOGY\*'. Below the search bar is a button with the text 'TOXICOLOGY\*' in a blue box. At the bottom of the menu, the text 'CONSULT: EM Tox Consult (30435)' is displayed in a blue font.

**POISON**  
**Help**  
**1-800-222-1222**



The logo for Poison Help features the word 'POISON' in a bold, black, sans-serif font. Below it, the word 'Help' is written in a large, bold, black, sans-serif font. A red pill bottle with a white cap and a white skull and crossbones symbol is positioned between the 'e' and 'p' of 'Help'. Below 'Help', the phone number '1-800-222-1222' is displayed in a bold, black, sans-serif font.



# REFERENCES

1. The Role of the Nephrologist in Management of Poisoning and Intoxication: Core Curriculum 2022. Mullins, Michael E. et al. American Journal of Kidney Diseases, Volume 79, Issue 6, 877 – 889
2. Nelson LS, Howland MA, Lewin NA, et al. eds. Goldfrank's Toxicologic Emergencies. 11th ed. McGraw-Hill; 2018.
3. Ghannoum M, Nolin TD, Lavergne V, Hoffman RS; EXTRIP Workgroup. Blood purification in toxicology: nephrology's ugly duckling. Adv Chronic Kidney Dis. 2011;18(3):160-166.



# Thank you!

