



ASSISTANT CLINICAL PROFESSOR, EMERGENCY MEDICINE

UCSF

CALIFORNIA POISON CONTROL CENTER - SF DIVISION







Disclosures

None



HR 134 BP 161/112 T 37.4 RR 21



Diaphoretic Tremulous Restless Hallucinosis

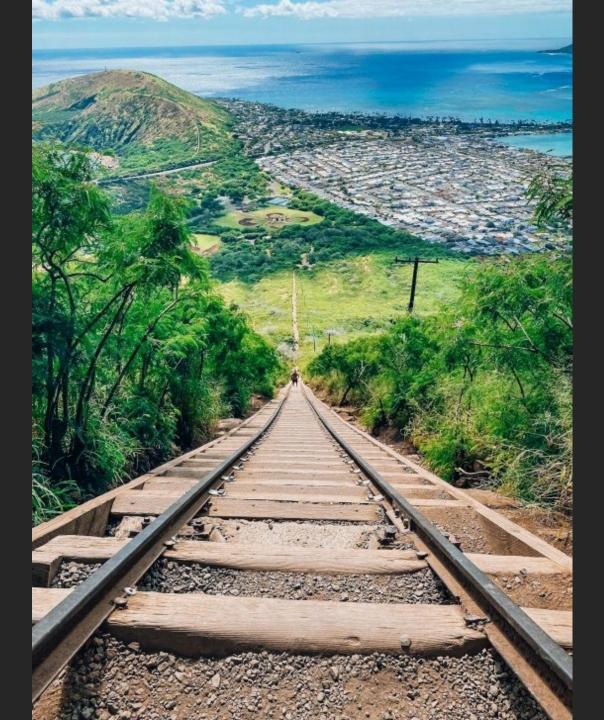


EtOH 51 Lactate 4.4 Bicarb 15 Anion Gap 22











Naïve-sober





Baseline normal



= GABA receptor with $\alpha_{\rm 1}$ subunit



= GABA receptor with α_4 subunit



= NMDA receptor

= Ethanol



Alcohol Withdrawal Syndrome

• Clinical diagnosis

CIWA?

Tactile disturbances

Ask, 'Have you any itching, pins and needles sensations, any burning, any numbness, or do you feel bugs crawling on or under your skin?'

| None | |
|---|----|
| Very mild itching, pin and needles, burning, or numbness | +1 |
| Mild itching, pin and needles, burning, or numbness | +2 |
| Moderate itching, pin and needles, purning, or numbness | +3 |

Headache/fullness in head

Ask 'Does your head feel different? Does it feel like there is a band around your head?' Do not rate for dizziness or lightheadedness.
Otherwise, rate 'severity.'

| Not Present | |
|-------------------|----|
| Very mild | +1 |
| Mild | +2 |
| Moderate | +3 |
| Moderately severe | +4 |
| Severe | +5 |
| Very severe | +6 |
| Extremely severe | +7 |

Orientation/clouding of sensorium

Ask 'What day is this? Where are you? Who am

| Oriented, can do serial additions | |
|--|----|
| Can't do serial additions or is uncertain about date | +1 |
| Disoriented for date by no more than 2 calendar days | +2 |
| Disoriented for date by more than 2 calendar days | +3 |
| Disoriented to place or person | +4 |

you seeing anything that is disturbing to you? Are you seeing things you know are not there?'

| very minu sensitivity | TI |
|----------------------------------|----|
| Mild sensitivity | +2 |
| Moderate sensitivity | +3 |
| Moderately severe hallucinations | +4 |
| Severe hallucinations | +5 |
| Extremely severe hallucinations | +6 |
| Continuous hallucinations | +7 |



The ASAM
Clinical Practice
Guideline on
Alcohol Withdrawal
Management

→ Recommendation I.6: Alcohol withdrawal severity assessment scales (including the Clinical Instrument Withdrawal Assessment for Alcohol, Revised [CIWA-Ar]) should not be used as a diagnostic tool because scores can be influenced by conditions other than alcohol withdrawal.

Alcohol Withdrawal Syndrome



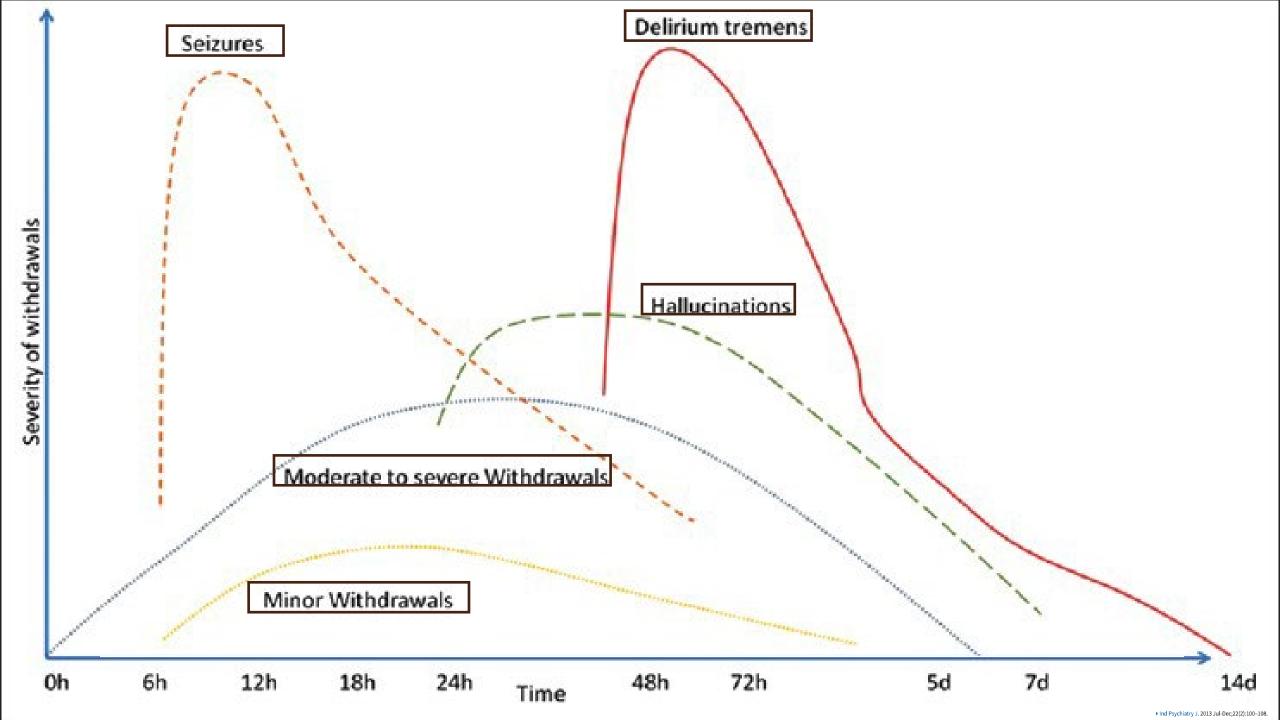
Cessation or Decrease after chronic or heavy EtOH use



Autonomic hyperactivity
Tremor
Nausea/vomiting
Anxiety or agitation
Hallucinosis
Seizures



Not attributable to other causes



Severe Alcohol Withdrawal Syndrome



History of prior severe AWS



Degree of autonomic hyperactivity



Detectable ethanol



What precipitated the AWS?

- Intercurrent illness?
- Loss of access to EtOH?
- Presentation for unrelated concern



History & Exam: Peals & Pitfalls



- AWS follows cessation or decrease in EtOH use
- Severe AWS can be predicted by history of similar or elevated EtOH at presentation



- Don't miss intercurrent illness accompanying AWS
- Not all patients follow the predictable course
- Don't rely on a scoring system to make the diagnosis

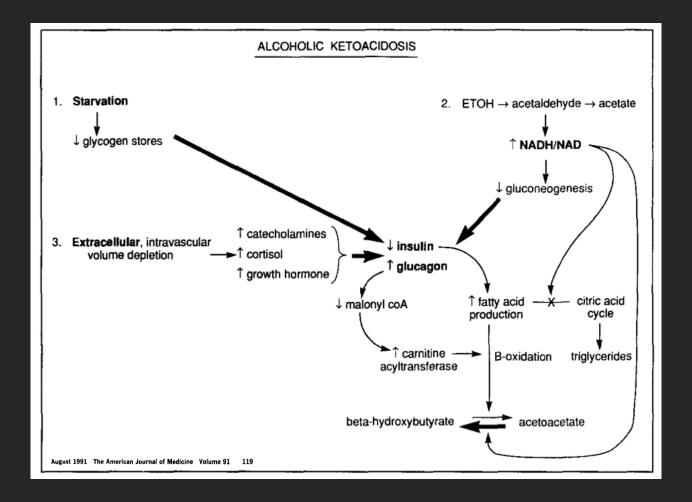
Evaluation



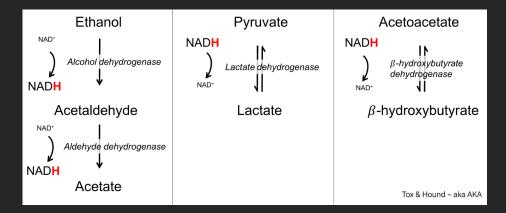
Evaluation

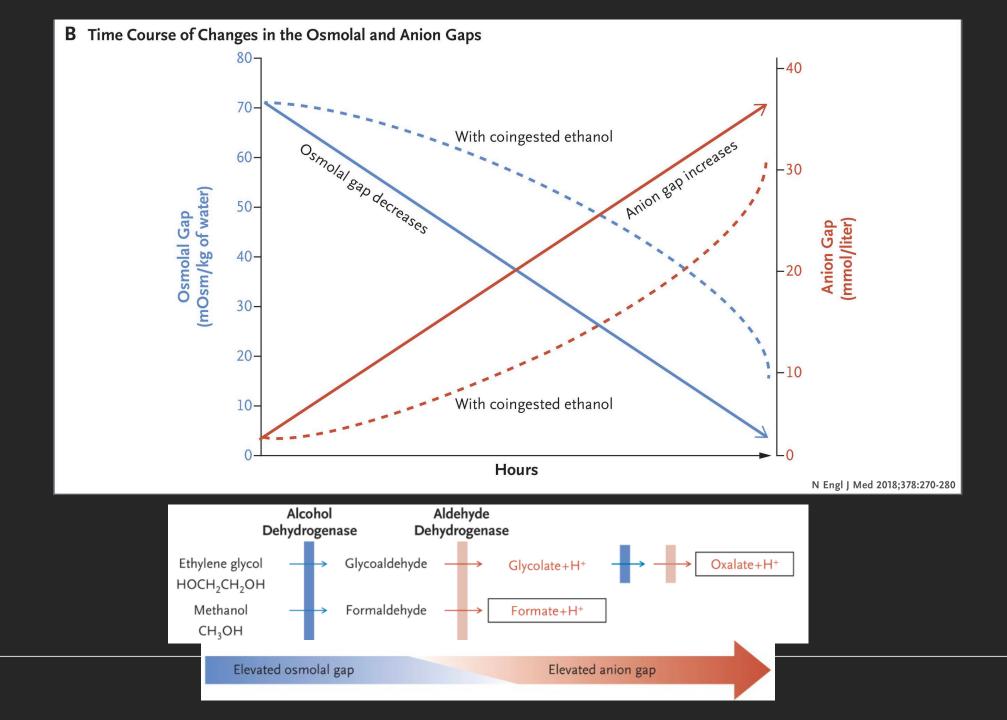
- CBC, CMP, Mg, INR, EtOH
- ECG
- Lipase
- Beta-hydroxybutyrate, lactate
- UA, UDS
- CT H, CXR





B-HB? Lactate!?





Treatment of Metabolic Derangements



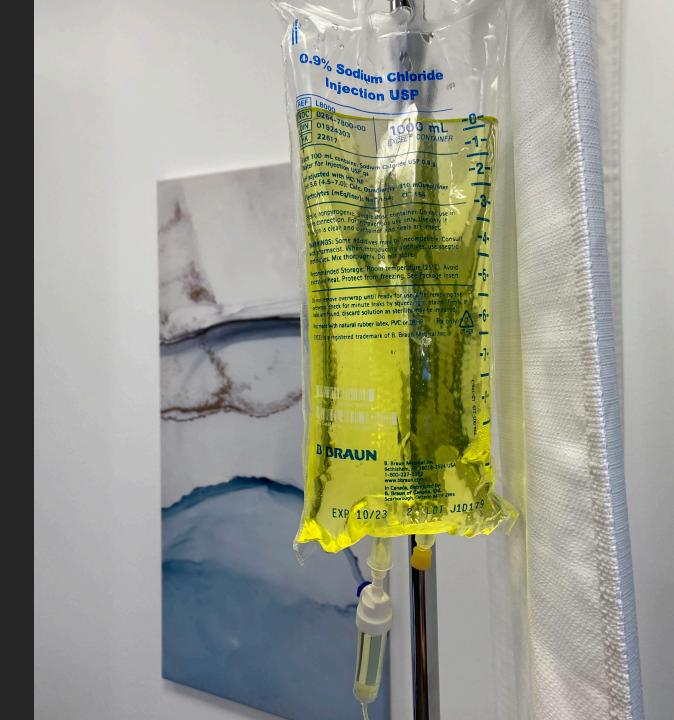
IVF



PO or IV calories (e.g., 1L D5W + 200 ml/hr)



Thiamine 100 mg IV



Treatment of Metabolic Derangements

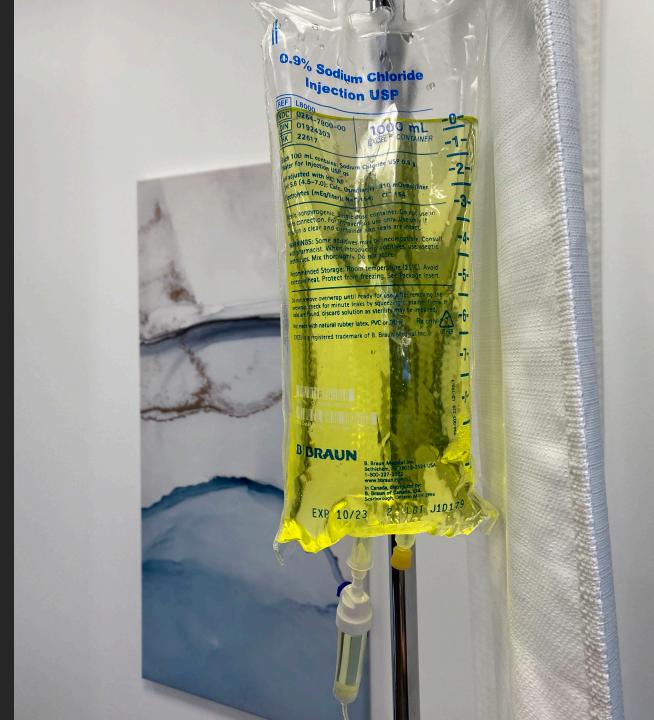


Unpeeling the Evidence for the Banana Bag: Evidence-Based Recommendations for the Management of Alcohol-Associated Vitamin and Electrolyte Deficiencies in the ICU



Treatment of Metabolic Derangements

- Saline (or dextrose in water solution) - 1L
- Thiamine 100 mg
- Folic acid 1 mg
- Low-dose Multivitamin -1 ampule
- Magnesium sulfate 3 g





Wernicke Encephalopathy

- 1. Dietary deficiencies
- 2. Oculomotor abnormalities
- 3. Cerebellar dysfunction
- 4. Confusion or memory impairment





Evaluation & Metabolic Derangements Peals & Pitfalls



- EtOH use is associated with AKA and hyperlactemia
- Most admitted AWS patients benefit from IVF, calories, thiamine, and magnesium



- Don't miss Wernicke encephalopathy
- It's probably not a toxic alcohol





Treatment



Abstinent





Loss of Uncontrolled inhibition excitation



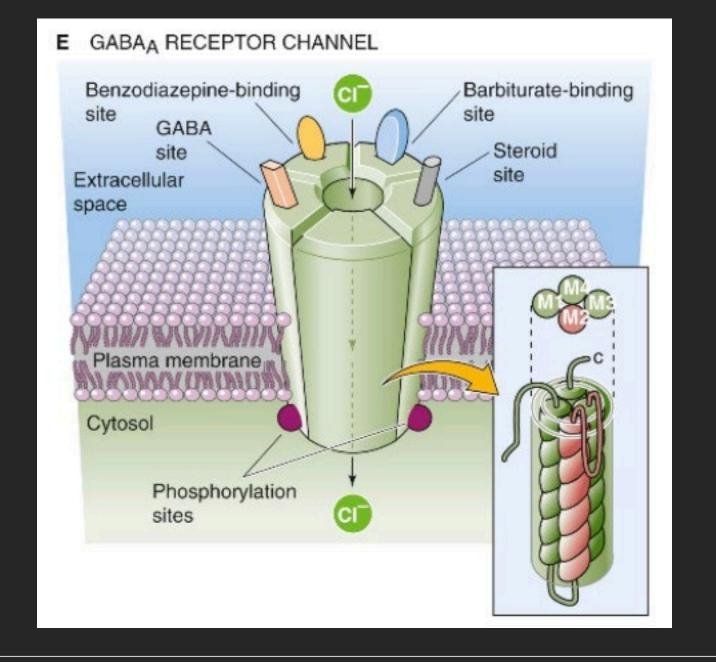
= GABA receptor with α_{1} subunit



= GABA receptor with $\alpha_{\rm 4}$ subunit



= NMDA receptor



Treatment of the Acute Alcohol Withdrawal State: A Comparison of Four Drugs

| | Inci | idence of Deliri | um Tremens and (| Convulsions | Amer. J. Psycho | iat. 125: 12, June 1969 |
|------------------------------------|------------------------------------|---------------------------------|--------------------------|--------------------|----------------------|-------------------------|
| | | | DRUG GROUP | | | |
| DISTURBANCE | CHLORDIA- ZEPOXIDE (N = 103) | CHLORPRO- Mazine (N = 98) | HYDROXYZINE (N = 103) | THIAMINE (N = 103) | PLACEBO (N = 130) | TOTAL (N = 537) |
| Delirium tremens | 1 | 4 | 2 | 4 | 7 | 18 |
| Convulsions Delirium tremens | 1 | 9 | 6 | 7 | 8 | 31 |
| and convulsions Total (percent in | 0 | 3 | 2 | 0 | 1 | 6 |
| parentheses) | 2 (2) | 16 (16) | 10 (10) | 11 (11) | 16 (12) | 55 (10) |



Treatment

- Diazepam (starting dose 10 mg IV)
- Lorazepam (starting dose 2 mg IV)
- Midazolam (starting dose 5 mg IM)

Re-dosing

• Goal: sleepy but breathing

| RASS (Richmond Agitation Sedation Scale) | | | | |
|--|----------------------|--|--|--|
| 4 | Combative | Overtly combative, violent, immediate danger to staff | | |
| 3 | Very agitated | Pulls or removes tubes or catheters; aggressive | | |
| 2 | Agitated | Frequent non-purposeful mvmt, fights ventilator | | |
| 1 | Restless | Anxious but movements not aggressive or vigorous | | |
| 0 | Alert and calm | | | |
| -1 | Drowsy | Sustained awakening to voice (≥10sec) | | |
| -2 | Light sedation | Briefly awakens with eye contact to voice (<10 sec) | | |
| -3 | Moderate sedation | Movement or eye opening to voice but no eye contact | | |
| -4 | Deep sedation | No response to voice but movement or eye opening to physical stimulation | | |
| -5 | Cannot be aroused | No response to voice or physical stimulation | | |



Phenobarbital

GABA_A agonist
 Binds at non-BZD site
 Opens Cl⁻ channel more effectively
 May have efficacy in BZD-resistant AWS



Phenobarbital

1 mL Single-dose Vial

1 mL Single-dose Vial





SAEM GRACE: Phenobarbital for alcohol withdrawal management in the emergency department: A systematic review of direct evidence

• "evidence that exists generally suggests that [phenobarbital] is a reasonable and appropriate approach [to treat AWS]."

Guidelines for Reasonable and Appropriate Care in the Emergency Department (GRACE-4): Alcohol use disorder and cannabinoid hyperemesis syndrome management in the emergency department

• "In adult ED patients (over the age of 18) with moderate to severe alcohol withdrawal who are being admitted to hospital, we suggest using phenobarbital in addition to benzodiazepines compared to using benzodiazepines alone."



Phenobarbital

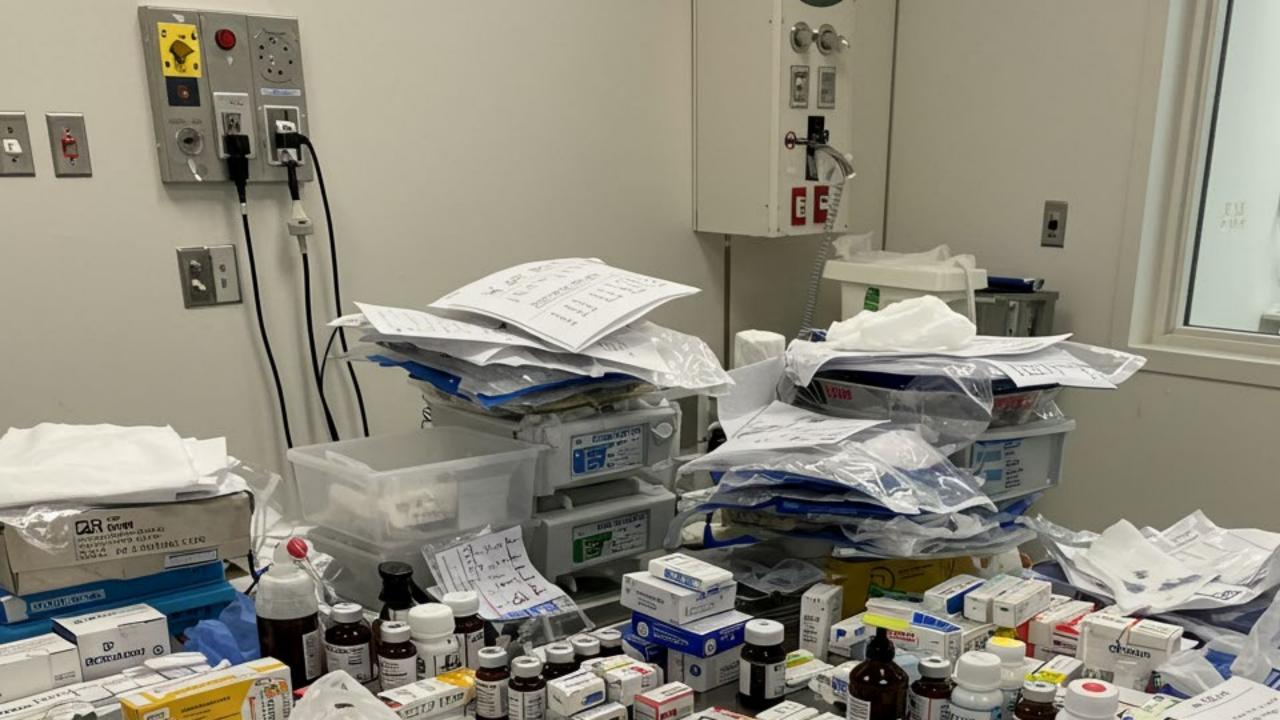
Anti-NMDA?
 Only at [PHB] > 170 ug/mL (therapeutic 10-40 ug/mL)

Figure 4-5 Benzodiazepines and Their Phenobarbital Withdrawal Equivalents

https://www.ncbi.nlm.nih.gov/books/NBK64116/

| Generic name | Trade name | Therapeutic dose range (mg/day) | Dose equal to 30mg of phenobarbital for withdrawal (mg) $^{\!**}$ | Phenobarbital conversion constant | |
|------------------|------------|---------------------------------|---|-----------------------------------|--|
| Benzodiazepines | | | | | |
| alprazolam | Xanax | 0.75–6 | 1 | 30 | |
| chlordiazepoxide | Librium | 15–100 | 25 | 1.2 | |
| diazepam | Valium | 4–40 | 10 | 3 | |
| lorazepam | Ativan | 1–16 | 2 | 15 | |

260 mg PHB ~ 87 mg diazepam ~ 17 mg lorazepam





Adjuncts











Treatment: Peals & Pitfalls



- GABA_A medications are cornerstone
- PHB might reduce need for ICU admission or intubation vs BZD



- Don't be afraid of large doses
- Don't rely on non-GABA medications

