

# Alcohol Use Disorder: Screening, Diagnosis, and Treatment in the Outpatient Clinic

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# Disclosure

- I have no actual or potential conflicts of interest in relation to this program/presentation.
- I will be discussing off-label use of gabapentin to treat AUD. I will not be discussing any other unapproved uses of pharmaceuticals or devices.

# Learning Objectives:

At the conclusion of this activity, participants will be able to:

- Accurately and appropriately screen for and diagnose alcohol use disorder (AUD)
- List the standard-of-care, FDA-approved medication treatments for alcohol use disorder
- List behavioral interventions for alcohol use disorder
- Implement harm reduction for patients who are not currently interested in discontinuing substance use

# Background:

- 28.9 million, or 10.2%, of US adults (age 12+) have an alcohol use disorder (AUD)
- 178,000 people die from alcohol related causes annually
- 4<sup>th</sup> leading cause of preventable death in US
- 7.9% of people with AUD receive ANY treatment
- ~2% of people with AUD receive an FDA approved medication
- Alcohol contributes to ~7.1% of all ED visits nationwide
- Only 1 out of 6 US adults report ever having been asked by a healthcare professional about their drinking behavior

www.niaaa.nih.gov

Kranzler and Soyka. JAMA. 2018. 320 (8): 815-824 Centers for Disease Control and Prevention (CDC). Alcohol and Public Health: Alcohol-Related Disease Impact (ARDI). Annual Average for United States 2011–2015 Alcohol-Attributable Deaths Due to Excessive Alcohol Use, All Ages. Available at: <u>https://nccd.cdc.gov/DPH\_ARDI/Default/Default/Default.aspx</u>.

### Alcohol Use in Primary Care





Each drink shown above represents one U.S. standard drink and has an equivalent amount (0.6 fluid ounces) of "pure" ethanol. Each beverage portrayed above represents one standard drink (or one alcoholic drink equivalent), defined in the United States as any beverage containing 0.6 fl oz or 14 grams of pure alcohol. The percentage of pure alcohol, expressed here as alcohol by volume (alc/vol), varies within and across beverage types. Although the standard drink amounts are helpful for following health guidelines, they may not reflect customary serving sizes.

"Rethinking Drinking." NIAAA. Publication No. 21-AA-3770. Revised July 2022.



## Learning Objectives:

• Accurately and appropriately screen for alcohol use disorder (AUD)

# Single Item Screening Question

We ask all our adult patients about substance use and mood because these factors can affect your health. Please ask your doctor if you have any questions. Your answers on this form will remain confidential

Alcohol: One drink =



### United States AUDIT

1.	How often do you have a drink containing alcohol?	Never	Less than monthly	Monthly	Weekly	2 -3 times a week	4 - 6 times a week	Daily
2.	How many drinks containing alcohol do you have on a typical day when you are drinking?	1 drink	2 drinks	3 drinks	4 drinks	5-6 drinks	7-8 drinks	10 or more drinks
3.	How often do you have X (5 for men; 4 for women & men over age 65) or more drinks on one occasion?	Never	Less than monthly	Monthly	Weekly	2-3 times a week	4-6 times a week	Daily
4.	How often during the last year have you found that you were not able to stop drinking once you had started?	Never	Less than monthly	Monthly	Weekly	Daily or almost daily		
5.	How often during the last year have you failed to do what was normally expected of you because of drinking?	Never	Less than monthly	Monthly	Weekly	Daily or almost daily		
6.	How often during the last year have you needed a first drink in the morning to get yourself going after a heavy drinking session?	Never	Less than monthly	Monthly	Weekly	Daily or almost daily		AUDIT:
7.	How often during the last year have you had a feeling of guilt or remorse after drinking?	Never	Less than monthly	Monthly	Weekly	Daily or almost daily		0 - 6: Women; All ages >65 0 - 7: Men <65 <b>II – Risky</b>
8.	How often during the last year have you been unable to remember what happened the night before because of your drinking?	Never	Less than monthly	Monthly	Weekly	Daily or almost daily		7 - 15: Women; All ages >65 8 - 15: Men <65 III – Harmful
9.	Have you or someone else been injured because of your drinking?	No		Yes, but not in the past year		Yes, in the past year		16 – 19: All adults IV – Severe ≥20: All adults
10.	Has a relative, friend, doctor, or other health care worker been concerned about your drinking or suggested you cut down?	No		Yes, but not in the past year		Yes, in the past year		
		0	1	2	3	4	5	6

# Brief Intervention and Referral to Treatment

US AUDIT Score	Risk Zone	Recommended Intervention
0 – 6: Women; All ages ≥65 0 – 7: Men <65	Low	Positive reinforcement, alcohol education
7 – 15: Women; All ages ≥65 8 – 15: Men <65	Risky	Brief intervention, alcohol education, provide patient feedback on risks Consider: DSM-V assessment for AUD
16 – 19: All adults	Harmful	Brief intervention, continued monitoring and follow up Consider: DSM-V assessment for AUD, referral to treatment, pharmacotherapy
≥20: All adults	Severe	Brief intervention, continued monitoring and follow up, DSM-V assessment for AUD, referral to treatment, pharmacotherapy, consider medically managed withdrawal

Johnson, Vinson and Seale. Alcohol Clin Exp Res, 2013.

Centers for Disease Control and Prevention, Planning and Implementing Screening and Brief Intervention for Risky Alcohol Use: A Step-by-Step Guide for Primary Care Practices. 2014. https://auditscreen.org/about/audit-decision-tree/



# Learning Objectives:

• Accurately and appropriately diagnose alcohol use disorder (AUD)

### DSM-5



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#### 11 Criteria

5 Main Categories:

- Craving
- Loss of **C**ontrol
- **C**onsequences
  - **C**ompulsion
- Pharmacologic

9 Substances\*

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The 4 C's

### DSM-5 Criteria for Alcohol Use Disorder

Loss of Control/Craving	Pharmacological				
1. Using in larger amounts or for longer than intended	10. Tolerance*				
2. Repeated unsuccessful efforts to cut back or control use	11. Withdrawal*				
3. Great deal of time spent using, obtaining, recovering from use					
4. Craving or strong desire to use					
Consequences/Compulsive Use					
5. Recurrent use in physically hazardous situations					
6. Recurrent/persistent physical or psychological difficulties from use					
7. Continued use despite social problems/interpersonal issues due to use					
8. Failure to fulfill major obligations at work/school/home due to recurrent use					
9. Important things given up or reduced due to continued use					

Diagnostic and Statistical Manual of Mental Disorders 5, APA, 2013.

### AUD Severity



### AUD Remission Status

- Early remission: >3 months but <12 months
- Sustained remission: ≥12 months



# Learning Objectives:

- List the standard-of-care, FDA-approved medication treatments for alcohol use disorder
- List behavioral interventions for alcohol use disorder
- Implement harm reduction for patients who are not currently interested in discontinuing substance use

# Determine Patients' Treatment Goals

- Determination of Initial Treatment Goals
  - Initial goals of treatment of alcohol use disorder
    - abstinence from alcohol use
    - reduction or moderation of alcohol use
    - other elements of harm reduction

### Starting the Conversation

- Approach it with non-stigmatizing attitude and language
  - Normalize screening
- Ask permission
- Highlight why it is important to discuss
  - 4<sup>th</sup> leading cause of preventable death
  - Goal is patient health reduce morbidity and mortality
  - Effective treatments

# AUD Treatment - Behavioral Interventions

- Offer one or more of the following interventions considering patient preference and provider training/competence:
  - For brief interventions:
    - Motivational Interviewing
  - For comprehensive interventions:
    - Talk Therapy
    - Mutual Aid (SMART Recovery, 12-Step approach)
    - Contingency Management

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### FRAMES Model for Brief Intervention

- **Feedback** "you scored in the "risky" category regarding alcohol use, this may put you at higher risk for alcohol-related harms including ... "
- **Responsibility** (placed on patient)- "Do you want to make changes to the amount you drink?"
- Advice "Based on your age, it's recommended to have no more than x drinks per week"
- Menu "Here are the treatment options available..."
- **Empathy** Approach the patient in a nonjudgemental way
- **Self-Efficacy** Empower them to make change

### Psychosocial interventions for AUD

Talk Therapy

- Cognitive Behavioral Therapy (CBT)
- Dialectical Behavioral Therapy (DBT)
- Acceptance and Commitment Therapy (ACT)

Self-help/Peer Recovery Peer Recovery Specialists Contingency Management



This Photo by Unknown Author is licensed under CC BY-NC-ND

UpToDate. Accessed 1/2021.

### Self-help/Peer Recovery

- Alcoholics Anonymous <u>https://www.aa.org/</u>
- LifeRing <u>https://lifering.org/</u>
- Refuge Recovery <u>https://www.refugerecovery.org</u>/
- Recovery Dharma <u>https://recoverydharma.org</u>/
- SMART Recovery <u>https://www.smartrecovery.org</u>/
- Women for Sobriety <u>https://womenforsobriety.org</u>/

### Treatment Levels for AUD

- Medically Managed Withdrawal (Detox)
  - Manage acute withdrawal syndromes
  - 3-7 day stays
  - 24/7 nursing
- Residential Treatment
  - Sleep at facility; must be able to complete ADLs independently; minimal medical staff
  - Programing 6-8 hours daily
  - 30-90 days
  - · Cannot be in withdrawal or actively intoxicated
- Intensive Outpatient Treatment
  - In-Person and Virtual Options
  - Few days a week, few hours each day
  - Cannot be in withdrawal or actively intoxicated
- 1:1 Counseling/Therapy
  - CBT
  - ACT
  - DBT

### Alcohol Withdrawal

- Patients may experience alcohol withdrawal syndrome with sudden discontinuation of drinking.
- Severe forms of alcohol withdrawal include seizures, hallucinations, delirium, and rarely death.
- Risk factors for severe withdrawal include:
  - $\odot$  >8 drinks per day
  - $\odot$  History of severe withdrawal in the past year
  - Co-occurring unstable medical and psychiatric conditions (i.e., epilepsy, uncontrolled diabetes)
  - $\circ$  Physiologic dependence on other substances

Tiglao, Am Fam Physician. 2021

### Alcohol Withdrawal

- Patients should be counseled on the risk for alcohol withdrawal if they plan to stop drinking.
- Patients at high risk for alcohol withdrawal syndrome should be referred to inpatient withdrawal management (detox/inpatient) or to the emergency room/hospital.
  - CIWA triggered symptom management
  - Don't forget to start MAUD!



### General Approach to AUD Treatment – Pharmacotherapy

- Tier 1
  - naltrexone or
  - acamprosate
- Tier 2
  - disulfiram
  - Other off-label medications:
    - gabapentin



### Medication Overview

	Naltrexone	Acamprosate	Disulfiram $\Theta_{\bigcirc}$
Mechanism of Action	Blocks opioid receptors	NMDA receptor modulator	Inhibitor of aldehyde dehydrogenase
Dose	50 mg PO daily 380 mg IM monthly	666 mg PO TID	250 mg PO
Contraindications	Concurrent opioids, decompensated liver failure	CrCl <30	Med interactions, recent drink, severe CAD, psychosis, liver dysfunction
Side Effects	Flu-like symptoms	GI upset, diarrhea	Drowsiness, optic neuritis, peripheral neuropathy, hepatotoxicity

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	Naltrexone
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Dose	50 mg PO daily 380 mg IM monthly
Contraindications	Concurrent opioids, decompensated liver failure
Side Effects	Flu-like symptoms

Jonas DE et al., Pharmacotherapy for adults with alcohol use disorders in outpatient settings: a systematic review and meta-analysis. JAMA. 2014 May 14;311(18):1889-900.

Bahji A, Bach P, Danilewitz M, Crockford D, Devoe DJ, El-Guebaly N, Saitz R. Pharmacotherapies for Adults With Alcohol Use Disorders: A Systematic Review and Network Meta-analysis. J Addict Med. 2022 Nov-Dec 01;16(6):630-638.

Adapted with permission from Kristin Prewitt, MD, MPH

#### FDA approved since:

1994 (oral) 2005 (IM)

#### Evidence for use:

Systematic Review and Meta-Analysis (Jonas et al., 2014)

53 studies on naltrexone (n= 9140), 44 RCTs

Return to any drinking, NNT = 20 (95% CI 11 to 500)

Return to heavy drinking, NNT= 12 (95% CI 8 to 26)

Systematic Review and Network Meta-Analysis (Bahji et al., 2022)

54 RCTs on oral naltrexone

Improved total abstinence, RR = 1.15 (95% CI 1.01-1.32)

Reduced heavy drinking, RR = 0.81 (95% CI 0.73-0.90)

	Naltrexone
Mechanism of Action	Blocks opioid receptors
Dose	50 mg PO daily 380 mg IM monthly
Contraindications	Concurrent opioids, decompensated liver failure
Side Effects	Flu-like symptoms

#### Mechanism of Action:

Blocks opioid receptors resulting in reduced dopamine release at the nucleus accumbens

#### Dose:

50mg PO daily 380mg IM monthly



	Naltrexone
Mechanism of Action	Blocks opioid receptors
Dose	50 mg PO daily 380 mg IM monthly
Contraindications	Concurrent opioids, decompensated liver failure
Side Effects	Flu-like symptoms

#### Starting:

- Moderate to severe alcohol use disorder

- LFTs < 5x ULN and no

decompensated liver failure

- 7 days since last short-acting opioid or

10 days since last long-acting opioid

#### Monitoring: - LFTs



- Stop if LFT > 5x ULN

#### Side effects:

- Nausea, vomiting, headache, and dizziness



### Medication Overview

	Naltrexone	Acamprosate	Disulfiram
	$\square$	(0,00) (0,00) (0,00)	$\Theta_{\Theta}$
Mechanism of Action	Blocks opioid receptors	NMDA receptor modulator	Inhibitor of aldehyde dehydrogenase
Dose	50 mg PO daily 380 mg IM monthly	666 mg PO TID	250 mg PO
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Adapted with permission	from Kristin Prewitt, MD, MPI	4	

### Acamprosate



Jonas DE et al. Pharmacotherapy for adults with alcohol use disorders in outpatient settings: a systematic review and meta-analysis. JAMA. 2014 May 14;311(18):1889- '900.

Bahji A, Bach P, Danilewitz M, Crockford D, Devoe DJ, El-Guebaly N, Saitz R. Pharmacotherapies for Adults With Alcohol Use Disorders: A Systematic Review and Network Meta-analysis. J Addict Med. 2022 Nov-Dec 01;16(6):630-638.

Adapted with permission from Kristin Prewitt, MD, MPH

#### FDA approved for AUD in 2004

#### **Evidence for use:**

Systematic Review and Meta-Analysis (*Jones et al., 2014*) 27 studies on acamprosate (n= 7519), 22 RCTs

Return to any drinking, NNT = 12 (CI 8 to 26)

Return to heavy drinking, no improvement

Systematic Review and Network Meta Analysis (*Bahji et al., 2022*) 35 RCTs on acamprosate

> Improved total abstinence RR = 1.33 (95% CI 1.15-1.54)

Reduced heavy drinking RR = 0.78 (95% CI 0.70-0.86)

Better results for maintenance of abstinence rather than reduction in drinking by non-abstinent patients

### Acamprosate

(0) (0) (0) (0) (0) (0) (0) (0) (0) (0)		
	Acamprosate	
Mechanism of Action	NMDA receptor modulator	
Dose	666 mg PO TID	
Contraindications	CrCl <30	
Side Effects	GI upset, diarrhea	
ins. Neurobiology and		

Jeffery Wilkins. Neurobiology an Pharmacotherapy for Alcohol Dependence.

Adapted with permission from Kristin Prewitt, MD, MPH

#### **Mechanism of Action:**

- Amino acid derivative
- NMDA receptor modulator that acts by changing balance between GABA and glutamate
- Drug elimination renal

#### Dose:

- 666 mg PO TID - 333 mg PO TID if moderate renal impairment (CrCl 30-50)

### Acamprosate

	0;0 0;0	
	Acamprosate	Starting: -Not a candidate
Mechanism of Action	NMDA receptor modulator	<ul> <li>Already on TID</li> <li>May be more e has begun abstir</li> </ul>
Dose	666 mg PO TID	Monitoring:
Contraindications	CrCl <30	- Contraindicate - Monitor Cr/CrC
Side Effects	GI upset, diarrhea	<u>Side Effects:</u> diarrhea, GI ups

e for naltrexone (liver medications

effective after patient inence

d in CrCl<30

CI periodically



### Naltrexone vs Acamprosate

No statistically significant difference in outcomes for return to any drinking or return to heavy drinking in individuals receiving behavioral interventions and acamprosate or behavioral interventions and naltrexone

Table 2. Summary of Findings and Strength of Evidence From Double-Blind Randomized Clinical Trials Directly Comparing Acamprosate and Naltrexone<sup>a</sup>

Outcome	No. of Studies	No. of Participants <sup>b</sup>	Results Effect Size (95% CI) <sup>c</sup>	Strength of Evidence
Return to any drinking	3	800	RD: 0.02 (-0.03 to 0.08)	Moderate
Return to heavy drinking	4	1141	RD: 0.01 (-0.05 to 0.06)	Moderate

Jonas DE et al. Pharmacotherapy for adults with alcohol use disorders in outpatient settings: a systematic review and meta-analysis. JAMA. 2014 May 14;311(18):1889-900.

### Medication Overview

	Naltrexone	Acamprosate	Disulfiram	
	$\mathcal{D}$	(0) (0) (0) (0) (0) (0) (0) (0) (0) (0)	90	
Mechanism of Action	Blocks opioid receptors	NMDA receptor modulator	Inhibitor of aldehyde dehydrogenase	
Dose	50 mg PO daily 380 mg IM monthly	666 mg PO TID	250 mg PO	
Contraindications	Concurrent opioids, decompensated liver failure	CrCl <30	Med interactions, recent drink, severe CAD, psychosis, liver dysfunction	
Side Effects	Flu-like symptoms	GI upset, diarrhea	Drowsiness, optic neuritis, peripheral neuropathy, hepatotoxicity	
			Adapted with permission	from Kristin Prewitt, MD, MPH

### Medication Overview

	(0;0) (0;0)		
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### Disulfiram

	Disulfiram $\Theta_{\bigcirc}$	
Mechanism of Action	Inhibitor of aldehyde dehydrogenase	
Dose	250 mg PO	
Contraindications	Med interactions, recent drink, severe CAD, psychosis, liver dysfunction	
Side Effects	Drowsiness, optic neuritis, peripheral neuropathy, hepatotoxicity	

Bahji A, Bach P, Danilewitz M, Crockford D, Devoe DJ, El-Guebaly N, Saitz R. Pharmacotherapies for Adults With Alcohol Use Disorders: A Systematic Review and Network Meta-analysis. J Addict Med. 2022 Nov-Dec 01;16(6):630-638.

Adapted with permission from Kristin Prewitt, MD, MPH

Second line <u>FDA approved</u> for alcohol dependence since 1949

#### **Evidence for Use:**

Heterogenous efficacy - may be best for patients in supervised settings

 Recent Systematic Review & Meta-analysis (Bahji et al., 2022),13 studies

> *Total Abstinence* RR 1.71, CI 1.39-2.10

Reduced Heavy Drinking RR 0.19, CI 0.1-0.25

### Disulfiram



### Disulfiram

	<u>Starting:</u>			
	Disulfiram ⊖	<ul> <li>Abstinence at least 12 hours and/or breath or blood alcohol level is zero</li> <li>No severe CAD, advanced liver disease, pregnancy/nursing, psychosis</li> </ul>		
Mechanism of	Inhibitor of aldehyde dehydrogenase	<ul> <li>Informed consent (severity of reaction, 2 week tail effect)</li> </ul>		
Action		<ul> <li>Many medication interactions</li> </ul>		
Dose	250 mg PO	- chlordiazepoxide, diazepam, warfarin, TCAs, metronidazole, phenytoin, theophylline, isoniazid, rifampin, among others		
Contraindications	Med interactions, recent drink, severe CAD, psychosis, liver dysfunction	Monitoring: Baseline LFTs, monitor periodically		
Side Effects	Drowsiness, optic neuritis, peripheral neuropathy, hepatotoxicity	Drowsiness is most common; all others are rare but serious, including hepatotoxicity (1:25,000), psychosis		

### Medication Overview

	Naltrexone	Acamprosate	Disulfiram $\Theta_{\bigcirc}$
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Side Effects	Flu-like symptoms	GI upset, diarrhea	Drowsiness, optic neuritis, peripheral neuropathy, hepatotoxicity

### **Off-Label Medications**

- Gabapentin
  - Mechanism of action: GABA analog reducing glutamate and increasing GABA
  - Dose: 600mg TID
  - Contraindications: myasthenia gravis, myoclonus, CrCl <30mL/min
  - <u>Side Effects</u>: dizziness, drowsiness, respiratory depression (when used with other CNS depressants, e.g. opioids, alcohol, benzodiazepines), nausea, xerostomia, nystagmus
  - <u>Efficacy</u>: decreased drinking days, may be best for patients with withdrawal symptoms

Total Abstinence, RR = 1.66 (95% CI, 1.04-2.67)

Bahji A, Bach P, Danilewitz M, Crockford D, Devoe DJ, El-Guebaly N, Saitz R. Pharmacotherapies for Adults With Alcohol Use Disorders: A Systematic Review and Network Meta-analysis. J Addict Med. 2022 Nov-Dec 01;16(6):630-638.

### Harm Minimization/Safer Drinking





# Questions?

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