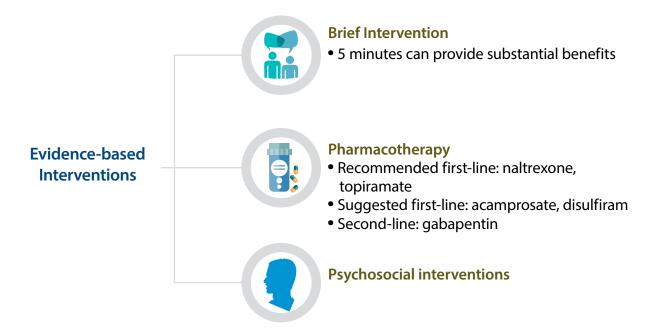


Alcohol Use Disorder (AUD) Leading the Charge in the Treatment of AUD



Key Takeaways

There are a variety of effective evidence-based pharmacotherapy and psychosocial interventions available to treat AUD.



All VA clinicians have the opportunity to be advocates for our Veterans' recovery from unhealthy alcohol use. Identifying alcohol-related risk and encouraging Veterans to obtain treatment for AUD is the right investment of VA resources.

Alcohol Use Disorder (AUD) Leading the Charge in the Treatment of AUD

A VA Clinician's Guide



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Risks of Excessive Alcohol

Alcohol consumption is one of the leading preventable causes of death in the U.S.¹

Rates of heavy drinking and binge drinking are increasing in the U.S. correlating with increasing alcohol related death rates.

Alcohol related deaths increased by 49% between 2006 and 2019.^{2,3}

- Males by 35% (2005-2018)
- Females by 76% (2000-2018)



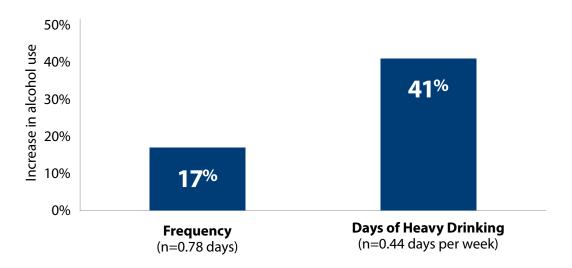
cdc.gov/alcohol

Alcohol Use Among Women is Increasing!



Studies show that among U.S. women who drink, about **one in four has engaged in binge drinking in the last month**, averaging about three binge episodes per month and five drinks per binge episode.⁴

Figure 1. Significant Increase in Alcohol Use Among Women 2019 – 2020⁵



Overall, in 2020 frequency of alcohol use increased by 0.74 days (14%) over the 2019 baseline of 5.48 days per week. Several subgroups showed significant increases: women (0.78 days, 17% increase), adults aged 30 to 59 (0.93 days, 19% increase), non-Hispanic whites (0.66 days, 10% increase). Among women, days of heavy drinking increased 0.18 days from a baseline of 0.44 days per week in 2019 (41% increase); alcohol-related problems measured on a validated, commonly used scale also grew (39% increase).⁵

Table 1. Drinking Levels⁶

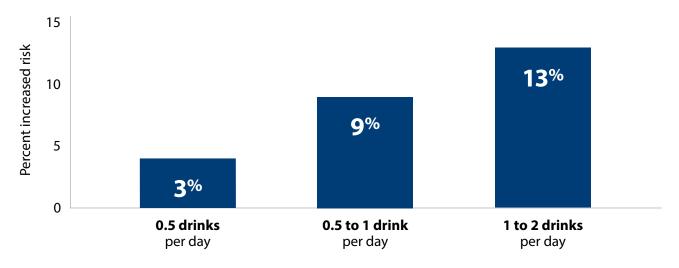
	Drinking in moderation	Binge drinking*	Heavy/Excessive alcohol use
Males	0 – 2 drinks per day	≥ 5 drinks at the same time or within a couple of hours of each other	> 4 drinks on any day or > 14 drinks per week
Females and/or Age > 65	0 – 1 drink per day	≥ 4 drinks at the same time or within a couple of hours of each other	> 3 drinks on any day or > 7 drinks per week

^{*}In the past month. Older adults and women have increased sensitivity to alcohol because they typically metabolize it at a slower rate. This makes them more susceptible to the adverse consequences associated with alcohol consumption.

Alcohol use is a modifiable risk factor for many medical and mental health conditions

For example, breast cancer risk is associated with even low levels of consumption and increases as consumption increases.⁷

Figure 2. Risk of Breast Cancer Increases as Alcohol Consumption Increases⁷



Excessive or heavy drinking (table 1) accounts for most of the morbidity and mortality seen and has immediate and long-term effects that increase the risk of many harmful health conditions.⁸

Figure 3. Health Risks Associated with Heavy or Excessive Alcohol Use⁹⁻¹⁴

Cancer

- Breast
- Colorectal
- Esophagus
- Liver
- Larynx
- Pancreatic
- Oropharynx

Vulnerability to infections

- Upper respiratory
- Pneumonia
- Tuberculosis



Neuropsychiatric disorders

- Anxiety
- Depression
- Insomnia
- Dementia
- Neurocognitive impairment
- Suicide risk



Other chronic diseases

- High blood pressure
- Cardiovascular diseases (e.g., stroke, cardiomyopathy, atrial fibrillation)
- Neuropathy



Gastrointestinal diseases

- Digestive issues
- Liver cirrhosis
- Pancreatitis



Increased reproductive problems

- Male: Erectile dysfunction, decreased fertility
- Female: Risk of birth defects

Cause for concern



Women are:

- The fastest-growing segment of alcohol consumers in the United States⁵
- At greater risk for adverse health outcomes associated with excessive

alcohol use15

- Liver: risk of cirrhosis in women is greater than in men and increases with as little as one drink a day and increases exponentially with the number of drinks consumed per day^{7,16}
- Brain: faster onset of alcohol-related cognitive decline and shrinkage of the brain^{7,17}
- Heart: increased risk for cardiomyopathy at lower levels of consumption and over fewer years of drinking⁷
- Breast Cancers: increased risk associated with even low levels of consumption⁷

Addressing Implicit Bias and Stigma

Society often views substance use disorders as a moral weakness. As a result, people facing these issues are often plagued with shame. If stigma is experienced individuals are less likely to seek out treatment or remain engaged in treatment.^{18,19}

Societal views can shape the values and beliefs of healthcare professionals, leading to stigma and stereotyping. Stereotyping often occurs without our conscious knowledge and is known as implicit bias.

Stigma towards substance use disorders is common amongst healthcare professionals.²⁰ To provide the best care, become aware of your own biases and seek ways to mitigate them.²¹

Start by thinking of AUD as a disease like any other chronic illness with physical components, such as diabetes. This helps to shift the focus from the behavior of a person to the disease.



Did you know?

Implicit biases exist in all of us, even in those who are committed to being fair and impartial.



Use the following steps to mitigate the effects of your own unconscious bias and model behaviors that set the tone and expectations of others.²²

Figure 4. Take CHARGE² To Mitigate Your Own Bias²²

C

change your context: is another perspective possible?

- Avoid judgment
- Listen actively with curiosity
- Be open-minded about the patient's story



BE HONEST WITH YOURSELF: ACKNOWLEDGE AND BE AWARE

- Acknowledge your preconceived judgments
- Do not make or be afraid to correct assumptions (say, "oops")



AVOID BLAMING YOURSELF: KNOW THAT YOU CAN DO SOMETHING ABOUT IT

- We all harbor implicit associations, this allows us to function efficiently but can make us more prone to errors
- Stereotyping of patients can lead to suboptimal care resulting in negative healthcare outcomes



REALIZE WHEN YOU NEED TO SLOW DOWN

• Slowing down may help reframe your perspective



GET TO KNOW PEOPLE YOU PERCEIVE AS DIFFERENT FROM YOU

• Building relationships is key to understanding



ENGAGE: REMEMBER WHY YOU ARE DOING THIS

- People can and do recover
- Do not give up
- Even small steps are a success



EMPOWER PATIENTS (AND PEERS)

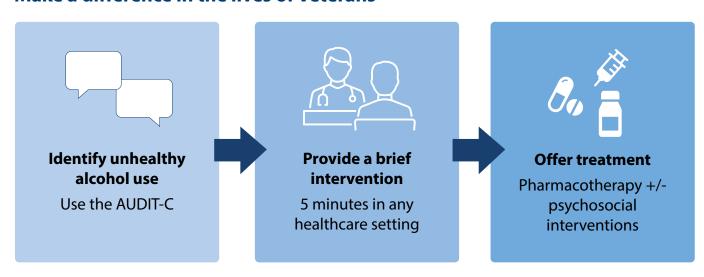
- Set meaningful short-term goals with the patient
- Accept that patients may not be ready and be willing to offer help again
- Step up to intervene if you witness bias

Language used to describe Veterans with substance use disorders can perpetuate stigma and stereotypes. You can counter stigma by using accurate, nonjudgmental, personcentered, language to describe AUD, those it affects, and its treatment.^{19,23}

Table 2. Changing the Conversation 19,23

Instead of	Use	Why
Alcoholic Drunk User	 Person with alcohol use disorder Person who misuses alcohol Person who engages in unhealthy and/or hazardous alcohol use 	 Person-first language: Indicates that the person has an illness vs. them being the illness Indicates that the disease is not permanent Avoids negative associations, punitive attitudes, and individual blame
Abuse	 Misuse Harmful use Unhealthy use Hazardous use Problem use Risky use 	The word abuse is associated with negative judgments and punishments
Habit	Active addictionSubstance use disorder	Habit signifies that resolution of the disorder is a matter of willpower and being able to stop the habitual behavior May undermine the seriousness of the disorder

Make a difference in the lives of Veterans

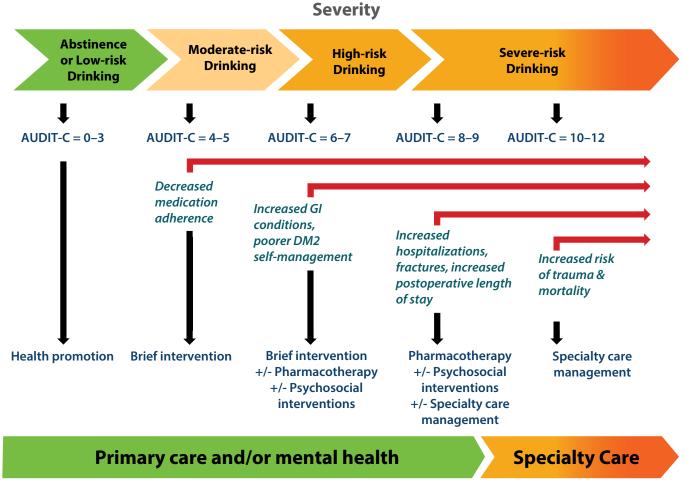


Identifying Unhealthy Alcohol Use

Screening

The Alcohol Use Disorders Identification Test-Consumption (AUDIT-C) is an alcohol screening assessment that can identify patients who are hazardous drinkers or have active alcohol use disorder.^{24,25} AUDIT-C scores have been correlated with rates of morbidity and mortality and can also be used as a tool to guide treatment.^{26,27}

Figure 5. Spectrum of Unhealthy Alcohol Use with AUDIT-C Score and Recommended Treatment and Setting Based on Severity²⁸⁻⁴¹



For VA, AUDIT-C score of 5 or more is considered positive for both men and women, and documentation of brief alcohol counseling is required. To better account for safer drinking levels in females, AUDIT-C item #3 is automatically tailored to the sex assigned at birth in the clinical reminder. This helps differentiate binge-drinking patterns in female Veterans and may improve detection of unhealthy drinking patterns. For males, item #3 queries consumption of \geq 6 drinks a day, and for females, it queries consumption of \geq 4 drinks.

DM2 = diabetes mellitus type II; GI = gastrointestinal.

Use the AUDIT-C to identify unhealthy alcohol use in our Veterans.

.....

Brief Alcohol Interventions

Screening for unhealthy alcohol use, followed by advice and education regarding alcohol-related risks, is effective at reducing drinking among individuals who have unhealthy drinking but do not have full-fledged AUD.^{24,42} For patients with an AUDIT-C of 4-7, offering a brief intervention has shown to:^{24,42,43}

- Decrease mean number of drinks per week
- Decrease number of heavy drinking episodes
- Increase the percentage of patients whose alcohol consumption is within recommended drinking limits
- Improve outcomes for all-cause mortality, hospitalization rates, and systolic blood pressure

Figure 6. Brief Interventions^{24,42,43}

Identify unhealthy alcohol use and provide at least 1 brief intervention lasting ≥5 minutes **ENHANCE REFER TO RAISE THE SUBJECT** PROVIDE FEEDBACK **NEGOTIATE PLAN** TREATMENT MOTIVATION Set a feasible Advise treatment Identify links Provide Ask permission to talk about between alcohol explicit advice drinking goal or suggest alcohol use. use and patient's to cut down and arrive at a referral, if shared decision. co-occurring and enhance appropriate Share you are health conditions. motivation to (e.g., AUDIT-C \geq 8). Encourage concerned they if present change and specificity are drinking (e.g., diabetes, decrease or (e.g., cutting above the mood disorders, abstain from recommended down to X insomnia). alcohol use. number of limits. Educate on safe drinks and Reflect patient's limits of alcohol responses that are documenting consumption. intended steps). consistent with changing drinking behavior.

Provide at least one 5-minute brief intervention to all Veterans with unhealthy drinking (AUDIT C 4-7).

Making the Diagnosis

Most Veterans who drink above the recommended limits (e.g., AUDT-C 4-7) will not have AUD. However, Veterans with AUDIT-C scores \geq 8 (>95% specificity DSM-5 severe AUD) are at a higher risk of AUD and need evaluation for diagnosis and management.^{43,44,45}

Table 3. Making the Diagnosis

ICD-10 Alcohol Dependence (3 or more criteria)	DSM-5 AUD Mild AUD = 2-3 symptoms, Moderate AUD = 4-5; Severe AUD = 6 or more
A. Persisting with drinking alcohol despite harmful consequences	1. Continued use despite social or interpersonal problems related to alcohol
	2. Continued use despite physical or psychological problem(s) related to alcohol
	3. Recurrent alcohol use in situations in which it is physically hazardous
B. Tolerance	4. Tolerance
C. Withdrawal symptoms or use of alcohol (or closely related substance) to relieve or avoid withdrawal symptoms	5. Withdrawal symptoms OR alcohol (or closely related substance – benzo) is taken to relieve or avoid withdrawal symptoms
D. Difficulties in controlling alcohol use; or a persistent desire or unsuccessful	6. Alcohol is often taken in larger amounts or over a longer period than was intended
efforts to reduce or control drinking	7. Persistent desire or unsuccessful efforts to cut down or control alcohol use
E. Higher priority given to alcohol use than other activities and obligations	8. Recurrent alcohol use resulting in a failure to fulfill obligations at home, school and work
A large amount of time spent in alcohol, related activity, or recovery	9. Large amount of time is spent obtaining alcohol, using alcohol, or recovering from its effects
.,,,,,	10. Activities given up because of alcohol use
F. Craving or compulsion	11. Craving or compulsion

ICD-10 Alcohol Abuse: Alcohol use that has clearly contributed to physical or psychological harm. ICD-10 Abuse = DSM-5 mild AUD; ICD-10 dependence = DSM-5 moderate and severe AUD

Alcohol Use Disorder Pharmacotherapy

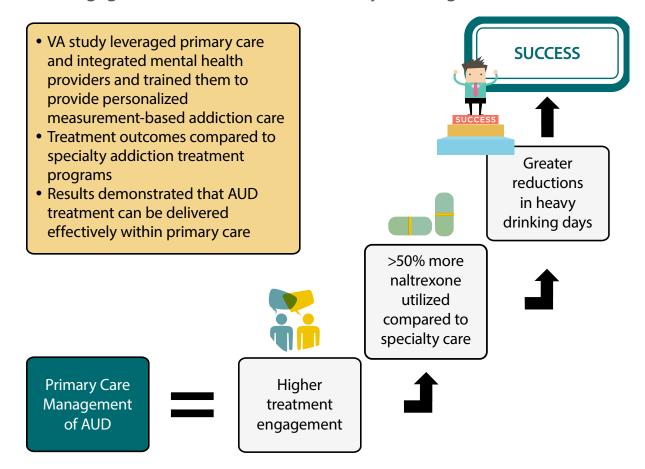
Medications to treat AUD are an effective treatment modality that can be utilized to reduce heavy drinking days, frequency of alcohol consumption, improve abstinence, and decrease cravings for alcohol.⁴³ They are often underutilized despite having greater efficacies than other medications used routinely as standards of care in other medical illnessess.^{46,47}

Offer Pharmacotherapy to patients with

- Moderate to severe AUD (AUDIT-C ≥8) OR
- AUDIT-C ≥6 and current alcohol use when contraindicated (e.g., previously diagnosed with AUD, presence of medical contraindications)

Medication treatment can be provided in primary care, general mental health, and specialty care settings.^{48–53} Primary care is the entry point for most people with AUD and is also the place where most clinical interventions should take place.^{48–53}

Figure 7. AUD Treatment in Primary Care Leads to Greater Rates of Treatment Engagement and Reductions in Heavy Drinking⁵¹



Management of AUD with pharmacotherapy in the primary care or general mental health setting should include medical management.

• Structured psychosocial intervention designed to be delivered by a medical professional (e.g., physician, nurse, physician assistant, clinical pharmacist practitioner) and is designed to support evidence-based medication treatment.^{24,43,49}

Figure 8. Components of Medical Management^{24,49*}

ENCOURAGE MONITOR EDUCATE Self-reported **Educate about AUD** To abstain from consequences and alcohol and use, urine drug treatments. other addictive test**, CDT, PEth, substances or set consequences, medication harm reduction adherence+, goals. treatment To attend mutual response, and help groups adverse effects. (community Praise small steps support groups towards recovery for recovery). goals. To adhere to Consider using prescribed medications. a measurementbased assessment To make lifestyle tool.++ changes that support recovery.

*Session structure varies according to the patient's substance use status and treatment adherence; **Ethyl glucuronide, ethyl sulfate (alcohol analytes) can be detected in the urine for 72 hours; *When adherence is inadequate develop a mutually agreed upon adherence plan; **Example: Brief Addiction Monitor (BAM); CDT = carbohydrate-deficient transferrin; PEth = phosphatidyl ethanol.

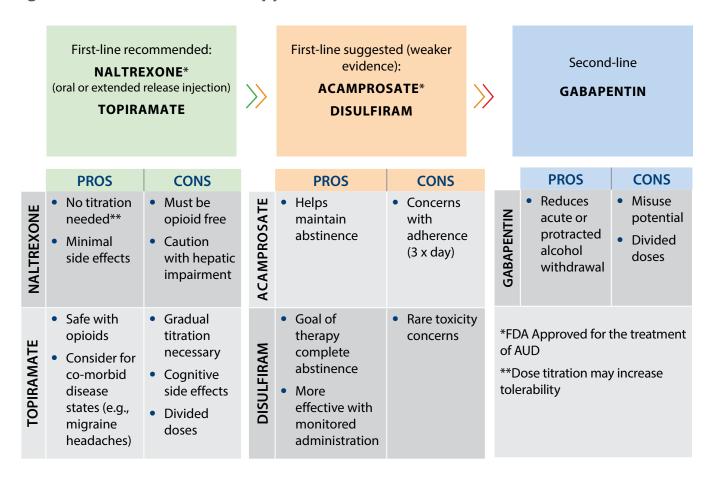
Not all patients will agree to stop drinking completely. If they are willing to reduce the amount of alcohol consumed this is a WIN and a positive step towards reducing alcohol-related problems Harm reduction focuses on strategies to reduce negative consequences of substance use. Consequences of substance use can be placed on a continuum from most to least harmful, and the goal is to move along this continuum by actions to reduce harm. Consider a harm reduction approach in individuals with fewer alcohol-related problems.

Work with the Veteran to develop alcohol consumption goals that focus on harm reduction strategies.

Example strategies:

- Have a plan to avoid or reduce intoxication
- Measure and document amount of alcohol consumed
- Arrange transportation to avoid driving intoxicated
- Carry condoms to facilitate safer sex

Figure 9. AUD Pharmacotherapy^{24,49,54}



Offer pharmacotherapy along with addiction-focused medical management to your Veterans with AUD.

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Length of Treatment

• ≥6 months; continuing pharmacotherapy • Greatest risk to return to drinking is up to and beyond one year may be during the first 90 days necessarv⁵⁴

First-line recommended pharmacotherapy for AUD

Naltrexone

Naltrexone is FDA approved and has the largest body of evidence to support its use. 43,49,54-60

Figure 10. Naltrexone

Naltrexone significantly reduces:

- Alcohol relapses
- Frequency and quantity of alcohol consumption
- Alcohol craving
- Health care utilization and cost

COMBINE Study

- Largest AUD multicenter trial in the U.S.
- When combined with medical management naltrexone reduced heavy drinking days

(HR, 0.72; 97.5% CI, 0.53-0.98; p = 0.02)

Meta-analysis

- Reduces the risk for heavy drinking (NNT = 8.1)
- Increases abstinence from alcohol (NNT = 17.4)

Naltrexone is also available as an monthly extended-release injection.

Naltrexone has real-world efficacy, is easy to use, and well tolerated. This medication should be considered the preferred first-line treatment for Veterans with AUD who are not on opioids and do not have severe liver disease.

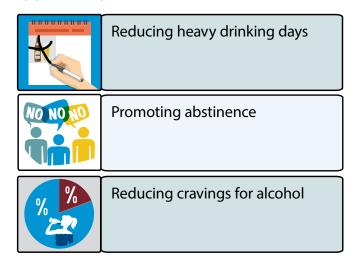


Topiramate

Topiramate is not FDA approved for AUD but is well-supported by evidence and may reduce heavy drinking and promote abstinence. One study found that for reducing heavy drinking days topiramate had a NNT = 5 and when adjusted for adverse events the NNT increased to 7.63

Topiramate is at least as effective as naltrexone and acamprosate, and evidence suggests that it may have a greater magnitude of effect. 56,64-68

Figure 11. Evidence Supports Topiramate Effectiveness at:



In patients with co-occurring Posttraumatic Stress Disorder (PTSD) and AUD, a small pilot trial found that topiramate:⁶⁹

- Reduced alcohol consumption, alcohol craving
- Reduced PTSD symptom severity—particularly hyperarousal symptoms

While traditionally the goal of treating AUD was abstinence, medications like naltrexone and topiramate that reduce the amount of alcohol consumed can lead to substantial reduction in alcohol-related problems.⁷⁰



First-line suggested pharmacotherapy for AUD

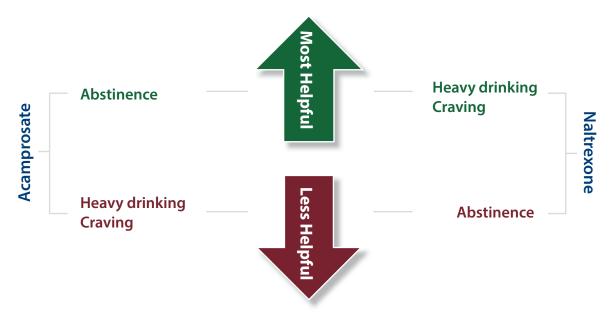
Acamprosate

An effective option for maintaining abstinence. 43,56,58,71-74

- Improved abstinence rates when used in combination with psychotherapy (NNT = 9)^{74,75}
- The risk of individuals returning to any drinking at 6 months is significantly lower than placebo (RR = 0.83, 95% CI = 0.78-0.89)⁵⁸

Naltrexone vs. Acamprosate

Figure 12. When are These Medications Most Helpful?⁵⁶



Naltrexone has a significant effect on the prevention of heavy drinking and to a lesser extent the maintenance of abstinence. Acamprosate supports abstinence; it does not influence alcohol consumption after the first drink.

Disulfiram

Disulfiram supports abstinence by creating an adverse physical reaction in which alcohol consumption is quickly followed by adverse effects (e.g., nausea, vomiting, headache, flushing). Disulfiram has been found to be most effective with monitored administration.^{76–79}

Disulfiram should only be used if: 43,80

- The patient is committed to a goal of complete abstinence from alcohol and is highly motivated and cognitively intact
- Veteran must be abstinent at least 24 to 48 hours before disulfiram is started



For motivated and informed patients, disulfiram can be an effective part of their recovery program, especially if there is monitored administration.^{43,76,81}

Second-line pharmacotherapy for AUD

Gabapentin

When first-line pharmacotherapy for AUD is not effective or is contraindicated then Gabapentin can be considered as second-line treatment option for patients with AUD.⁴³ It has been shown to:^{43,82-88}

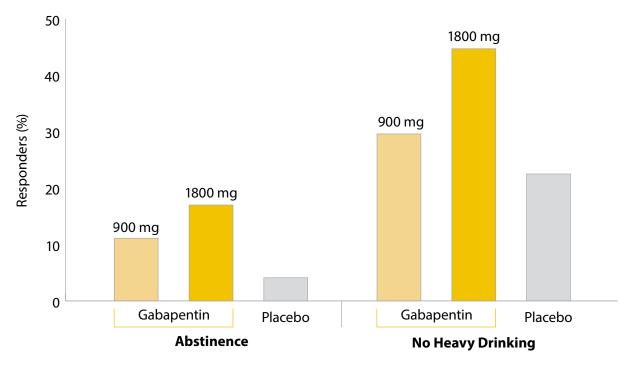
- Reduce heavy drinking
- Improve abstinence to a lesser extent
- Improve sleep
- Reduce acute or protracted alcohol withdrawal symptoms



Did you know?

Gabapentin may be more effective in patients with a history of or current alcohol withdrawal symptoms. A recent study found that individuals with high self-reported alcohol withdrawal symptoms had less relapse to heavy drinking (NNT = 3.1) and more total abstinence (NNT = 2.7) and for those who reported minimal withdrawal symptoms, gabapentin was no different than placebo.

Figure 13. Gabapentin in a Linear Dose Effect Improved Abstinence and Reduced Drinking⁸³



In this 12-week, double-blind trial (n = 150) significant linear dose effects were reported with abstinence rate, no heavy drinking, cravings, mood, and sleep. These effects were more pronounced in the gabapentin 1800 mg group (abstinence: NNT = 8; no heavy drinking: NNT = 5).

Table 4. AUD Pharmacotherapy Pearls and Clinical Precautions

Recommended First-line Treatment Options^{43,69,89,90}

Drug Name	Usual Dose	Benefits	Contraindications/ Precautions/ Limitations	Side Effects
Naltrexone hydrochloride	 50 mg orally once daily Starting at 25 mg daily may increase tolerability 	 Reducing heavy drinking Supports abstinence Abstinence ≥4 days prior to initiation may improve results 	 Cannot be used in patients on opioids Patients with physiologic opioid dependance should be opioid-free (including tramadol) a minimum of 7-10 days Do not use in severe hepatic impairment or acute hepatitis Should be stopped 48 to 72 hours prior to surgery 	 Abdominal cramps Abdominal pain Nausea Headache (Generally subsides with continued use)
Naltrexone Injectable suspension	380 mg intramuscular gluteal injection every 4 weeks or once a month	 Same as oral Injectable suspension 	 Same as oral Clinic administration by nurse CrCl < 50 mL/min use with caution, no dose adjustment recommended Kept refrigerated. Allow drug to reach room temperature (about 45 minutes) prior to preparation 	 Same as oral Injection site reactions

Despite early reports, research has not found increased hepatotoxicity associated with naltrexone when provided at usual doses. ⁹¹ CrCl = creatinine clearance; See <u>Clinical Pearls for Treatment of Alcohol Use Disorder</u> for specific monitoring recommendations.

Drug Name	Usual Dose	Benefits	Contraindications/ Precautions/ Limitations	Side Effects
Topiramate	 Initiate: 25 mg/day Max/target dose: 200 to 300 mg/day divided doses 5-6 week titration recommended Do not abruptly discontinue 	 Reduces heavy drinking Promotes abstinence Concomitantly treat seizure disorder or migraine prophylaxis Potential benefit in Veterans with AUD and co-occurring PTSD 	 Renal impairment and geriatric dosing: CrCl <70 mL/min, use 50% of the usual adult dosage Caution in patients with history of kidney stones Caution in use with metformin (increased metabolic acidosis risk) Use during pregnancy can cause cleft lip or palate May decrease the serum concentrations of contraceptives and reduce their effectiveness 	 Cognitive/memory impairment Paresthesia Weight loss Headache Fatigue Dizziness (Most side effects are dose dependent and may dissipate over time. If side effects don't improve after 4-6 weeks dose reduction is recommended to improve adherence.)90

Despite early reports, research has not found increased hepatotoxicity associated with naltrexone when provided at usual doses. ⁹¹ CrCl = creatinine clearance; See <u>Clinical Pearls for Treatment of Alcohol Use Disorder</u> for specific monitoring recommendations.

Suggested (Weaker Evidence) First-Line Treatment Options 43,89

Drug Name	Usual Dose	Benefits	Contraindications/ Precautions/ Limitations	Side Effects
Acamprosate	• 666 mg orally three times daily	 Increases abstinence Abstinence ≥4 days prior to initiation may improve results Can be used safely in liver disease 	 Higher pill burden, multiple daily dosing Renal impairment dosing: CrCl 30-50 mL/min: 333 mg orally 3 times daily CrCl< 30 mL/min: do not use; Contraindicated Reduce dose in patients with low body weight < 60kg 	 Diarrhea Nervousness Fatigue (Generally subsides with continued use)
Disulfiram	• 250 mg once daily (125-500 mg range; maximum daily dose 500 mg)	 Discourages drinking by causing unpleasant physiologic reaction May be more effective with monitored administration 	 Patient must be abstinent prior to therapy and committed to staying sober Reaction with alcohol can occur for up to 14 days after last dose Contraindicated in severe myocardial disease, psychosis, cognitive disorders, Severe hepatic dysfunction Products containing ethanol could cause disulfiram reaction (e.g., mouthwash) Metronidazole (risk of disulfiram-like reaction) 	 Somnolence Metallic taste Headache Major: Hepatotoxicity, peripheral neuropathy, psychosis, delirium If alcohol consumed: nausea, vomiting, sweating, headache, disorientation, dyspnea, hypotension, palpitations

CrCl = creatinine clearance; See <u>Clinical Pearls for Treatment of Alcohol Use Disorder</u> for specific monitoring recommendations.

2nd Line Treatment Option^{43,89}

Drug Name	Usual Dose	Benefits	Contraindications/ Precautions/ Limitations	Side Effects
Gabapentin	Initiated at 300 mg at bedtime and titrate to target dose of 1,800 mg per day in divided doses	 Reduces heavy drinking Improves abstinence Helps with insomnia, and acute withdrawal symptoms May be more effective in patients with withdrawal symptoms 	 Renal dosing if CrCL = 30 to 59 mL/min, 400 to 1400 mg/ day given in two divided doses; 15 to 29ml/min give 200 to 700 mg at bedtime; CrCl 15 mL/min give 100 to 300 mg/day given once daily; CrCL < 15 mL/min reduce daily dose in proportion to CrCL Misuse potential Use caution with CNS depressants 	 Somnolence/ fatigue Dizziness Ataxia Peripheral edema Weight gain

CrCl = creatinine clearance; See <u>Clinical Pearls for Treatment of Alcohol Use Disorder</u> for specific monitoring recommendations.

Combining Different Pharmacotherapies

There is little evidence to support combination pharmacotherapy. Short-term combinations may be used if patients experience a poor response to adequate trials of monotherapy combined with psychosocial interventions.^{49,92–96} In addition, medications like gabapentin may be used to manage residual withdrawal symptoms like insomnia.⁹²

Other Pharmacotherapy Options Still Under Investigation

Several trials have looked at other pharmacotherapy modalities for the treatment of AUD including: baclofen, ondansetron, and varenicline.

Consideration of these agents should only occur in patients for whom first-line and second-line pharmacotherapy is contraindicated or ineffective. These medications could be useful as adjunct agents in patients with comorbid AUD and disease states for which these agents are FDA-approved.

Figure 14. Baclofen^{43,97-109}

- Has been studied in patients with advanced liver disease and AUD; more patients able to maintain abstinence compared to placebo (71% vs. 29%) using low dose.
- Low dose (30–60 mg/day) may be more effective and tolerable than high-dose (up to 270 mg/day).
- One study found that 90 mg/day has greater efficacy in men with good tolerability, whereas women showed benefit on 30 mg/day and reported increased adverse effects at 90 mg/day.
- May be more effective in those with a higher daily alcohol intake.
- Further studies are needed to define future role.
- Baclofen may be a treatment option for patients with advanced liver disease who do not respond to or do not tolerate topiramate, acamprosate, or gabapentin.

May be effective at:

- Promoting abstinence
- Reducing heavy drinking days
- Reducing relapse rates
- Reducing cravings and anxiety

Significant risks:

- Withdrawal
- Sedation
- Misuse potential

Figure 15. Additional Medications Studied 99, 110-121

Ondansetron

May reduce heavy drinking and improve abstinence; however results vary

Preliminary evidence suggests better response with: early onset (<25 years old) AUD; specific polymorphism of the serotonin transporter and HTR3A/3B genes

Complex dosing (4 mcg/kg twice daily)

Varenicline

May reduce heavy drinking and cravings for alcohol; however results vary

May be more effective in Men, those who reduce tobacco smoking, and those with less-severe AUD at reducing heavy drinking days

Receiving any smoking cessation intervention (during treatment or later in recovery) is associated with 25% increased likelihood of long-term abstinence from alcohol and drugs

Psychosocial and Behavioral Interventions

For Veterans with alcohol use disorder it can be useful to offer, when available, one or more time limited psychosocial or behavioral interventions (Figure 16).^{43,46}

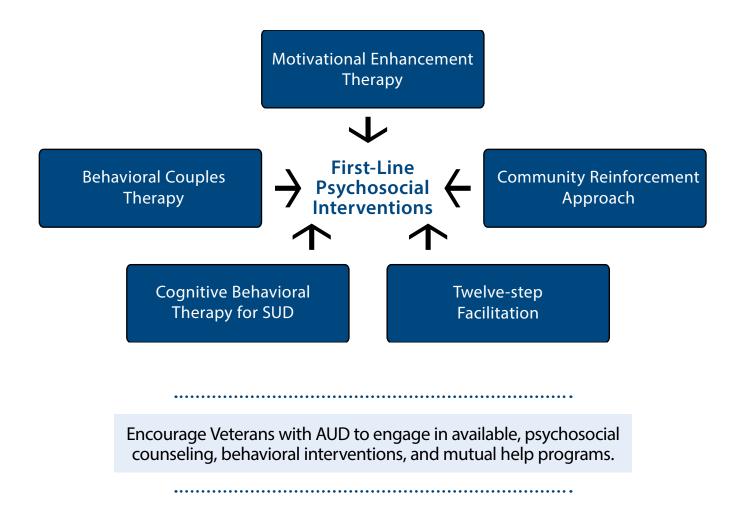
Psychosocial interventions:

- Improve outcomes of consumption, adherence and recovery
- When combined with pharmacotherapy may enhance treatment adherence and address problems (e.g., skills building) that medications won't address¹²²

In addition, encourage active involvement in group mutual help programs like Alcoholics Anonymous and SMART recovery.

Regular contact with the Veteran can positively influence treatment adherence and outcomes.

Figure 16. Effective Psychosocial Interventions⁴³



Assessing Treatment Effectiveness

Provide periodic monitoring of the Veteran's response to treatment. Ask about alcohol use and discuss Veteran's recovery goals. Consider using standardized and valid measurement instruments (e.g., Brief Addiction Monitor) and alcohol biomarkers (e.g., Carbohydrate-Deficient transferrin), whenever possible.

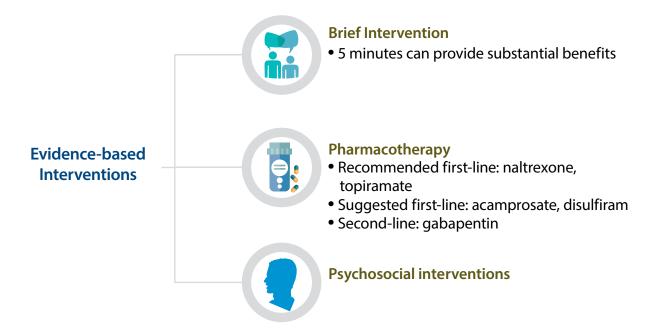
Monitor alcohol use and recovery goals to optimize treatment outcomes at periodic predetermined intervals.

Ask about:

Current alcohol use
Cravings
Medication side effects
Emerging symptoms
Alcohol biomarkers

Summary

There are a variety of effective evidence-based pharmacotherapy and psychosocial interventions available to treat AUD.



All VA clinicians have the opportunity to be advocates for our Veterans' recovery from unhealthy alcohol use. Identifying alcohol-related risk and encouraging Veterans to obtain treatment for AUD is the right investment of VA resources.

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Textual Equivalents for Graphic Diagrams

Figure 5: This figure presents a brief overview of recommended treatments by AUDIT-C score from low to high risk. Patients who practice abstinence or present a low-risk of drinking, AUDIT-C of 0 to 3, are recommended health promotion.

Patients with a moderate-risk of drinking, AUDIT-C of 4 to 5, are recommended brief intervention. Patients with an AUDIT-C Score greater than or equal to 4 are also at risk of decreased medication adherence.

Patients at high-risk of drinking, AUDIT-C of 6 to 7, are recommended to brief interventions with or without pharmacotherapy and psychosocial interventions.

Patients with an AUDIT-C Score greater than or equal to 6 are also at risk for increased GI conditions and poorer diabetes mellitus self-management.

Patients at severe-risk of drinking fall into two AUDIT C ranges: 8 to 9 and 10 to 12.

Patients with an AUDIT-C range of 8 to 9 are recommended pharmacotherapy with or without psychosocial interventions and specialty care management. Patients with an AUDIT-C Score greater than or equal to 8 are also at increased rates of hospitalizations, fractures, and prolonged postoperative hospital stays.

Lastly, patients in the highest AUDIT-C range of 10-12 are recommended specialty care management. This group is also at increased risk of trauma and mortality.

Figure 12: This graphic depicts the differences between acamprosate and naltrexone effectiveness for the treatment of alcohol use disorder. Up arrow shows that naltrexone has a significant effect on the prevention of heavy drinking and cravings and acamprosate is most helpful with abstinence. The down arrow shows that naltrexone is less helpful with maintenance of abstinence and acamprosate does not influence alcohol consumption after the first drink.

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This reference guide was created to be used as a tool for VA providers and is available to use from the Academic Detailing Service SharePoint.

These are general recommendations only; specific clinical decisions should be made by the treating provider based on an individual patient's clinical condition.

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