BENZODIAZEPINES FOR ANXIETY

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Learning Objectives

- 1. List common/serious adverse effects of benzodiazepines
- 2. Identify factors that influence whether a patient is or is not an appropriate candidate for short-term benzodiazepine use for anxiety.
- 3. Identify key educational points for a patient newly starting on benzodiazepine treatment for anxiety
- 4. List common/serious benzodiazepine withdrawal symptoms.
- 5. Develop a taper schedule for a patient who is discontinuing benzodiazepine treatment to minimize risk for withdrawal symptoms

Bridging Therapy- used in short-term to alleviate acute anxiety symptoms

Benzodiazepine- anxiety significantly impedes performance of ADL and no safety concerns

****Gabapentin, hydroxyzine, atypical antipsychotic**- alternatives to benzodiazepine therapy for acute treatment if safety concerns exist

Sleep aid- anxiety significantly impairing ability to sleep -trazodone, mirtazapine, TCA, hydroxyzine

**Not FDA-approved or well studied methods, but are sometimes used in practice due to lack of other options for acute anxiety treatment

Guidelines available from: American Psychiatric Association (APA), World Federation of Societies of Biological Psychiatry (WFSBP), and British Association for Psychopharmacology.

Benzodiazepines

- Act at GABA-A receptor complex in CNS
- Potentiate effects of GABA (main inhibitory neurotransmitter)
- Use: anxiety, insomnia, seizures, alcohol withdrawal, agitation, sedation
- Reduce emotional and somatic symptoms of anxiety
- Onset of action: several minutes to hours
- Duration of use: short-term, until maintenance treatments are expected to take effect

Benzodiazepines

Generic	Brand	Daily dose range (mg)	Elimination Half-life (hrs)	Active metabolite	Metabolic pathway
Lorazepam	Ativan	2 – 10	10 – 20	No	Conjugation
Clonazepam	Klonopin	0.5 – 4	20 – 50	No	Oxidation
Chlordiazepoxide	Librium	15 – 100	> 100	Yes	Oxidation
Oxazepam	Serax	30 – 120	5 – 10	No	Conjugation
Diazepam	Valium	4 – 40	> 100	Yes	Oxidation
Alprazolam	Xanax	0.75 – 4	12 – 15	Yes	Oxidation

**Conjugated agents preferred in patients on multiple meds, and patients with reduced hepatic function (elderly, alcohol use disorder, etc.)

Benzodiazepines

Adverse Effects

CNS Depression	Cognitive Impairment	Behavior and Mood
 Sedation Blurred Vision Ataxia Psychomotor impairment Disorientation Respiratory depression 	 Confusion Slowed mental processes Memory impairment Dementia 	 Paradoxical disinhibition Labile mood

Benzodiazepine Cautions

- Abuse potential (C-IV)
 - Enhance euphoric effects of opioids, synergistic with alcohol
 - Short-half life, high potency and lipophilic agents preferred
- Respiratory Depression
 - Increased risk of fatal overdose when co-ingested with other CNS depressants (alcohol, opioids, antihistamines, sedative hypnotics)
 - COPD and other lung conditions
- Elderly
 - Increased risk of falls and other accidents, drug interactions, cognitive dysfunction, dementia, paradoxical disinhibition, withdrawal effects
- Pregnancy category D (teratogenic effects)

Bridge Therapy with Benzodiazepine

Consider...

- Indication
 - Bridge therapy possibly indicated if high degree of functional impairment potentially detrimental to wait several weeks for maintenance therapy to become effective
 - Examples:
 - Frequent and severe panic attacks
 - Inability to leave home
 - Inability to attend or function at work or school
- Safety
 - Abuse potential
 - Mortality/morbidity

Benzodiazepine Taper

- Minimize withdrawal or rebound symptoms:
 - Anxiety, irritability, insomnia, tachycardia, increased BP, tremor, etc.
 - Seizures, delirium
- Slowly over period of weeks to months (depends on duration of use)
 - 10 25% reduction no quicker than every 2 weeks
 - May need to slow rate of taper once 50% of original dose is achieved
- If withdrawal symptoms, resume last tolerated dose and decrease rate of taper
- Consider switch to equivalent dose of agent with longer half-life to facilitate taper (clonazepam)

Example

- FR is a 54yo male whose physician wishes to taper him off of diazepam.
- Benzodiazepine regimen:
 - Diazepam 5mg po 4 times daily (~2.5 years)—was taking 5mg po TID for ~1 year prior to that
- What taper schedule would you recommend? For each step of the taper include dose/frequency and duration (how long to continue before next step down)

Example

Week 1: 5mg TID Week 3: 2.5mg QAM; 5mg QPM; 5mg Q bedtime Week 5: 2.5mg QAM; 2.5mg QPM; 5mg Q bedtime Week 7: 2.5mg TID Week 9: 2mg QAM; 2mg QPM; 2mg Q bedtime Week 11: 1mg QAM; 2mg QPM; 2mg Q bedtime Week 13: 1mg QAM; 1mg QPM; 2mg Q bedtime Week 15: 1mg TID Week 17: 1mg BID Week 19: 1mg daily Week 21: Stop



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