AGS BEERS CRITERIA FOR POTENTIALLY INAPPROPRIATE MEDICATION USE IN OLDER ADULTS

FROM THE AMERICAN GERIATRICS SOCIETY

This clinical tool, based on The AGS 2012 Updated Beers Criteria for Potentially Inappropriate Medication Use in Older Adults (AGS 2012 Beers Criteria), has been developed to assist healthcare providers in improving medication safety in older adults. Our purpose is to inform clinical decision-making concerning the prescr bing of medications for older adults in order to improve safety and quality of care.

Originally conceived of in 1991 by the late Mark Beers, MD, a geriatrician, the Beers Criteria catalogues medications that cause adverse drug events in older adults due to their pharmacologic properties and the physiologic changes of aging. In 2011, the AGS undertook an update of the criteria, assembling a team of experts and funding the development of the AGS 2012 Beers Criteria using an enhanced, evidence-based methodology. Each criterion is rated (quality of evidence and strength of evidence) using the American College of Physicians' Guideline Grading System, which is based on the GRADE scheme developed by Guyatt et al.

The full document together with accompanying resources can be viewed online at www.americangeriatrics.org.

INTENDED USE

The goal of this clinical tool is to improve care of older adults by reducing their exposure to Potentially Inappropriate Medications (PIMs).

- This should be viewed as a guide for identifying medications for which the risks of use in older adults outweigh the benefits.
- These criteria are not meant to be applied in a punitive manner.
- This list is not meant to supersede clinical judgment or an individual patient's values and needs. Prescribing and managing disease conditions should be individualized and involve shared decision-making.
- These criteria also underscore the importance of using a team approach to prescribing and the use of non-pharmacological approaches and of having economic and organizational incentives for this type of model.
- Implicit criteria such as the STOPP/START criteria and Medication Appropriateness Index should be used in a complementary manner with the 2012 AGS Beers Criteria to guide clinicians in making decisions about safe medication use in older adults.

The criteria are not applicable in all circumstances (eg, patient's receiving palliative and hospice care). If a clinician is not able to find an alternative and chooses to continue to use a drug on this list in an individual patient, designation of the medication as potentially inappropriate can serve as a reminder for close monitoring so that the potential for an adverse drug effect can be incorporated into the medical record and prevented or detected early.

First-generation antihistamines (as single agent or as part of combination products)	Avoid.
 Brompheniramine Carbinoxamine Chlorpheniramine Clemastine Cyproheptadine 	Highly anticholinergic; clearance reduced with advanced age, and tolerance develops when used as hypnotic; increased risk of confusion, dry mouth, constipation, and other anticholinergic effects/toxicity.
DexbrompheniramineDexchlorpheniramineDiphenhydramine (oral)	Use of diphenhydramine in special situations such as acute treatment of severe allergic reaction may be appropriate.
 Doxylamine Hydroxyzine Promethazine Triprolidine 	QE = High (Hydroxyzine and Promethazine), Moderate (All others); SR = Strong
Antiparkinson agents Benztropine (oral)	Avoid.
■ Trihexyphenidyl	Not recommended for prevention of extrapyramidal symptoms with antipsychotics; more effective agents available for treatment of Parkinson disease.
	QE = Moderate; SR = Strong

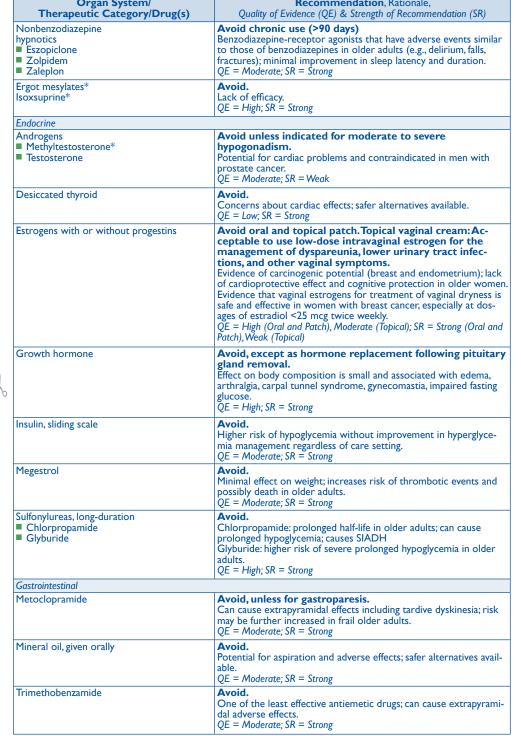
Table I (continued from page I)	
Antispasmodics Belladonna alkaloids Clidinium-chlordiazepoxide Dicyclomine Hyoscyamine Propantheline Scopolamine	Avoid except in short-term palliative care to decrease oral secretions. Highly anticholinergic, uncertain effectiveness. QE = Moderate; SR = Strong
Dipyridamole, oral short-acting* (does not apply to the extended-release combination with aspirin) Ticlopidine*	Avoid. May cause orthostatic hypotension; more effective alternatives available; IV form acceptable for use in cardiac stress testing. QE = Moderate; SR = Strong Avoid. Safer, effective alternatives available. QE = Moderate; SR = Strong
Nitrofurantoin	Avoid for long-term suppression; avoid in patients with CrCl <60 mL/min. Potential for pulmonary toxicity; safer alternatives available; lack of efficacy in patients with CrCl <60 mL/min due to inadequate drug concentration in the urine. QE = Moderate; SR = Strong
Alpha _i blockers Doxazosin Prazosin Terazosin	Avoid use as an antihypertensive. High risk of orthostatic hypotension; not recommended as routine treatment for hypertension; alternative agents have superior risk/benefit profile. QE = Moderate; SR = Strong
Alpha agonists Clonidine Guanabenz* Guanfacine* Methyldopa* Reserpine (>0.1 mg/day)*	Avoid clonidine as a first-line antihypertensive. Avoid others as listed. High risk of adverse CNS effects; may cause bradycardia and orthostatic hypotension; not recommended as routine treatment for hypertension. QE = Low; SR = Strong
Antiarrhythmic drugs (Class Ia, Ic, III) Amiodarone Dofetilide Dronedarone Flecainide Ibutilide Procainamide Propafenone Quinidine Sotalol	Avoid antiarrhythmic drugs as first-line treatment of atria fibrillation. Data suggest that rate control yields better balance of benefits and harms than rhythm control for most older adults. Amiodarone is associated with multiple toxicities, including thyroid disease, pulmonary disorders, and QT interval prolongation. QE = High; SR = Strong
Disopyramide*	Avoid. Disopyramide is a potent negative inotrope and therefore may induce heart failure in older adults; strongly anticholinergic; other antiarrhythmic drugs preferred. QE = Low; SR = Strong
Dronedarone	Avoid in patients with permanent atrial fibrillation or heart failure. Worse outcomes have been reported in patients taking drone-darone who have permanent atrial fibrillation or heart failure. In general, rate control is preferred over rhythm control for atrial fibrillation. QE = Moderate; SR = Strong
Digoxin >0.125 mg/day	Avoid. In heart failure, higher dosages associated with no additional benefit and may increase risk of toxicity; decreased renal clearance may increase risk of toxicity. QE = Moderate; SR = Strong

PAGE I Table 1 (continued on page 2) PAGE 2 Table 1 (continued on page 3)

Organ System/	Recommendation, Rationale,
Therapeutic Category/Drug(s)	Quality of Evidence (QE) & Strength of Recommendation (SR)
Nifedipine, immediate release*	Avoid.
	Potential for hypotension; risk of precipitating myocardial ischemia. QE = High; SR = Strong
Spironolactone >25 mg/day	Avoid in patients with heart failure or with a CrCl <30 mL/min.
	In heart failure, the risk of hyperkalemia is higher in older adults if taking >25 mg/day. QE = Moderate; SR = Strong
Central Nervous System	
Tertiary TCAs, alone or in combination:	Avoid.
 Amitriptyline Chlordiazepoxide- amitriptyline Clomipramine Doxepin >6 mg/day 	Highly anticholinergic, sedating, and cause orthostatic hypotension; the safety profile of low-dose doxepin (≤6 mg/day) is comparable to that of placebo.
Imipramine Perphenazine-amitriptyline Trimipramine	QE = High; SR = Strong
Antipsychotics, first- (conventional) and second- (atypical) generation (see online for full list)	Avoid use for behavioral problems of dementia unless non-pharmacologic options have failed and patient is threat to self or others.
	Increased risk of cerebrovascular accident (stroke) and mortality in persons with dementia. QE = Moderate; SR = Strong
Thioridazine	Avoid.
Mesoridazine	Highly anticholinergic and greater risk of QT-interval prolongation. QE = Moderate; SR = Strong
Barbiturates	Avoid.
Amobarbital* Butabarbital* Butalbital	High rate of physical dependence; tolerance to sleep benefits; greater risk of overdose at low dosages.
 Mephobarbital* Pentobarbital Phenobarbital Secobarbital 	QE = High; SR = Strong
Benzodiazepines Short- and intermediate-acting:	Avoid benzodiazepines (any type) for treatment of insomnia, agitation, or delirium.
 Alprazolam Estazolam Lorazepam Oxazepam Temazepam Triazolam 	Older adults have increased sensitivity to benzodiazepines and decreased metabolism of long-acting agents. In general, all benzodiazepines increase risk of cognitive impairment, delirium, falls, fractures, and motor vehicle accidents in older adults.
Long-acting: Chlorazepate Chlordiazepoxide Chlordiazepoxide-amitriptyline Clidinium-chlordiazepoxide	May be appropriate for seizure disorders, rapid eye movement sleep disorders, benzodiazepine withdrawal, ethanol withdrawal, severe generalized anxiety disorder, periprocedural anesthesia, end-of-life care.
ClonazepamDiazepamFlurazepamQuazepam	QE = High; SR = Strong
Chloral hydrate*	Avoid. Tolerance occurs within 10 days and risk outweighs the benefits in light of overdose with doses only 3 times the recommended dose. QE = Low; SR = Strong
Meprobamate	Avoid.

	tentially Inappropriate Medication Use in Older Adults
Organ System/ Therapeutic Category/Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Nonbenzodiazepine hypnotics Eszopiclone Zolpidem Zaleplon	Avoid chronic use (>90 days) Benzodiazepine-receptor agonists that have adverse events similar to those of benzodiazepines in older adults (e.g., delirium, falls, fractures); minimal improvement in sleep latency and duration. QE = Moderate; SR = Strong
Ergot mesylates* Isoxsuprine*	Avoid. Lack of efficacy. QE = High; SR = Strong
Endocrine	
Androgens ■ Methyltestosterone* ■ Testosterone	Avoid unless indicated for moderate to severe hypogonadism. Potential for cardiac problems and contraindicated in men with prostate cancer. QE = Moderate; SR = Weak
Desiccated thyroid	Avoid. Concerns about cardiac effects; safer alternatives available. QE = Low; SR = Strong
Estrogens with or without progestins	Avoid oral and topical patch. Topical vaginal cream: Acceptable to use low-dose intravaginal estrogen for the management of dyspareunia, lower urinary tract infections, and other vaginal symptoms. Evidence of carcinogenic potential (breast and endometrium); lack of cardioprotective effect and cognitive protection in older womer Evidence that vaginal estrogens for treatment of vaginal dryness is safe and effective in women with breast cancer, especially at dosages of estradiol <25 mcg twice weekly. QE = High (Oral and Patch), Moderate (Topical); SR = Strong (Oral and Patch), Weak (Topical)
Growth hormone	Avoid, except as hormone replacement following pituitary gland removal. Effect on body composition is small and associated with edema, arthralgia, carpal tunnel syndrome, gynecomastia, impaired fasting glucose. QE = High; SR = Strong
Insulin, sliding scale	Avoid. Higher risk of hypoglycemia without improvement in hyperglycemia management regardless of care setting. QE = Moderate; SR = Strong
Megestrol	Avoid. Minimal effect on weight; increases risk of thrombotic events and possibly death in older adults. QE = Moderate; SR = Strong
Sulfonylureas, long-duration Chlorpropamide Glyburide	Avoid. Chlorpropamide: prolonged half-life in older adults; can cause prolonged hypoglycemia; causes SIADH Glyburide: higher risk of severe prolonged hypoglycemia in older adults. QE = High; SR = Strong
Gastrointestinal	·
Metoclopramide	Avoid, unless for gastroparesis. Can cause extrapyramidal effects including tardive dyskinesia; risk may be further increased in frail older adults. QE = Moderate; SR = Strong
Mineral oil, given orally	Avoid. Potential for aspiration and adverse effects; safer alternatives available. QE = Moderate; SR = Strong
Trimethobenzamide	Avoid. One of the least effective antiemetic drugs; can cause extrapyramidal adverse effects. QE = Moderate; SR = Strong





	otentially Inappropriate Medication Use in Older Adults
Organ System/ Therapeutic Category/Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Pain Medications	
Meperidine	Avoid. Not an effective oral analgesic in dosages commonly used; may cause neurotoxicity; safer alternatives available. QE = High; SR = Strong
Non-COX-selective NSAIDs, oral Aspirin >325 mg/day Diclofenac Diflunisal Etodolac Fenoprofen Ibuprofen Ketoprofen Meclofenamate Mefenamic acid Meloxicam Nabumetone Naproxen Oxaprozin Piroxicam Sulindac Tolmetin	Avoid chronic use unless other alternatives are not effective and patient can take gastroprotective agent (proton-pump inhibitor or misoprostol). Increases risk of GI bleeding/peptic ulcer disease in high-risk groups, including those ≥75 years old or taking oral or parenteral corticosteroids, anticoagulants, or antiplatelet agents. Use of proton pump inhibitor or misoprostol reduces but does not eliminate risk. Upper GI ulcers, gross bleeding, or perforation caused by NSAIDs occur in approximately 1% of patients treated for 3-6 months, and in about 2%-4% of patients treated for 1 year. These trends continue with longer duration of use. QE = Moderate; SR = Strong
Indomethacin Ketorolac, includes parenteral	Avoid. Increases risk of GI bleeding/peptic ulcer disease in high-risk groups (See Non-COX selective NSAIDs) Of all the NSAIDs, indomethacin has most adverse effects. QE = Moderate (Indomethacin), High (Ketorolac); SR = Strong
Pentazocine*	Avoid. Opioid analgesic that causes CNS adverse effects, including confusion and hallucinations, more commonly than other narcotic drugs; is also a mixed agonist and antagonist; safer alternatives available. QE = Low; SR = Strong
Skeletal muscle relaxants Carisoprodol Chlorzoxazone Cyclobenzaprine Metaxalone Methocarbamol Orphenadrine	Avoid. Most muscle relaxants poorly tolerated by older adults, because of anticholinergic adverse effects, sedation, increased risk of fractures; effectiveness at dosages tolerated by older adults is questionable. QE = Moderate; SR = Strong
	ons: ACEI, angiotensin converting-enzyme inh bitors; ARB, angiotensin

*Infrequently used drugs. Table I Abbreviations: ACEI, angiotensin converting-enzyme inh bitors; ARB, angiotensin receptor blockers; CNS, central nervous system; COX, cyclooxygenase; CrCI, creatinine clearance; GI, gastrointestinal; NSAIDs, nonsteroidal anti-inflammatory drugs; SIADH, syndrome of inappropriate antidiuretic hormone secretion; SR, Strength of Recommendation; TCAs, tricyclic antidepressants; QE, Quality of Evidence

TABLE 2: 2012 AGS Beers Criteria for Potentially Inappropriate Medication Use in Older Adults Due to Drug-

Disease or D	rug-Syndrome Interactions I hat May Exaceri	date the Disease or Syndrome
Disease or Syndrome	Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Cardiovascular		
Heart failure	NSAIDs and COX-2 inhibitors	Avoid.
	Nondihydropyridine CCBs (avoid only for systolic heart failure) Diltiazem	Potential to promote fluid retention and/or exacerbate heart failure.
	■ Verapamil	QE = Moderate (NSAIDs, CCBs, Dronedarone), High (Thiazolidinediones (glitazones)), Low (Cilostazol); SR = Strong
	Pioglitazone, rosiglitazone	(5 /// (7 //
	Cilostazol Dronedarone	

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TABLE 2: 2012 AGS Beers Criteria for Potentially Inappropriate Medication Use in Older Adults Due to Drug-

	rug-Syndrome Interactions That May Exacerl	ropriate Medication Use in Older Adults Due to Drug- bate the Disease or Syndrome
Disease or Syndrome	Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Syncope	Acetylcholinesterase inhibitors (AChEls) Peripheral alpha blockers Doxazosin Prazosin Tertazosin Tertiary TCAs Chlorpromazine, thioridazine, and olan-	Avoid. Increases risk of orthostatic hypotension or bradycardia. QE = High (Alpha blockers), Moderate (AChEls, TCAs and antipsychotics); SR = Strong (AChEls and TCAs), Weak (Alpha blockers and antipsychotics)
Central Nervou	zapine	
Chronic seizures or epilepsy	Bupropion Chlorpromazine Clozapine Maprotiline Olanzapine Thioridazine	Avoid. Lowers seizure threshold; may be acceptable in patients with well-controlled seizures in whom alternative agents have not been effective.
Delirium	Thiothixene Tramadol All TCAs Anticholinergics (see online for full list) Benzodiazepines Chlorpromazine Corticosteroids H ₂ -receptor antagonist Meperidine Sedative hypnotics	QE = Moderate; SR = Strong Avoid. Avoid in older adults with or at high risk of delirium because of inducing or worsening delirium in older adults; if discontinuing drugs used chronically, taper to avoid withdrawal symptoms. QE = Moderate; SR = Strong
Dementia & cognitive impairment	Thioridazine Anticholinergics (see online for full list) Benzodiazepines H ₂ -receptor antagonists Zolpidem Antipsychotics, chronic and as-needed use	Avoid. Avoid due to adverse CNS effects. Avoid antipsychotics for behavioral problems of dementia unless non-pharmacologic options have failed and patient is a threat to themselves or others. Antipsychotics are associated with an increased risk of cerebrovascular accident (stroke) and mortality in persons with dementia. QE = High; SR = Strong
History of falls or fractures	Anticonvulsants Antipsychotics Benzodiazepines Nonbenzodiazepine hypnotics Eszopiclone Zaleplon Zolpidem TCAs/SSRIs	Avoid unless safer alternatives are not available; avoid anticonvulsants except for seizure. Ability to produce ataxia, impaired psychomotor function, syncope, and additional falls; shorter-acting benzodiazepines are not safer than long-acting ones. QE = High; SR = Strong
Insomnia	Oral decongestants Pseudoephedrine Phenylephrine Stimulants Amphetamine Methylphenidate Pemoline Theobromines Theophylline Caffeine	Avoid. CNS stimulant effects. QE = Moderate; SR = Strong
Parkinson's disease	All antipsychotics (see online publication for full list, except for quetiapine and clozapine) Antiemetics Metoclopramide Prochlorperazine Promethazine	Avoid. Dopamine receptor antagonists with potential to worsen parkinsonian symptoms. Quetiapine and clozapine appear to be less likely to precipitate worsening of Parkinson disease. QE = Moderate; SR = Strong

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6) TABLE 2: 2012 AGS Beers Criteria for Potentially Inappropriate Medication Use in Older Adults Due to Drug-Disease or Drug-Syndrome Interactions That May Exacerbate the Disease or Syndrome Disease or **Recommendation**, Rationale, Quality of Evidence Drug(s) **Syndrome** (QE) & Strength of Recommendation (SR) **Gastrointestinal** Chronic Oral antimuscarinics for urinary inconti-Avoid unless no other alternatives. constipation nence Darifenacin Can worsen constipation; agents for urinary incon-■ Fesoterodine tinence: antimuscarinics overall differ in incidence of Oxybutynin (oral) constipation; response variable; consider alternative ■ Solifenacin agent if constipation develops. ■ Tolterodine ■ Trospium QE = High (For Urinary Incontinence), Moderate/Low (All Others): SR = Strong Nond hydropyridine CCB Diltiazem ■ Verapamil First-generation ant histamines as single agent or part of combination products Brompheniramine (various) Carbinoxamine Chlorpheniramine Clemastine (various) Cyproheptadine Dexbrompheniramine Dexchlorpheniramine (various) Diphenhydramine Doxylamine Hydroxyzine ■ Promethazine Triprolidine Anticholinergics/antispasmodics (see online for full list of drugs with strong anticholinergic properties)
Antipsychotics ■ Belladonna alkaloids Clidinium-chlordiazepoxide Dicyclomine Hyoscyamine Propantheline Scopolamine ■ Tertiary TCAs (amitriptyline, clomipramine, doxepin, imipramine, and trimipramine) History of Aspirin (>325 mg/day) Avoid unless other alternatives are not efgastric or Non-COX-2 selective NSAIDs fective and patient can take gastroprotective duodenal agent (proton-pump inhibitor or misoprostol). ulcers May exacerbate existing ulcers or cause new/additional ulcers. QE = Moderate; SR = StrongKidney/Urinary Tract Chronic kid-**NSAIDs** Avoid. ney disease stages IV May increase risk of kidney injury. and V Triamterene (alone or in combination) May increase risk of acute kidney injury. QE = Moderate (NSAIDs), Low (Triamterene); SR = Strong (NSAIDs), Weak (Triamterene) Urinary Estrogen oral and transdermal (excludes Avoid in women. incontinence intravaginal estrogen) (all types) in Aggravation of incontinence. women QE = High; SR = Strong

Table 2 (continued from page 7)

TABLE 2: 2012 AGS Beers Criteria for Potentially Inappropriate Medication Use in Older Adults Due to Drug-Disease or Drug-Syndrome Interactions That May Exacerbate the Disease or Syndrome

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Disease or Syndrome	Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Lower urinary tract symptoms, benign prostatic hyperplasia	Inhaled anticholinergic agents Strongly anticholinergic drugs, except antimuscarinics for urinary incontinence (see Table 9 for complete list).	Avoid in men. May decrease urinary flow and cause urinary retention. QE = Moderate; SR = Strong (Inhaled agents), Weak (All others)
Stress or mixed urinary in- continence	Alpha-blockers Doxazosin Prazosin Terazosin	Avoid in women. Aggravation of incontinence. QE = Moderate; SR = Strong

Table 2 Abbreviations: CCBs, calcium channel blockers; AChEls, acetylcholinesterase inh bitors; CNS, central nervous system; COX, cyclooxygenase; NSAIDs, nonsteroidal anti-inflammatory drugs; SR, Strength of Recommendation; SSRIs, selective serotonin reuptake inhibitors; TCAs, tricyclic antidepressants; QE, Quality of Evidence

TABLE 3: 2012 AGS Beers Criteria for Potentially Inappropriate Medications to Be Used with Caution in

Older Adults	
Drug(s)	Recommendation, Rationale, Quality of Evidence (QE) & Strength of Recommendation (SR)
Aspirin for primary prevention of cardiac events	Use with caution in adults \geq 80 years old. Lack of evidence of benefit versus risk in individuals \geq 80 years old. $QE = Low$; $SR = Weak$
Dabigatran	Use with caution in adults ≥75 years old or if CrCl <30 mL/min. Increased risk of bleeding compared with warfarin in adults ≥75 years old; lack of evidence for efficacy and safety in patients with CrCl <30 mL/min QE = Moderate; SR = Weak
Prasugrel	Use with caution in adults ≥75 years old. Greater risk of bleeding in older adults; risk may be offset by benefit in highestrisk older patients (eg, those with prior myocardial infarction or diabetes). QE = Moderate; SR = Weak
Antipsychotics Carbamazepine Carboplatin Cisplatin Mirtazapine SNRIs SSRIs TCAs Vincristine	Use with caution. May exacerbate or cause SIADH or hyponatremia; need to monitor sodium level closely when starting or changing dosages in older adults due to increased risk. QE = Moderate; SR = Strong
Vasodilators	Use with caution. May exacerbate episodes of syncope in individuals with history of syncope. QE = Moderate; SR = Weak
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Table 3 Abbreviations: CrCl, creatinine clearance; SIADH, syndrome of inappropriate antidiuretic hormone secretion; SSRIs, selective serotonin reuptake inhibitors; SNRIs, serotonin-norepinephrine reuptake inhibitors; SR, Strength of Recommendation; TCAs, tricyclic antidepressants; QE, Quality of Evidence

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