Text brittanygoldstein340 to 37607



SELECTING AN ANTIDEPRESSANT: MANY TO CHOOSE, LITTLE TO LOSE

BRITTANY GOLDSTEIN, MD

UW MEDICINE

DEPARTMENT OF PSYCHIATRY AND BEHAVIORAL SCIENCES
CONSULTATION-LIAISON PSYCHIATRY FELLOW







SPEAKER DISCLOSURES

Brittany Goldstein, MD has no conflicts of interest to disclose.



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Planner disclosures

The following series planners have no relevant conflicts of interest to disclose; other disclosures have been mitigated.

Mark Duncan MD Rick Ries MD Kari Stephens PhD Barb McCann PhD Anna Ratzliff MD PhD Betsy Payn MA PMP Esther Solano Cara Towle MSN RN



OBJECTIVES

- 1. Review the diagnostic criteria for major depressive disorder
- 2. Provide guiding principles for initiating antidepressants for depression
- 3. Review differences between antidepressants
- 4. Practice selecting antidepressant for example cases

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MAJOR DEPRESSIVE DISORDER

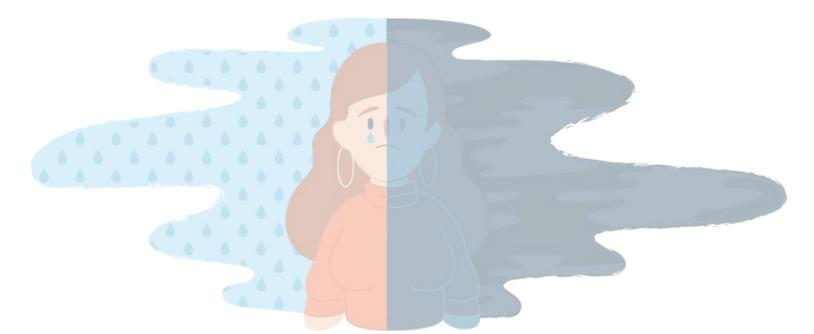
- Change in mood (depressed or anhedonic) ≥ 2 weeks
- 5 symptoms total
 - Depressed mood
 - Anhedonia
 - Sleep change
 - Psychomotor change
 - Appetite/weight change
 - Energy low
 - Guilt, worthlessness
 - Poor concentration
 - Suicidal ideation
- Most of the day for most days
- Change from baseline or chronic symptoms





ANTIDEPRESSANTS FOR DEPRESSION

- First-line
 - Mild-moderate: psychotherapy +/- medication
 - Severe: medication +/- therapy
 - Severest (psychosis, catatonia): medication + consider ECT





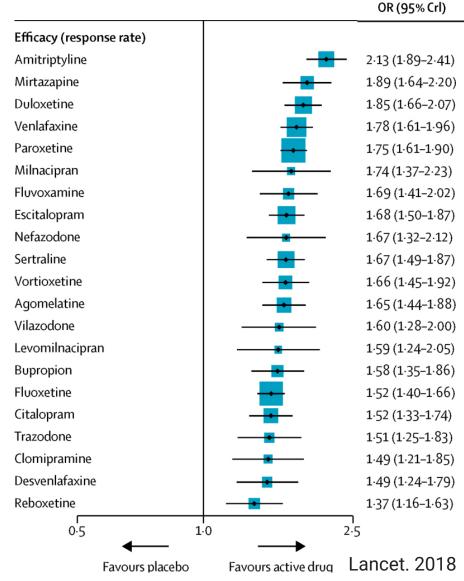
ANTIDEPRESSANTS FOR DEPRESSION

Set expectations



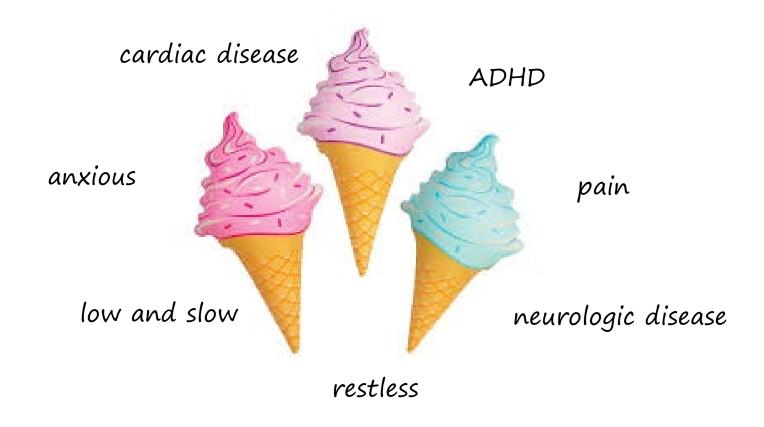


Psychother Psychosom. 2009



ANTIDEPRESSANTS FOR DEPRESSION

Depression flavor and comorbidities





PATIENT WITH DEPRESSION



- Standard depression
- With or without anxiety.
- Treatment naïve.



SELECTIVE SEROTONIN REUPTAKE INHIBITORS

- Multiple indications
 - On-label: depressive disorders, anxiety disorders, trauma disorders
 - Off-label: irritability/mild agitation, dysthymia, premature ejaculation
- Safe
- Sexual side effects
- Discontinuation syndrome
- Serotonin syndrome in overdose



SSRI

SHT2C NET SERT	Fluoxetine	Most activating Longest half-life DDI	Pt who is needs a pep in their step, poor medication adherence, otherwise healthy (low med burden) Pt who is not highly anxious or restless, polypharmacy
	Sertraline	Most GI SE Wide therapeutic range Start low, go slow Rare DDI	Pt who is comorbid cardiovascular dz, polypharmacy Pt who is not comorbid GI
S-citalopram SERT	Escitalopram	Clean Easy titration Rare DDI	Pt who is sensitive to side effects, wants a quick titration, polypharmacy
M1 NET SERT	Citalopram	QTc risk	Pt who is not cardiac comorbidity, other QTc prolonging medications
	Paroxetine	Messy→ most side effects Shortest half life DDI	Pt who is anxious, not sleeping, not eating, otherwise healthy (low med burden), good adherence Pt who is not sensitive to SE, polypharmacy

PATIENT WITH DEPRESSION



- Comorbid pain syndrome
- Comorbid ADHD

SEROTONIN AND NOREPINEPHRINE REUPTAKE INHIBITORS

- Multiple indications
 - On-label: depressive disorders, anxiety disorders, pain syndromes
 - Off label: pain syndromes, ADHD
- Safe
- Sexual side effects
- Discontinuation syndrome
- Serotonin syndrome in overdose



SNRI

duloxetine			
SERT	Duloxetine	FDA indication for pain	Pt who has comorbid pain syndrome, stress urinary incontinence
venlafaxine	Venlafaxine	SE: 个BP Short half-life	Pt who has vasomotor symptoms of menopause, good adherence Remember to monitor BP at initiation and increase, extended release formulation
desvenlafaxine	Desvenlafaxine	SE: ↑BP	Pt who has 2D6 inhibitor medications Remember to monitor BP at initiation and increase
S-milnacipran (levo) NET	Levomilnacipran	Greatest NE:5HT Most activating SE: 个BP	Pt who has comorbid pain or ADHD Pt who is not anxious Remember to monitor BP at initiation and increase

PATIENT WITH DEPRESSION

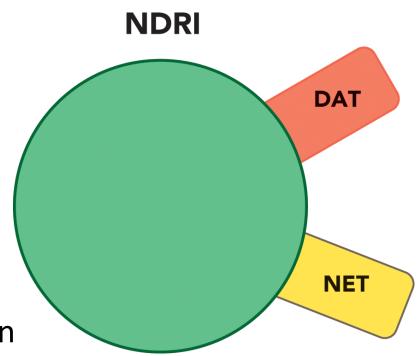


- Fatigued.
- Sleeping and eating a lot.
- Not anxious or restless.



BUPROPION

- Activating
- Monotherapy or augmentation
- Multiple indications
 - On-label: depressive disorders, smoking cessation
 - Off label: weight loss, ADHD, fatigue, SSRI-induced sexual dysfunction
- Extended release formulation
- Rare discontinuation syndrome
- Lowers seizure threshold in high risk patients
- Overdose: seizure, elevated BP and HR, arrhythmia





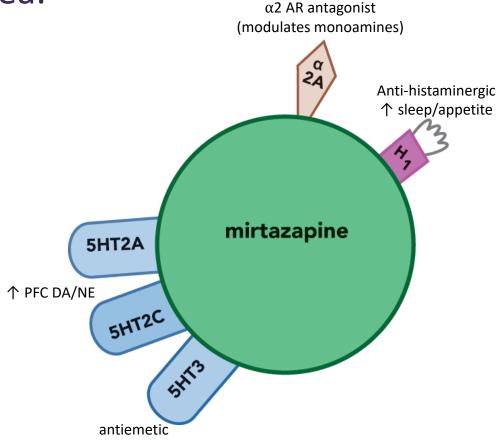
PATIENT WITH DEPRESSION



- Not sleeping
- Not eating

MIRTAZAPINE

- Increases sleep, appetite. Improves nausea.
- Dose dependent effects
 - 7.5mg-15mg = sleep and eat receptors
 - >15mg = mood/anxiety receptors
- No sexual side effects.
 Might improve sexual side effects.
- Monotherapy or augmentation
- Safe
- Discontinuation syndrome





PATIENT WITH DEPRESSION

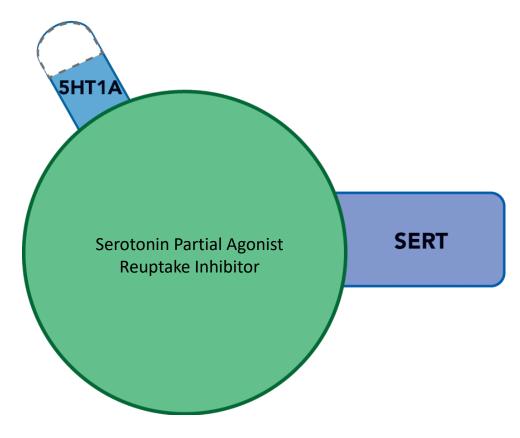


- Tried other things
- Other medications caused side effects
- Other medications had insufficient effectiveness



VILAZODONE

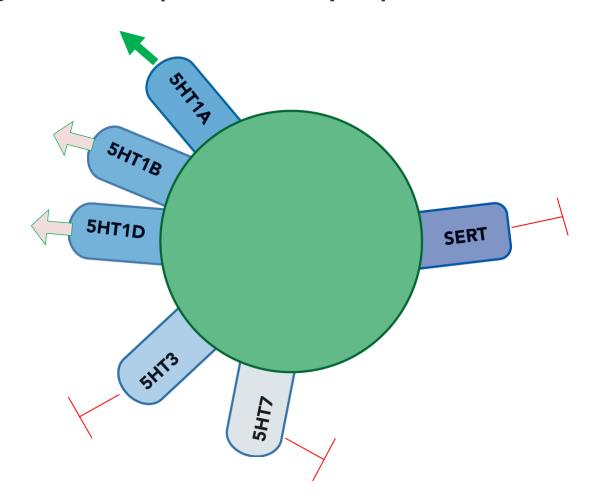
- Less sexual side effects
- More GI side effects





VORTIOXETINE

• Improves cognitive depressive symptoms





AUGMENTATION

Partial antidepressant effects with persistent	Consider adding
Hypersomnia, anergia, poor concentration, sexual side effects	bupropion
Insomnia, restlessness, low appetite, sexual side effects	mirtazapine
Anxiety, sexual side effects	buspirone
Insomnia	trazodone
Depression, anxiety	low dose second generation (atypical) antipsychotic
Depression, SI	lithium
Depression, hypersomnia, anergia, hyperphagia	T3 or levothyroxine



PATIENT WITH DEPRESSION

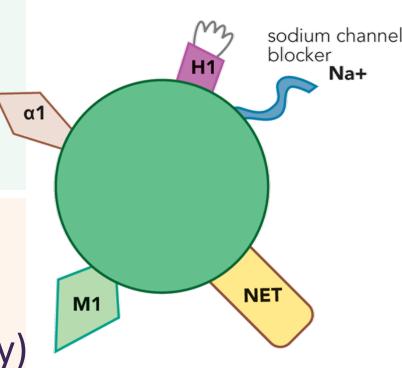


- Patient with a history of recurrent major depressive disorder complains of a recurrent depressive episode.
 Patient has been on numerous antidepressants from numerous classes. None have been effective, even in combination with each other. Their current depressive episode is significantly impairing their daily function though not imposing an imminent safety risk.
- Treatment resistant depression



TRICYCLIC ANTIDEPRESSANTS

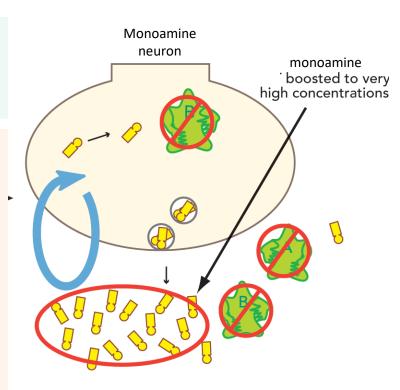
- Treatment resistant depression
- Multiple indications:
 - High doses: mood & anxiety disorders
 - Low doses: sleep, pain, functional GI
- Messy → side effects
- Discontinuation syndrome
- Lethal overdose (cardiotoxicity, neurotoxicity)





MONOAMINE OXIDASE INHIBITORS

- Treatment resistant depression
- MDD with "atypical features"
- High risk of serotonin syndrome and drug-drug interactions
- Dietary restrictions (low tyramine), risk of hypertensive crisis
- Messy → side effects
- Discontinuation syndrome, cannot cross-titrate
- Lethal overdose





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Practice

SELECT THAT ANTIDEPRESSANT



Patient with MDD and on multiple cardiac medications due to comorbid CAD s/p CABG, CHF, HTN, and HLD.

Which antidepressant would you choose first:

- A. Citalopram
- B. Amitriptyline
- C. Sertraline
- D. Venlafaxine



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Which antidepressant would you choose first:

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Patient with MDD including passive SI and comorbid ADHD who is also on cyclobenzaprine and ibuprofen for chronic myofascial pain.

Which antidepressant class would you choose first:

- A. SSRI
- B. SNRI
- C. TCA
- D. Alpha-2 adrenergic antagonist



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Patient with MDD already on the maximum dose of sertraline who had partial relief though still complains of excessive fatigue and has noticed the onset of anorgasmia since starting an SSRI.

What would you do next:

- A. Switch to paroxetine
- B. Increase sertraline dose
- C. Add quetiapine to sertraline
- D. Add bupropion to sertraline



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Patient experiencing a moderate major depressive episode and moderate-severe generalized anxiety disorder with panic attacks.

Which medication would you start:

- A. Escitalopram
- B. Buspirone
- C. Bupropion
- D. Selegiline



Patient experiencing a moderate major depressive episode and moderate-severe generalized anxiety disorder with panic attacks.

Which medication would you start:

- A. Escitalopram
- B. Buspirone
- C. Bupropion
- D. Selegiline



Patient with MDD who has trouble falling asleep and has lost 6 pounds in the past 1 month.

Which antidepressant would you choose:

- A. Venlafaxine
- B. Mirtazapine
- C. Bupropion
- D. Fluoxetine



Patient with MDD who has trouble falling asleep and has lost 6 pounds in the past 1 month.

Which antidepressant would you choose:

- A. Venlafaxine
- **B.** Mirtazapine
- C. Bupropion
- D. Fluoxetine



REFERENCES

- Stahl, S. M. (2000). Essential psychopharmacology: neuroscientific basis and practical application (2nd ed.). Cambridge University Press.
- Schatzberg, A. F., & DeBattista, C. (2019). *Schatzberg's Manual of Clinical Psychopharmacology* (9th ed.). American Psychiatric Publishing.
- Taylor, D., Barnes, T. R. E., & Young, A. (2021). The Maudsley prescribing guidelines in psychiatry (14th edition.). Wiley Blackwell.
- Rothmore J. Antidepressant-induced sexual dysfunction. Med J Aust. 2020 Apr;212(7):329-334. doi: 10.5694/mja2.50522. Epub 2020 Mar 15. PMID: 32172535.
- Rutherford BR, Sneed JR, Roose SP. Does study design influence outcome? The effects of placebo control and treatment duration in antidepressant trials. Psychother Psychosom. 2009;78(3):172-81. doi: 10.1159/000209348. Epub 2009 Mar 24. PMID: 19321970; PMCID: PMC3785090.
- Cipriani A, et al. Comparative efficacy and acceptability of 21 antidepressant drugs for the acute treatment of adults with major depressive disorder: a systematic review and network meta-analysis. Lancet. 2018 Apr 7;391(10128):1357-1366. doi: 10.1016/S0140-6736(17)32802-7. Epub 2018 Feb 21. PMID: 29477251; PMCID: PMC5889788.
- Gutlapalli SD, et al. The Risk of Fatal Arrhythmias Associated With Sertraline in Patients With Post-myocardial Infarction Depression. Cureus. 2022 Sep 8;14(9):e28946. doi: 10.7759/cureus.28946. PMID: 36237772; PMCID: PMC9547663.