Psychopharmacology for Therapists & Psychologists

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Outline

- Disclaimers
- Brief overview of brain anatomy and function
- Review main types of psychiatric disorders, their presentations & symptoms
- Review psychotropic medication classes and therapeutic effects
- Describe potential side effects, averse effects signs and symptoms, do not miss signs
- Novel Treatments
- Medication Assisted Treatment (MAT)

Collaborative Effort

- Time constraints during visits, especially insurance driven
- Appointment availability
- May not see prescriber as often as for therapy
- Many times embarrassed, have better rapport with therapist/psychologist
- Help client advocate for themselves
- Motivational interviewing, are you better with no meds?
- Progress farther in therapy

Some Vocabulary

- Psychopharmacology- the study of the effects of drugs on affect, behavior and cognition
- Agonist- drug that binds to and activates a receptor
- Antagonist- drug that binds to but does not activate instead blocks the receptor
- Efficacy- maximal therapeutic effect the drug can achieve
- Half-life- time necessary for half the drug to be removed from the system
- Neurotransmitters- chemical messengers
- Potency- the amount of drug needed to achieve maximum effect
- Receptors- molecules situated on the cells

Side Effect presentations/complaints in session

- Why are they getting up and moving around all the time?
- Lip smacking, sticking tongue out?
- Vivid dreams
- No sex drive or unable to have an orgasm
- Food tastes weird, Coke tastes flat
- Irritable
- Apathetic

Side effects (cont.)

- Shuffling feet
- Vision changes
- Gl disturbances
- Grinding teeth
- Sunburn type rash
- Seizures
- Weight gain

Brain Anatomy & Function

"Your brain is a three pound universe that processes 70,000 thoughts each day using 100 billion neurons that connect at more than 500 trillion points through synapses that travel 300 miles/hour." Cleveland Clinic

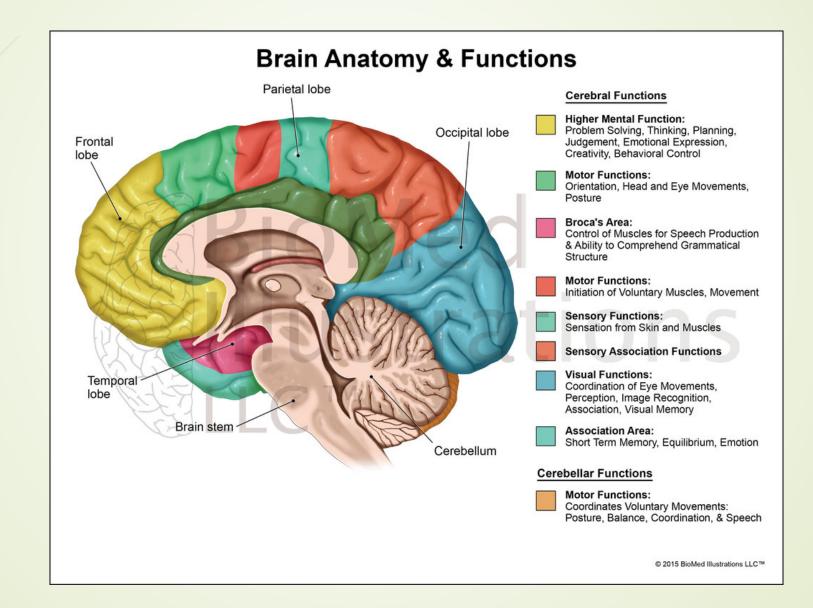
https://healthybrains.org/brain-facts/

Brain Anatomy & Function

Three main parts:

- Cerebrum: largest part of the brain, remembering, thinking, feeling, problem solving and movement control are all done here
- Cerebellum: located under cerebrum and in the back of the skull. Controls coordination and balance.
- Brain Stem: located below the cerebrum and in front of the cerebellum. It connects the spinal cord to the brain. It is responsible for controlling breathing, heart rate, digestion and blood pressure.

Brain Anatomy & Function



Antidepressants, do they work?

Antidepressants and psychological therapies – of which the most frequently used is CBT (cognitive behavior therapy) – have similar success rates. Around 60% of people respond by about two months to the drugs with about a 50% reduction in their symptoms - an improvement in mood, better sleep and so on. But, he said, "about 80% of people stop antidepressants within a month".

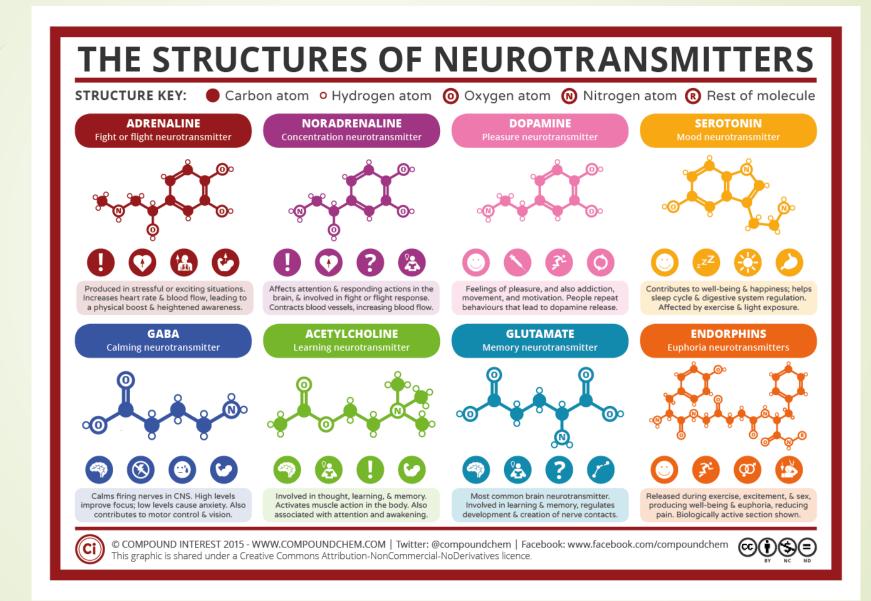
Comparative efficacy and acceptability of 21 antidepressant drugs for the acute treatment of adults with major depressive disorder: a systematic review and network meta-analysis. Over 500 trials examined

Andrea Cipriani, MD Psychiatry Oxford University

Monoamine Neurotransmitters

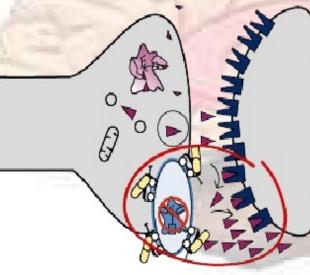
- There are three primary transmitters related to the monoamine hypothesis of depression:
- Serotonin regulates mood, sleep, appetite, adds in attention, learning, libido, pain control, temperature regulation
- Norepinephrine concentration, arousal, learning, memory
- Dopamine regulating movement, working memory, attention, reward motivated behavior

Neurotransmitters

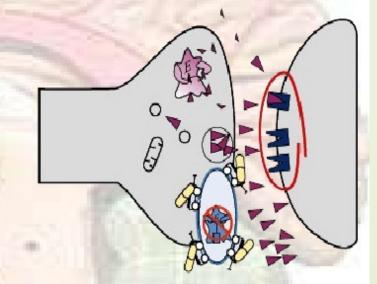


Neurotransmitter Hypothesis

Neurotransmitter Receptor Hypothesis of Antidepressant Action



Antidepressant blocks the reuptake pump, causing more NT to be in the synapse



Increase in NT causes receptors to down-regulate

Stahl S M, Essential Psychopharmacology (2000)

Evidence Supporting Monoamine Hypothesis

- A simplified theory of depression
- Idea that "normal" amounts of the neurotransmitters were depleted by stress, genetics, drug use or some disease process
- Certain drugs that were known to deplete these neurotransmitters could induce depression
- Antidepressants at the time of hypothesis were known to have an affect on the increase of the monoamines
- New evidence points more towards changes in gene expression in neurons targeted by the monoamines. "It is clear that antidepressant agents in current use require an intact monoamine system for their therapeutic effect". Delgado PL. Depression: the case for a monoamine deficiency. J Clin Psychiatry. 2000;61 Suppl 6:7–11.

Depression Facts

- Depression is a very common mental disorder. Globally, there are more than 264 million people of all ages suffer from depression
- Suicide is the second leading cause of death for 15-29 year old's
- Depression is a leading cause of disability worldwide and is a major contributor to the overall global burden of disease.
- Women are affected by depression more than men.

https://www.who.int/news-room/fact-sheets/detail/depression

Depression

- Intense feelings of sadness, despair, hopelessness
- Unable to experience pleasure with activities that are usually pleasurable
- Crying spells
- Change in appetite
- Sleep pattern changes
- Suicidal thoughts

Anxiety-GAD, Panic, OCD (formerly under anxiety)

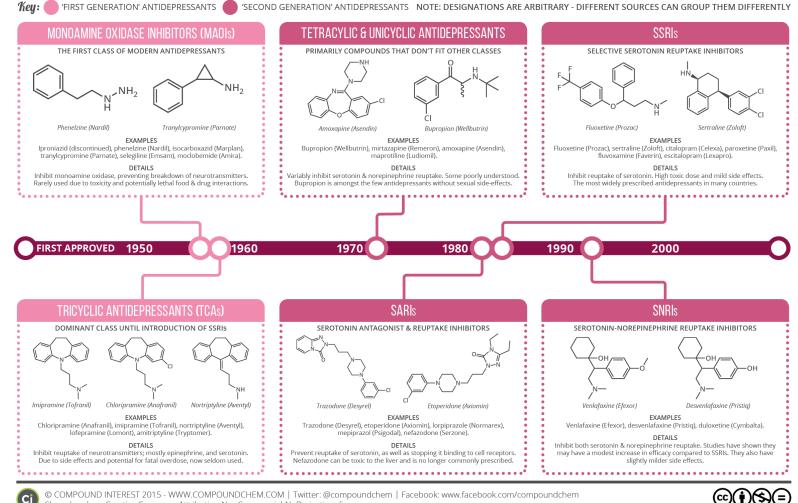
- GAD- persistent worry/anxiety in multiple facets of life, overthinking plans/ solutions, difficulty with uncertainty, indecisiveness, unable to let go, restless, fatigue, GI upset, irritable
- Panic- intense fear peaks in 10 minutes, palpitations, sweating, trembling, difficulty breathing, chest pain, GI upset, lightheaded, fainting, derealization, fears of dying or going crazy, numbing/tingling, especially lips, fear of it happening again
- OCD- Recurrent/persistent thoughts, urges, impulses that are intrusive/ unwanted, usually cause distress.

PTSD

- Recent studies from VA have concluded therapy hedges out medication but that both are preferred and more efficacious.
- Direct/indirect exposure/witnessing trauma, near death or sexual trauma, reexperienced nightmares, flashbacks, intrusive thoughts, triggers, negative thoughts that began/worsened after the trauma, hyperarousal, last longer than a month, distress/function in social/work life and not due to medication/substance use
- SSRI's first line
- Benzodiazepines decrease efficacy of CPT/PE/EMDR (VA requires vets to be off BZD before initiating treatment)

Classes of Antidepressants

MAJOR CLASSES OF ANTIDEPRESSANT DRUGS



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Antidepressants/Antianxiety

- Tricyclic antidepressants (TCA)- amitriptyline, doxepin. Worked very well but had many drug interactions and highly lethal in OD
- Monoamine oxidase inhibitors (MAOI). Drug and food interactions with tyramine, Silence of the Lambs diet (red wine, fava beans, aged meats and cheeses)
- Selective Serotonin Reuptake Inhibitors (SSRI's)- Prozac was first SSRI approved in 1987. First line medication for depression/ anxiety. More tolerated, low lethality.
- Serotonin and Noradrenaline Reuptake Inhibitors (SNRI)-Cymbalta

Antidepressants/Antianxiety

- Norepinephrine and Dopamine Reuptake Inhibitor (NDRI)- Wellbutrin, also used for ADHD and smoking cessation (Zyban), combined with naltrexone is called Contrave, weight loss drug.
- Norepinephrine Reuptake Inhibitor (NRI)- Strattera most well-known to treat ADHD as non-stimulant
- Tetracyclic- Remeron, generally safer than tricyclics

SSRIs/SNRIs

- SSRI-Prozac, Zoloft, Lexapro, Celexa, Paxil
- Most prescribed, first line MDD/GAD, panic d/o, bulimia also used in OCD.
- Generally well tolerated
- SNRI- Effexor, Cymbalta
- Newer, increased efficacy and overall tolerability to TCAs
- Should take for minimum 9 months once effective, stopping sooner has increased chance of recurrent depression

Side Effects/Adverse Effects of SSRI/SNRI

- GI upset (nausea, diarrhea) usually in the beginning, take with food, start low dose
- Sexual dysfunction (SSRI's)
- Sedating or insomnia
- Headache (SSRI's, trazodone and Wellbutrin)
- Dry mouth
- Teeth grinding (SSRI's)
- Irritability

Side Effects/Adverse Effects of SSRI/SNRI

- Tremor (lithium, antipsychotics)
- Hyponatremia (especially in the older population and with Zoloft)
- GI bleeding
- Serotonin syndrome (emergency)
- Discontinuation syndrome- usually from abrupt cessation

Other depression/anxiety meds

- Wellbutrin mainly depression, targets anhedonia, anergia, focus, concentration. Off label for ADHD in children. Also called Zyban (smoking cessation). Some evidence helping curb cravings for stimulants (meth and cocaine).
- Remeron- helps with sleep, higher doses used for MDD/GAD, combined with Effexor (California Rocket Fuel), known to cause weight gain.
- Trazodone- "mild antidepressant" at higher doses, commonly used off label for sleep, helps induce sleep, take and go to bed, read, don't wait for sedation. Common SE are headache and vivid dreams.

Other depression/anxiety meds

- Vistaril- prescription anti-histamine, non habit forming, fast, effective, some sedating effects
- Clonidine- blood pressure med, used for anxiety, as well as opiate withdrawals, FDA approved for ADHD in children
- Anti-psychotics, low dose Seroquel for anxiety/panic

Key Points for Patients

Antidepressants are effective

- No evidence that they lose efficacy over time
- Not known to cause long term side effects
- Non addictive

Benzodiazepines AKA Benzos

- Xanax, Valium, Ativan, Klonopin
- Affects gaba receptors, drug effect felt is similar to alcohol
- Can lead to short term memory & balance loss
- Very effective for short term panic, anxiety, sleep
- Also used to break catatonia and to manage side effects from antipsychotics
- Highly Addictive
- High lethality when combined with alcohol/opiates
- Abrupt cessation at higher does can cause seizures

Bipolar

Episodes of mania/hypomania lasting a week (type I and II), grandiosity, lack of need for sleep, racing thoughts, pressured speech, increase in goal-directed activity, impulsive, high risk behaviors later followed by periods of depression, anhedonia, anergia, recurrent thoughts of death/suicidal ideation all while sober

Borderline Personality Disorder

Difficulty regulating emotion: chronic pit/sense of emptiness, fears of abandonment, idolizing/devaluing relationships, unstable self-image, impulsivity, explosive anger, dissociation, self harm, suicide attempts.

"Mood Stabilizers"

- Include a few different classes of medications
- Used to treat bipolar, border line PD and treatment resistant depression
- Many are anti-convulsant medications with different mechanisms of action
- Lithium-gold standard for bipolar, one of a few medications with indications for suicidality, used in conjunction with antidepressants for treatment resistant depression. Possible that it is altering the balance of NT signaling in the hypothalamus

Mood Stabilizers

- Lamictal- known as a maintenance med to prevent mania, also used for borderline PD, takes time to ramp up due to protocols to prevent Stevens Johnson Syndrome
- Depakote-often used for augmentation (lithium)
- Tegretol
- Antipsychotics (Neuroleptics) (Clozaril, Zyprexa, Seroquel, Abilify) Used to treat/extinguish acute mania and for bipolar depression.

Schizophrenia

- Two or more of the following, each lasting the majority of time in a one month period (hallucinations, delusions, disorganized speech, grossly disorganized/catatonic behavior and negative symptoms.
- Effecting work/social/self care aspects
- Continuous signs of episode for at least 6 months (must include 1 months of the symptoms above, unless its successfully treated)

Antipsychotics (Neuroleptics)

- Two classes, first and second generation antipsychotics (SGA's)
- Used to treat psychosis (delusions, hallucinations, paranoia and disordered thought)
- First generation "typicals", used mainly for positive symptoms such as delusions and hallucinations. Includes Haldol and Thorazine which was first antipsychotic, developed 1950, still used in some cases.
- Affects dopamine receptors, first generation consider toxic to brain by some researchers
- First generation have higher risk of EPS (Extra Pyramidal Symptoms): muscle spasms, tardive dyskinesia (irregular, jerky movements), parkinsonism (rigidity, speech changes), akathisia (restlessness, unable to sit still, need to move around to feel relief)

Antipsychotics (Neuroleptics)

- Second Generation "atypicals", used to treat negative symptoms flat affect, apathy anhedonia as well as positive symptoms
- Less EPS symptoms but still possible
- Greater metabolic effects, weight gain, hypertension, increase risk of diabetes
- Increased risk of stroke in older patients
- Some claim more effective at treatment of schizophrenia but very limited evidence

ADHD

- Inattention and impulsivity/hyperactivity are the core symptoms
- Dx based on clinical assessment, age of onset and social impairment (ASRS v1.1, Wender Utah Rating Scale screening tools for Adults, Vanderbilt children)
- Strong genetic component
- Neurologic basis
- Affects both genders, worldwide prevalence, persists through adulthood in significant percentage of cases, however evidence of age-dependent decline
- Impacts multiple areas of function
- Highly treatable, most treatable disorder in psychiatry
- Stimulants are first line treatment
- Faraone et al. Nature Reviews Disease Primers 2015

Stimulants/Non-stimulants

- Methylphenidate and amphetamine based stimulants, MPH typically tried first with children, Ritalin the most common
- Work by increasing dopamine & norepinephrine levels in the prefrontal cortex in turn raises motivation, concentration/focus
- Many approved medications, multiple delivery forms, tablets, capsules, liquids, chewable
- With children need to be aware of diversion by parents/guardian

Non Stimulants

- Atomoxetine (Stratera) SNRI
- Clonidine (Kapvay) blood pressure drug
- (Buproprion) Wellbutrin
- Guanfacine (Intuniv)- blood pressure drug
- Fish oil

Side Effects/Adverse Effects

- Decreased appetite
- Insomnia
- Headache
- Moodiness/Irritability
- Tics
- Psychosis, rare but possible even at approved doses
- Cardiac/hypertension rare but possible

Mass General Hospital <u>https://advances.massgeneral.org/neuro/journal.aspx?</u> id=1315

Controlled Substances

Schedule I

Schedule I drugs, substances, or chemicals are defined as drugs with no currently accepted medical use and a high potential for abuse. Some examples of Schedule I drugs are: heroin, lysergic acid diethylamide (LSD), marijuana (cannabis), 3,4-methylenedioxymethamphetamine (ecstasy), methaqualone, and peyote

Schedule II

Schedule II drugs, substances, or chemicals are defined as drugs with a high potential for abuse, with use potentially leading to severe psychological or physical dependence. These drugs are also considered dangerous. Some examples of Schedule II drugs are:

(Vicodin), cocaine, methamphetamine, methadone, hydromorphone (Dilaudid), meperidine (Demerol), oxycodone (OxyContin), fentanyl, Dexedrine, Adderall, and Ritalin

Schedule III

Schedule III drugs, substances, or chemicals are defined as drugs with a moderate to low potential for physical and psychological dependence. Schedule III drugs abuse potential is less than Schedule I and Schedule II drugs but more than Schedule IV. Some examples of Schedule III drugs are:

(Tylenol with codeine), ketamine, anabolic steroids, testosterone

Controlled Substances

Schedule IV

Schedule IV drugs, substances, or chemicals are defined as drugs with a low potential for abuse and low risk of dependence. Some examples of Schedule IV drugs are:

Benzodiazepines (Xanax, Valium, Ativan), Soma, Darvon, Darvocet, Ambien, Tramadol

Schedule V

Schedule V drugs, substances, or chemicals are defined as drugs with lower potential for abuse than Schedule IV and consist of preparations containing limited quantities of certain narcotics. Schedule V drugs are generally used for antidiarrheal, antitussive, and analgesic purposes. Some examples of Schedule V drugs are:

(Robitussin AC), Lomotil, Motofen, Lyrica

https://www.dea.gov/drug-scheduling

Novel Treatments

- Ketamine rapid antidepressant effects, nasal administration, Esketamine has FDA approval, however IV ketamine appears more efficacious
- Psilocybin found in certain mushrooms, blocks serotonin uptake, about to enter phase 3 trials for nasal spray treatment of PTSD.
- MDMD (ecstasy)- being researched for PTSD
- DMT (Ayahuasca root)- possible benefits for PTSD, a few states allow its use for religious purposes
- LSD assisted psychotherapy researched for PTSD

MAT-Medication Assisted Treatment

- Suboxone opiate use
- Methadone opiate use, requires daily visits in beginning to clinic
- Naltrexone/Vivitrol FDA alcohol/opiate use, decreases cravings and blocks dopamine reward pathway. Off label gambling/compulsive sexual behavior/video gaming/binge eating/self harm
- Antabuse alcohol use, reduces cravings, causes significant distress if taken with alcohol
- Acamprosate-cravings, needs three times daily dosing, odd dosage milligram number concerns some people (666mg)
- Gabapentin- alcohol/opiate cravings
- Topamax- alcohol, marijuana, food cravings
- Clonidine opioid withdrawal/cravings