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A REVIEW: DEPRESSION AND ITS MANAGEMENT

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ABSTRACT

Depression is one type of mental disorder that frequently leads to diminished functioning and a low quality of life. During the early phases of a major depressive episode, the aim of treatment is to assist the patient in achieving remission and ultimately getting back to their normal level of activity. Although alternative potential pharmacological therapies are still competing for practitioners' attention, the most common medication for treating acute depression is still a selective serotonin antidepressant The second most popular strategy for assisting patients in overcoming the acute phase, sustaining remission, and avoiding relapses is depression-focused psychotherapy. Sometimes the most effective physical treatment for depression is electroconvulsive therapy; alternative methods have disadvantages, and research is ongoing to determine the precise causes of these drawbacks. It is necessary to codify and streamline treatments for depression, a persistent public health issue, in order to improve present practice. Internationally, a number of treatments with differing degrees of efficacy have been proposed.

KEYWORDS: Internationally, a number of treatments with differing degrees of efficacy have been proposed.

INTRODUCTION

A chronic sense of melancholy, emptiness, and loss of joy are symptoms of depression, a mental condition. It is not the same as the emotional swings that people routinely go through in life. Significant life events, including losing a job or experiencing a bereavement, might set off depression. But the bad emotions that a patient may experience momentarily in reaction to a challenging life event are not the same as depression. Despite a change in circumstances, depression frequently endures and results in deep, persistent feelings that are out of proportion to the individual's circumstances. It is a persistent issue rather than a one-time occurrence. Major depression is the most common type of depression, however there are others. It is characterized by bouts where symptoms persist for a minimum of two weeks. Weeks, months, or even years can pass during a depressive episode. It is a chronic condition that improves and then recurs for many people.[1]

However, the National Institute of Mental Health (NIMH) claims, everyone experiences occasional sadness and unhappiness, but these emotions often go away with time. Clinical depression and major depressive disorder are not the same as depression. Severe symptoms may hinder your capacity for rational thought, emotional regulation, and day-to-day activities like eating, sleeping, and working. This can affect anyone, regardless of age, race, wealth, culture, or educational attainment. Research suggests that

psychological, genetic, and environmental variables may all have an impact on depression. [2]

Subcategories of Depression

- Clinical depression (major depression disorder):
 - The intensity of depression can vary, ranging from mild, transient episodes to severe, persistent depression. Clinical depression, often known as serious depressive illness or major depression, is the most severe type of depression. It is not comparable as depression brought on by a medical ailment, such a thyroid issue, or a loss, like the death of a loved one. [4]
- Persistent depressive disorder(PDD): PDD is the term used to describe mild to severe chronic depression. It involves spending most of the time, on typical days, feeling down or gloomy for a period of time or more. PDD is common and can impact individuals of any age. The most effective treatment consists of medication, therapy, and leading a healthy lifestyle.^[5]
- Disruptive Mood Dysregulation disorder(DMDD): Children and teens with annoying mood dysregulation disorder (DMDD) are constantly irritated and furious, and they often have extreme outbursts of anger. There is more to DMDD symptoms than just a "bad mood." With DMDD, the There are serious signs Children with DMDD can have serious problems at school, at home, and often with their peers. They also typically require mental

health care including appointments with doctors and hospital stays. Furthermore, these kids are more prone to experience anxiety and despair later in life [6]

• Premenstrual dysphoric disorder (PMDD): Premenstrual dysphoric disorder (PMDD) is a severe
and sometimes incapacitating complication of
premenstrual syndrome (PMS). Although the
physical and mental symptoms of PMS and PMDD
are similar, PMDD is distinguished by severe
alterations in mood that can disrupt daily activities
and damage relationships with others. Both PMDD
and PMS symptoms often begin up to ten days
before the period begins and last for the first few
days of the period. [7]

Depressive Disorders Due to Various Medical Conditions

A number of medical conditions might cause bodily changes that can lead to depression. Examples include hypothyroidism, Parkinson's disease, cancer, and heart disease. If the underlying illness is addressed, depression usually improves.^[3]

Additionally, Major depressive illness comes in several forms, such as.

- Seasonal affective disorder (seasonal depression):
 - Seasonal depression, sometimes referred to as a form of seasonal affective disorder (SAD), is one kind of depression. It is caused by the change of the seasons and often begins in late October. Symptoms include weight gain, oversleeping, poor energy, sadness, and indifference in daily activities. Among the therapies include conversation therapy, light therapy, and antidepressants.^[8]
- Prenatal depression and postpartum depression:
 - Depression which occurs during pregnancy is known as prenatal depression. Chronic or severe sadness is a symptom of depression. Anxiety, exhaustion, and difficulty sleeping might also result from it. You can distance yourself from friends and family if you suffer from this mood condition. Activities you used to like may no longer pique your interest. One may experience prenatal depression at any point during your pregnancy. Postpartum depression (PPD) is a type of depression that appears after giving birth. It's crucial to remember that the "baby blues" are not the same as pregnant or postpartum depression. Usually, the "baby blues" go away in two to three weeks. In the meanwhile, without treatment, postpartum and prenatal depression do not go away. Changes in lifestyle can sometimes alleviate depression symptoms. However, if they don't, medical professionals use medicine and therapy to manage this issue. You are not alone if you suffer from prenatal depression. There are therapies for depression, which is a prevalent medical disease.[9
- **Atypical depression:** Atypical depression is a type of depression in which you experience positive

events that temporarily lift your spirits. Additional indicators of atypical depression include increased appetite, hypersomnia, and rejecting hypersensitivity. It can be treated with psychotherapy and medications. [10]

Depression-Related Disorders

- Anxiety: Despite being distinct disorders, anxiety and depression frequently coexist. They are treated similarly as well. [11] Strong, uncontrolled feelings of worry, panic, anxiety, and/or anxiety are hallmarks of anxiety disorders. These emotions may last for a long period and interfere with day-to-day tasks. [12] Feeling depressed or experiencing the blues occasionally is normal. And everyone experiences anxiety occasionally; it's a common reaction to stressful circumstances. On the other hand, persistent or severe anxiety and depression may indicate a mental illness. [11] Usually, talk therapy, often known as psychotherapy, antidepressant medications, or both can help with the symptoms of both disorders. Lifestyle changes that encourage regular exercise, less stress, greater social support, and better sleep may also be helpful. If you suffer from any of these conditions, alcohol, smoke, or recreational drugs. They can worsen both issues and make therapy more difficult.[12]
- PTSD: It is possible to experience depression and post-traumatic stress disorder (PTSD) simultaneously. Because of their similar symptoms, people may mistake them for one another. PTSD can result from traumatic experiences (Trusted Source). Within three months after the upsetting incident, symptoms may appear. Traumatic experiences may also be followed by depression. According to research, between 30% and 50% of PTSD sufferers also suffer from depression. Furthermore, compared to those without PTSD, those with PTSD have a 3 to 5 times higher risk of developing depression. [13]

Depression and a diagnosis of post-traumatic stress disorder (PTSD) frequently co-occur. Anxiety, flashbacks, and reliving terrible events are hallmarks of post-traumatic stress disorder (PTSD). The illness appears after a stressful event, like a natural disaster, car crash, assault, abuse, or conflict. Conversely, depression is typified by poor mood, fluctuations in energy levels, and a loss of interest and enjoyment.

It is perhaps not unexpected that these two illnesses might coexist because depression is also a common reaction following a stressful or traumatic incident. According to research, 6.8% of people may experience PTSD at some point in their lives. In any given year, 8.3% of American adults are thought to suffer from serious depressive disorder. According to one study, 35.2% of individuals with PTSD also satisfied the diagnostic requirements for major depressive disorder,

however numbers vary. Among veterans with PTSD, 68% also have depression. [14]

• **BPD**: - Emotional instability is a feature of both depression and Borderline Personality Disorder (BPD), albeit they might show up in various ways. People with depression frequently express a lingering sense of melancholy and emptiness, but people with BPD may experience intense and quickly shifting emotions. Emotions can be like a rollercoaster ride for those with BPD, with overwhelming highs and lows. They can find themselves feeling ecstatic one minute, and then furious or depressed the next. People with BPD may find it difficult to control their emotions and preserve healthy relationships as a result of these abrupt emotional changes.

On the other hand, people who are depressed could experience a persistent sense of hopelessness and heaviness. They might experience more static feelings, such as a persistent sense of melancholy that lasts for weeks, months, or even years. People who suffer from depression may find it difficult to find motivation or joy in their everyday life as a result of this ongoing melancholy. Giving people with these diseases the right care and support requires an understanding of and ability to distinguish between these emotional experiences. To create individualized treatment plans, therapists and other mental health experts must thoroughly evaluate and pinpoint each patient's unique emotional patterns and difficulties.^[15]

Eating Disorders: - Eating disorders and depressive disorders are both complex and diverse conditions. Studies have shown a correlation between eating disorders and a greater incidence of depressed symptoms and disorders. Personality disorders, anxiety disorders, and drug addiction in bulimia nervosa, or obsessivecompulsive disorders and anxiety-related disorders in anorexia nervosa, are less prevalent mental syndromes that have a substantial impact on the onset and maintenance of eating disorders. More research will be required since few, studies have examined how fasting might alter one's physical, endocrine, or psychological makeup to resemble symptoms thought to be associated with depression. The evidence for a common etiology for bulimia nervosa is at best questionable, whereas for anorexia nervosa it is weak. Women who display depressed or self-critical qualities will be differentially recruited into eating disorders, as dieting-induced weight reduction is the main cause of eating disorders in contemporary instances. The hypothesis that best fits the data would suggest a link between an eating disorder and all of the depressive illnesses from no depression to serious depression. Patients with bulimic anorectic and bulimia nervosa would have slightly higher rates of depression than patients with restricting anorexia nervosa, but each eating disorder would have its own pathophysiology and psychopathology. [16]

Etiology of depression

The etiology of severe depressive disease is complicated and involves both hereditary and environmental factors. Even while melancholy can strike anybody without a familial history of sadness, Some evidence suggests that hereditary factors are less likely to affect late-onset depression than early-onset depression. There are several biological risk factors for depression in the elderly. Depression has been linked to increased rates of neurodegenerative diseases (such as Parkinson's and Alzheimer's), stroke, autoimmune disorders, seizures, cancer, macular degeneration, and chronic pain. The initial symptoms of depression are brought on by problems and occurrences in life. Numerous stresses. including as traumatic experiences like dying or the passing of someone you love, a lack of or weakened social support, the strain of taking care of others, monetary challenges, relationship issues, and arguments, can cause depression.^[17]

Pathophysiology of depression

Pathophysiology: Environmental Factors and Genetic Predisposition

There is a genetic component to major depression that passes in families. However, given the variety of symptoms, developmental and environmental variables have to be taken into account when identifying the elements that contribute to severe depression. One viewpoint on mood disorders is the connection between susceptible genes and environmental variables, pressures in life combined with a possible malfunction in the serotonin (5-HT) pathway. The serotonin transporter may modulate the serotonergic reaction to stress and aids in the absorption of serotonin at the synapse. Compared to people homozygous for the 1 gene, Significant sadness and suicidal ideation in response to stress were more common in people has the s allele present in two copies. Additionally, after four or more stressful events, people with 2 s alleles were twice as likely to have severe depressive episodes (McCance & Huether, 2014). [18]

• Pathophysiology: Neurochemical dysregulation

The body's neurotransmitters can be elevated by antidepressant drugs which supports a different viewpoint known as the monoamine hypothesis of depression. According to this theory, depression results from a deficiency in the brain's norepinephrine, dopamine, and/or serotonin concentrations. Increasing the levels of monoamine neurotransmitters at synapses is the aim of antidepressant therapies (McCance & Huether, 2014). [18]

• Pathophysiology: Neuroendocrine Dysregulation

The underlying cause of depression is linked to two theories: neuroendocrine system malfunction. The hypothalamic-pituitary-adrenal system and stress are the subjects of the first one. A person's capacity to control stress is significantly impacted by the HPA system. A link between depression and a malfunctioning system is suggested by the discovery that 30 to 70 percent of those

with severe depression had persistent stimulation of their HPA system and chronic cortisol production. The body secretes pro-inflammatory cytokines as a result of chronic cortisol release, which leads to inflammation and immunosuppression. There is also a neurotrophic theory for depression. It is believed to concentrate on the hippocampus's neuronal atrophy, which suppresses cell proliferation and, as a result, lowers the amount of BDNF. It has been suggested as an addition to the depression monoamine hypothesis. The second hormonal anomaly occurs in the thyroid, pituitary, hypothalamus systems. Twenty to thirty percent of cases of severe depression have been discovered to have abnormalities in the hypothalamic-pituitary-thyroid (HPT) system. while the exact nature of this dysfunction is unknown. In response to TRH challenge, thyrotropinreleasing hormone increases, thyroid stimulating hormone decreases, and the typical nocturnal rise in Thyroid stimulating hormone (TSH) levels is halted. Relapse risk is increased by all of these factors (McCance & Huether, 2014). [18]

• Pathophysiology: Neuroanatomic and Function Abnormalities

Depressive individuals' post-mortem brains usually have decreased serotonin 5-HT1a Serotonin carrier binding in the cerebral cortex and hippocampus, as well as receptor type affinity in the limbic, frontal, and temporal cortex, indicating a raphe-serotonin system issue. It is possible to inhibit the raphe-serotonin system by activating the locus cerulean-norepinephrine system. This implies a secondary function in the regulation of serotonin. Some depressed suicide victims have changes in their frontal brain that are related to norepinephrine receptors. Depression-related sleep and arousal issues, as well as issues with focus or attention, may be associated with changes in norepinephrine systems.

Individuals with unipolar illnesses have been found to have less glial cells due to changes in the frontal and limbic regions (including the amygdala). decreases in frontal lobe volume and prefrontal cortex function. It has also been shown that individuals with depression exhibit abnormalities in glucose metabolism and cerebral blood flow. Similar to those observed in schizophrenia, speech issues and delays in cognitive processing may arise from dorsolateral prefrontal abnormalities in depression. Mnemonic and attentional problems that accompany mood disorders may be linked to dorsomedial frontal dysfunction. The metabolism and blood flow in the frontal brain are elevated. In those who are depressed, it has a favorable correlation with unpleasant emotions (McCance & Huether, 2014). [18]

Causes of Depression

The illness of depression is complicated. There are several potential reasons, even though the precise one is unclear. When they have a serious illness, some people develop depression. Other people may experience depression as a result of significant life changes, such as

moving or losing a loved one. Some people may have a familial history of depression, while others may feel a great deal of loneliness and sadness for no apparent cause. [19]

- Abuse: Experiencing physical, sexual, or emotional abuse might raise your chance of later getting depression.
- Age: Older adults are more susceptible to depression. Additionally, two factors that might raise the risk include living alone and lacking social support.
- Death or loss: Even while it's normal to feel sad or grieve after losing a loved one, these emotions may make you more susceptible to melancholy.
- **Conflict**: If you regularly have disagreements or personal confrontations with friends or family, you may be more prone to depression.
- **Some drugs**: Corticosteroids, the antiviral drug interferon-alpha, and isotretinoin, which is used to treat acne, are among the drugs that might increase your risk of getting depression.
- Gender: The likelihood of depression in women is around double that of males. Nobody knows for sure why. One explanation might be the hormonal shifts that women experience at different stages of their lives.
- Genes: A family descent from depression may raise your risk. Since depression is thought to be a complex characteristic, several genes, each with a little effect, are probably involved rather than a single gene driving illness risk. Unlike conditions like Huntington's disease or cystic fibrosis, which are purely hereditary, depression, like other mental illnesses, has a more complicated genetic composition.
- Important events: Even happy life events, such as marriage, finishing school, or beginning a new career, can cause depression. Divorce, retirement, losing a job or other source of income, and moving can all have an effect. However, depressive disorder is never just a "normal" response to difficult life circumstances.
- Other personal problems: Your chance of getting clinical depression may be raised by problems such as being excluded from familial or friends or social isolation resulting from other mental diseases.
- Major illnesses: Depression can occasionally coexist with a serious disease or be brought on by another medical condition.
- **Drug abuse**: More than 30% of those who struggle with substance addiction also have major or severe depression. Even if they temporarily improve your mood, both alcohol and drugs will make your depression worse. [19]

Symptoms of Depression

A useful starting point for learning about mental illness is the Diagnostic and Statistical Manual of Mental

Disorders (DSM). Mental health professionals classify and diagnose mental diseases using the DSM.

The DSM lists the following as signs of major depressive disorder.

- Depressed mood
- Lack of interest or enjoyment
- Weight gain or loss
- Excessive sleeping or difficulty falling asleep
- Speaking or moving more slowly or more quickly than usual
- Fatigue
- Feelings of regret or worthlessness
- Incapacity to concentrate or form opinions
- Suicidal thinking.^[20]
- A decline in sexual desire
- Abnormally sluggish or agitated motions. [21]

A person may be diagnosed with depression by a doctor if they have five or more of these symptoms in a twoweek period. Along with aggravating the symptoms of chronic pain, headaches, and digestive disorders, depression can also result in other symptoms, such as restlessness and irritability.[21]

Diagnosis of Depression

Mental health professionals often use the Guidelines for the Diagnosis and Statistic Manual of Mental Disorders, 5th Edition (DSM-5-TR) to diagnose major depressive disorder. For a diagnosis with depression, a person must have signs for the majority of the time, or almost every day for a minimum of two weeks. One of the signs must be a depressed mood or a loss of enthusiasm or pleasure in most activities. Children and teens may feel agitated rather than depressed. [22]

In order to be diagnosed with depression, a person needs to.

- Possess a minimum of five signs of depression
- For a minimum of two weeks, put up with the signs and symptoms for the majority of each day.
- Symptoms include depression and a decline in enjoyment or interest in almost all activities. Children and adolescents are subject to slightly different criteria. Instead of being in a depressed state, they might appear irritated.

Depression symptoms include

- A mood of depression
- Lack of enjoyment or interest in nearly every
- Notable variations in weight
- Alterations in appetite
- Sleep disorders
- Weariness and a lack of energy
- Frantic gestures or actions, including fumbling or talking quickly

- Difficulties focusing or thinking
- An excessive, improper, or deluded sense of guilt
- A feeling of devaluation
- Recurring death-related ideas, such as suicidal thoughts or thoughts

To rule out additional psychological illnesses that could be generating the same symptoms is another stage in the diagnosing process, like:

- Bipolar illness
- Problems related to substance or alcohol usage
- Psychotic illnesses, such as schizophrenia

To aid in the development of a treatment plan, a medical expert will also make an effort to assess the severity of the depression. Loss of interest in activities may be a sign of severe depression, according to a 2018 study. [22]

Gender-Based Analysis of Depression

1. 2019 (based on data from NMHS 2015–16)

10.3% of males (based on the prevalence of mental illnesses among men)

Women: 11.2% (the prevalence of depression in women is slightly greater).

2. 2020 (according to the WHO COVID-19 Survey)

Men: 9.0% (a little drop from overall rates among young)] 10.2% of women (prevalence greater among women, particularly during lockdown)

3. 2021 (according to the report from Sapiens Labs)

Men: 11.0% (pandemic stress-related general rise)

Women: 12.5%; women have been found to have higher rates of mental anguish.

2022 (according to the Sapien Labs report and WHO data)

Men: 12.0% (an increase in mental anguish in general).

Women: 15.0% (particularly high given that pandemicrelated variables have disproportionately affected women)

5. 2023 (according to the report from Sapien Labs)

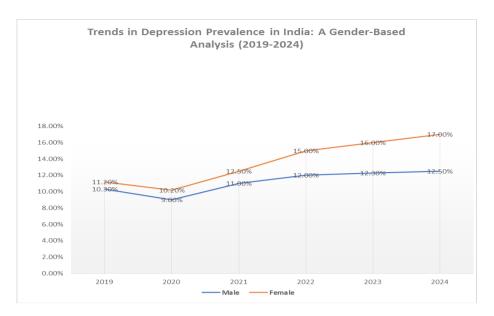
Males: 12.3% (depression rates among males have been steadily rising).

Women: 16.0% (the greatest percentage among women, steadily rising over time)

6. 2024 (Predicted Amounts according to Current **Data Patterns**)

Men: 12.5% (greater because of lifestyle variables and

Women: 17.0% (depression rates among women are still on the rise, particularly due to persistent socioeconomic issues). [24][25][26][27]



Treatment for depression

Similar to diabetes, depression can affect some people for the rest of their lives. Individuals with diabetes can lead long, healthy lives if they take their prescriptions and modify their lifestyle. The same is true with depression. Changes in lifestyle, support groups, medication, and therapy are common ways to manage depression. Making lifestyle adjustments like eating a healthy diet, getting adequate sleep, and practicing mindfulness and physical activity might help some people manage their depression. Others discover that taking medicine makes it easier for them to deal with depression on a daily basis. For many people, therapy also changes their lives. Individuals with depression frequently employ a mix of these therapies. However, it's about determining which treatments are most effective for you.[28]

Depression happens to be one of the commonest mental conditions to treat. Ultimately, between 70% and 90% of people with depression benefit from therapy. In order to diagnose depression, a doctor will do a thorough diagnostic examination that includes a thorough interview to discuss your symptoms and your personal, and medical history. Additionally. family, should examination comprehensive physical he conducted to check for underlying medical issues such as vitamins, hormone levels of imbalances, neurological diseases, and drug or alcohol addiction that might mimic sadness. Laboratories and imaging examinations may be included in the evaluation as a component of the healthcare screening. The evaluating physician will take each of these factors into account when diagnosing a patient and recommending a personalized treatment plan. [29]

Psychotherapy

Psychotherapy, sometimes referred to as psychological treatment or counseling, can help people with depression acquire new patterns of thought and conduct and help them overcome depressing behaviors. In one-on-one or

group settings, psychotherapy takes place under the supervision of a qualified and licensed mental health practitioner. Psychotherapy can be administered online through telehealth or in person. A provider may use mobile or digital technology, such as apps or other tools, to enhance or assist therapy.

Interpersonal therapy and cognitive behavioral therapy are evidence-based treatments for depression. Some persons who suffer from depression may also benefit from short-term use of alternative psychotherapies, such as psychodynamic therapy.^[30]

- Cognitive behavioral therapy (CBT): By educating people to challenge and change negative beliefs and behaviors, cognitive behavioral therapy (CBT) assists patients with anxiety and depression. Recent advancements in cognitive behavioral therapy include incorporating mindfulness principles and customizing the treatment to target specific issues, such sleeplessness. [30]
- Interpersonal therapy (IPT): IPT is concerned with how relationships and life experiences impact emotion and vice versa. IPT aims to help people build social support networks, abilities to communicate in partnerships, and reasonable expectations so they may better manage crises or other Problems that might be causing or exacerbating depression. [30]

Pharmacological Treatment

Depression can be effectively treated with antidepressant drugs. It may occasionally be necessary to try several antidepressant drugs before determining which one is most effective for the patient. The person seeking treatment is likely to benefit from medications that have helped a close family member. It often takes two to four weeks for antidepressants to start to affect symptoms. Usually, improvements in appetite, sleep, and focus come before a mood rise. Before concluding that the drug is not functioning, it is crucial for those undergoing

therapy to give it some time. Although adverse effects are possible with antidepressants, many of them may go away with time. It's crucial to discuss the adverse effects with a healthcare professional and avoid stopping the antidepressant on your own. Abruptly stopping the medicine might be dangerous and result in unpleasant withdrawal symptoms. It would be preferable to take the drug as directed for the entire recommended duration, and then, with a doctor's assistance, to cease it gradually and safely.[31] Antidepressants are a useful therapeutic tool. Some people may be at risk from them, particularly kids, teenagers, and young adults. Children are rarely taken antidepressants, and teenagers are not often treated with them as their first line of treatment. Some people may experience adverse responses when taking antidepressants for the first time. Before it starts to work, some people could get upset. Others might attempt suicide or have suicidal thoughts. Antidepressant users should be regularly watched, particularly when they initially begin taking the medication. It's crucial to keep in mind that, for most individuals, the risks of untreated depression are far higher than the risks of taking antidepressant medications under a doctor's close supervision.[31]

Antidepressants are often classified into many categories by doctors. These consist of. [32]

> Selective serotonin reuptake inhibitors (SSRIs)

SSRIs are a First-line Trusted Source treatment option for Depression.

Most commonly used SSRIs are.

- Fluoxetine
- o Fluvoxamine
- Citalopram
- o Escitalopram
- o Paroxetine

> Serotonin-norepinephrine reuptake inhibitors (SNRIs)

SNRIs are the more modern category of antidepressants than SSRIs. However, they work identically.

Along with depression doctor prescribe SNRIs for.

- Menopausal symptoms
- o Obsessive-compulsive disorder (OCD)
- o Attention deficit hyperactivity disorder (ADHD)
- o Anxiety disorders
- o Fibromyalgia
- Persistent neuropathic pain

Examples of most commonly used SNRIs include.

- Duloxetine, also known as Cymbalta
- Venlafaxine, also known as Effexor XR
- Pristiq (desvenlafaxine)

> Tricyclic antidepressants (TCAs)

Doctors recommend TCAs for.

- Depression
- o Fibromyalgia
- anxiety

- o Insomnia
- Chronic pain

Examples of TCAs includes.

- Amoxapine
- Amitriptyline
- Camopramine (Anafranil)
- o Desipramine (Norpramin)
- O Doxepin (Sinequan)
- o Imipramine (Tofranil)
- Nortriptyline (Pamelor)
- Protriptyline (Vivactil)
- Trimipramine (Surmontil) are among the medications.

Monoamine oxidase inhibitors (MAOIs)

Since MAOIs have negative side effects and interact with other medications, doctors recommend them as a first-line treatment for depression. They might be a possibility, though, if the depression is not improving with treatment.

Examples of MAOIs includes.

- Phenelzine (nardil)
- o Transylcypromine (parnate)
- Isocarboxazid (marplan)
- Selegiline (emsam, eldepryl)

Mechanism of Action

Targeting certain neurotransmitters, each antidepressant modifies mood and behavior in a somewhat different way. Every antidepressant now available is believed to increase serotonin, norepinephrine, or both in the synapses. These neurotransmitters can be increased in a number of methods, but antidepressant drugs concentrate on nerve terminal reuptake.^[33]

Selective serotonin reuptake inhibitors (SSRIs)

5-hydroxytryptamine/serotonin (5HT) is absorbed into presynaptic terminals by SERT, and neuronal absorption is the main mechanism that inhibits neurotransmission via 5HT. SSRIs prolong and improve serotonergic neurotransmission while blocking reuptake. Continuous SSRI treatment has long been associated with increases in cyclic AMP signaling and nuclear transcription factor phosphorylation, as well as enhanced neurogenesis and trophic factor expression, including BDNF. SSRIs are currently the first-line therapy for depression. [33]

> Serotonin-norepinephrine reuptake inhibitors (SNRIs)

By preventing serotonin and norepinephrine from being reabsorbed in the synapse, serotonin and norepinephrine reuptake inhibitors (SNRIs) increase postsynaptic receptor activation. SNRIs bind to the norepinephrine and serotonin transporters in different ways.

Milnacipran and levomilnacipran are more selective in blocking norepinephrine reuptake than serotonin reuptake when compared to other selective serotoninnorepinephrine reuptake inhibitors such as duloxetine, venlafaxine, and desvenlafaxine. [33]

> TCAs, or Tricyclic antidepressants,

TCA, like amitriptyline, prevents serotonin and norepinephrine from being reabsorbed at the postsynaptic neuronal membrane's membrane. Histamine H1 and muscarinic M1 receptors are likewise favored by amitriptyline. As a result, TCA may have sedative and anticholinergic effects. [33]

➢ Monoamine oxidase inhibitors (MAOIs)

Despite being among the initial antidepressant to be discovered, monoamine oxidase inhibitors (MAOIs) are not regarded as the first-line therapy for depression because of their adverse effects and interactions with other medications. The monoamine oxidizing enzyme, which catabolizes dopamine, serotonin, and norepinephrine, is inhibited by MAOIs. [33]

Effectiveness of the antidepressant drugs

The best drugs for addressing the symptoms of depression are antidepressants. However, antidepressants may benefit some people more than others, as is the case with many other drugs.

Research indicates that the antidepressant's "effectiveness" or benefit is often correlated with the degree of the depression; the more severe the depression, the higher the benefit. In most cases, antidepressants work well for moderate, severe, and long-term depression. They often don't assist with moderate depression. It is crucial to keep in mind that psychotherapy is an additional crucial component of depression treatment. Your symptoms will often improve the most when you combine psychotherapy with depression medication. [34][35][36][37][38][39][40][41]

Antidepressant classes	Effectiveness					
	Adults	Male	Female	Age (18-30)	Age (30-60)	Age 60+
SSRIs	60-70%	65%	60-75%	65%	60%	50-55%
SNRIs	65-75%	70%	65-75%	70%	65%	60-65%
TCA	50-60%	55%	50-55%	60%	50-55%	40-55%
MAOIs	55-65%	60%	55-60%	60%	55%	50%
Atypical Antidepressants	60-70%	65%	60-70%	70%	65%	55-60%

CONCLUSION

A major, long-term illness, depression can have an impact on many facets of a person's life. It is deadly when it leads to suicide thoughts. Depression is not something that can be overcome by thinking. Depression is not an indication of frailty or personal failure. Early therapy may improve the likelihood of recovery, and it is curable. Depression may be challenging to cure, therefore a patient ought to seek psychotherapy from a doctor who specialises in depression as well as be willing to attempt a range of therapies. The combination of medication and therapy often yields the best results. Other kinds of drugs, including antidepressants, can occasionally be helpful in treating depression. Many of these are intended to treat other illnesses, such bipolar disorder or anxiety. One obvious example is atypical antipsychotics. Many individuals with a diagnosis of depression also exhibit signs of other mental and physical illnesses. Depression may also benefit from treatment of these other illnesses. Additionally, there are other options for treating depression than medication. For many people, therapy is beneficial. Making lifestyle adjustments can also be beneficial; increasing your exercise, getting more sleep, eating foods that make you feel healthy and energized, and engaging in fun hobbies are all wonderful places to start. Through regulatory changes, digital projects like Tele MANAS, and increasing access to treatments through the National Suicide Prevention Strategy, Ayushman Bharat HWCs, and NMHP, India has achieved significant strides in the

field of mental healthcare. India needs to invest in digital mental health solutions, increase worker training, and bolster awareness efforts going forward. Individual wellbeing, economic expansion, and national development all depend on a psychologically healthy India, which calls for a comprehensive social strategy to make mental healthcare available, inclusive, and stigma-free.

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