

# Major Depressive Disorder Across the Lifespan: Children, Adolescents, and Adults — Comprehensive Clinical Guide (2025)

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Definitions • Epidemiology • Etiology • Clinical Features • Diagnosis • Treatment • Prognosis • References  
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## Definition & Diagnostic Criteria

- MDD: persistent depressed mood and/or anhedonia with  $\geq 5/9$  DSM-5 symptoms for  $\geq 2$  weeks, causing impairment.
- Symptoms: depressed/irritable mood, anhedonia, appetite/weight change, sleep disturbance, psychomotor changes, fatigue, guilt/worthlessness, poor concentration, suicidality.
- In children/adolescents: irritability may substitute for depressed mood.
- Chronic form: Persistent Depressive Disorder ( $\geq 2$  yrs adults;  $\geq 1$  yr youth).

## Epidemiology & Course

- Prevalence:  $\sim 1\text{--}2\%$  in children; up to  $5\text{--}8\%$  in adolescents;  $6\text{--}7\%$  annual in adults; lifetime prevalence  $\sim 16\%$  adults.
- Trends: rising youth depression, especially post-COVID-19.
- Course: untreated episodes last  $6\text{--}12$  months; recurrence common ( $>60\%$  adolescents within 5 yrs).
- Early onset predicts adult recurrence and chronicity.

## Etiology & Risk Factors

- Multifactorial: genetic vulnerability (heritability  $\sim 40\%$ ), neurotransmitter dysregulation, HPA-axis, inflammation, neuroplasticity deficits.
- Risk modifiers: early trauma, neglect, chronic stress, bullying, loss, parental psychopathology.
- Cognitive: negative schemas, rumination.
- Adolescents: developmental transitions and self-perception distress increase vulnerability.

## Clinical Features

### Children

- Irritability, somatic complaints, school refusal, poor peer relations.

### Adolescents

- Anhedonia, academic decline, social withdrawal, suicidal ideation more common.

### Adults

- Depressed mood, anhedonia, psychomotor changes, fatigue, impaired concentration, guilt, suicidality.

### Associated

- Anxiety, ADHD, substance use, somatic symptoms, sleep problems.

## Assessment & Diagnosis

- Clinical interview: onset, duration, impairment, suicidality, comorbidity.
- Tools: PHQ-9 (adults), PHQ-A/CDI (youth).
- Collateral info from parents/teachers in youth.
- Rule out: bipolar disorder, medical causes (thyroid, anemia), substance/medication effects.

## Treatment — Psychotherapy

- CBT: strong evidence across ages.
- Interpersonal Therapy (IPT): effective, esp. in adolescents.
- Behavioral activation, family therapy, supportive therapy.
- Digital/online therapies emerging for youth.

## Treatment — Pharmacotherapy

### Children/Adolescents

- Fluoxetine first-line; escitalopram also used.
- Monitor closely for suicidal ideation (FDA black-box warning).

### Adults

- SSRIs, SNRIs, bupropion, mirtazapine, TCAs, MAOIs (latter less common).
- Augmentation: antipsychotics, lithium, thyroid hormone.

### General

- • Combine with psychotherapy for moderate-severe cases; continue treatment 6–12 months after remission.

### Advanced/Resistant Cases

- • ~40% of youth show treatment resistance.
- • Options: ECT (adults, severe/refractory), rTMS, ketamine/esketamine, investigational neuromodulation.
- • Multimodal integration often needed.

### Prognosis & Outcomes

- • Many remit, but relapse/recurrent risk is high.
- • Early onset associated with poorer prognosis.
- • Functional impairment: academic, occupational, social, increased suicide risk.
- • Treatment reduces relapse, improves QoL, decreases suicide and morbidity.

### Clinical Snapshots

#### Child/Adolescent

- ☐ Depressed/irritable mood + ≥4 other symptoms, ≥2 wks
- ☐ Use PHQ-A/CDI; assess suicidality, comorbidity
- ☐ First-line: CBT/IPT; fluoxetine if moderate-severe

#### Adult

- ☐ Depressed mood/anhedonia + ≥4 others, ≥2 wks
- ☐ Rule out bipolar/medical/substance causes
- ☐ First-line: SSRI/SNRI or CBT; augment if resistant

### References (selected, 2019–2025)

- • American Psychiatric Association. DSM-5-TR criteria for MDD.
- • Cheung, A. H., Zuckerbrot, R. A., et al. (2023). GLAD-PC guidelines for adolescent depression. *Pediatrics*.
- • Feeney, J. C., et al. (2022). Antidepressants in children/adolescents: meta-analysis. *J Child Psychol Psychiatry*.
- • Franklin, T. B., et al. (2024). Depression: Nature Reviews Disease Primers.
- • JAMA Psychiatry (2018–2024). Epidemiology and treatment of adult MDD.
- • March, J., et al. (2020). Treatment-resistant depression in youth. *Child Adolesc Psychiatr Clin*.
- • WHO (2023). Depression fact sheet and global burden data.



## References (selected, 2019–2025)

1. American Psychiatric Association. (2022). DSM-5-TR: Diagnostic and statistical manual of mental disorders, text revision.
2. Cheung, A. H., Zuckerbrot, R. A., et al. (2023). Guidelines for Adolescent Depression in Primary Care (GLAD-PC). Pediatrics.
3. Feeney, J. C., et al. (2022). Antidepressants in children and adolescents: meta-analysis. J Child Psychol Psychiatry.
4. Franklin, T. B., et al. (2024). Major depressive disorder. Nature Reviews Disease Primers.
5. Merikangas, K. R., et al. (2018). Prevalence and treatment of MDD in the US. JAMA Psychiatry.
6. March, J., et al. (2020). Treatment-resistant depression in youth. Child Adolesc Psychiatr Clin.
7. World Health Organization. (2023). Depression fact sheet.