

Mood Stabilizers: Advanced Clinical Reference



Comparative Pharmacology and Clinical Applications

This comprehensive reference provides detailed comparisons between mood stabilizers for psychiatric prescribers, with evidence-based clinical pearls and monitoring recommendations.



Mechanism of Action and Pharmacodynamics

AGENT	PRIMARY MECHANISMS	SECONDARY MECHANISMS	CLINICAL IMPLICATIONS
Lithium (Chokhawala et al., 2024)	<ul style="list-style-type: none"> • Na⁺/K⁺ ATPase inhibition • GSK-3β inhibition • Inositol monophosphatase inhibition 	<ul style="list-style-type: none"> • BDNF upregulation • Anti-inflammatory effects • Modulation of G-protein signaling 	<ul style="list-style-type: none"> • Narrow therapeutic index • Neuroprotective effects • Anti-suicidal properties
Valproate (Rahman et al., 2023)	<ul style="list-style-type: none"> • GABA potentiation • Voltage-gated Na⁺ channel blockade • Histone deacetylase inhibition 	<ul style="list-style-type: none"> • GSK-3β inhibition • Inhibition of protein kinase C • Inhibition of NMDA mediated neuronal excitation 	<ul style="list-style-type: none"> • Broad spectrum efficacy • Rapid onset of action • Multiple formulations available
Lamotrigine (Betchel & Saadabadi, 2023)	<ul style="list-style-type: none"> • Voltage-gated Na⁺ channel blockade • Inhibition of glutamate release 	<ul style="list-style-type: none"> • HCN channel modulation • Weak 5-HT₃ antagonism 	<ul style="list-style-type: none"> • Primarily depression preventing • Minimal cognitive effects • Gradual titration required
Carbamazepine (Maan & Saadabadi, 2023)	<ul style="list-style-type: none"> • Voltage-gated Na⁺ channel blockade • Adenosine A₁ receptor agonism 	<ul style="list-style-type: none"> • GABA potentiation • Adenosine reuptake inhibition • Modulation of Ca²⁺ channels 	<ul style="list-style-type: none"> • Strong CYP3A4 inducer • Autoinduction of metabolism • Multiple drug interactions
Oxcarbazepine (Preuss et al., 2021)	<ul style="list-style-type: none"> • Voltage-gated Na⁺ channel blockade • Ca²⁺ channel inhibition 	<ul style="list-style-type: none"> • Similar to carbamazepine but weaker 	<ul style="list-style-type: none"> • Fewer drug interactions than carbamazepine • Less autoinduction • Lower risk of severe cutaneous reactions








Comparative Efficacy in Bipolar Disorder




AGENT	ACUTE MANIA	ACUTE DEPRESSION	MAINTENANCE	MIXED STATES	RAPID CYCLING
 Lithium (Chokhwal a et al., 2024)	★★★★★☆ • First-line • Slower onset (7-14 days) • Less effective in mixed states	★★★★★☆ • Moderate efficacy • Better for classic bipolar I • May require adjunctive therapy	★★★★★★ • Gold standard • Superior long term outcomes • Unique anti suicide effects	★★★★★☆ • Limited efficacy • Often requires combination therapy	★★★★★☆ • Less effective • Often requires combination therapy
 Valproate (Rahman et al., 2023)	★★★★★☆ • First-line • Faster onset (5-7 days) • Effective for mixed states	★★★★★☆ • Limited evidence • Often requires adjunctive therapy	★★★★★☆ • Effective • Less evidence than lithium • Good for mixed states	★★★★★☆ • First-line for mixed states • Good for agitation	★★★★★☆ • Better than lithium • Often requires combination
 Lamotrigine (Betchel & Saadabadi, 2023)	★☆☆☆☆ • Not effective for acute mania • Not indicated	★★★★★☆ • First-line for bipolar depression • Gradual titration limits acute use	★★★★★☆ • Superior for preventing depression • Modest anti manic prophylaxis	★★★★★☆ • Effective for depressive component • Often used in combination	★★★★★☆ • Effective • Often used with other agents
 Carbamazepine (Maan & Saadabadi, 2023)	★★★★★☆ • Second-line • Effective • More side effects than newer agents	★★★★★☆ • Limited evidence • Not first-line	★★★★★☆ • Effective • Less evidence than lithium • More side effects	★★★★★☆ • Effective • Consider for treatment resistant cases	★★★★★☆ • Effective • Consider for treatment resistant cases
 Oxcarbazepine	★★★★★☆ • Limited evidence • Less effective than carbamazepine	★★★★★☆ • Limited evidence • Not first-line	★★★★★☆ • Limited evidence • Better tolerated than carbamazepine	★★★★★☆ • Limited evidence • Consider if carbamazepine effective but not tolerated	★★★★★☆ • Limited evidence • Consider if carbamazepine effective but not tolerated



Pharmacokinetics and Dosing Considerations

AGENT	HALF-LIFE	THERAPEUTIC LEVELS	DOSING STRATEGY	FORMULATIONS
 Lithium (Chokhawa et al., 2024)	<ul style="list-style-type: none">• 18-24 hours• Increased in elderly• Decreased in pregnancy	<ul style="list-style-type: none">• Acute mania: 0.8-1.2 mEq/L• Maintenance: 0.6-0.8 mEq/L• Elderly: 0.4-0.6 mEq/L	<ul style="list-style-type: none">• Start: 300mg BID/ TID• Titrate based on levels• Check levels 12h post-dose• Steady state: 5-7 days	<ul style="list-style-type: none">• Immediate release• Extended release• Liquid (citrate)
 Valproate (Rahman et al., 2023)	<ul style="list-style-type: none">• 8-16 hours• Shorter with enzyme inducers• Dose-dependent kinetics	<ul style="list-style-type: none">• 50-125 µg/mL• Acute mania: aim for >85 µg/mL• Free fraction varies with albumin	<ul style="list-style-type: none">• Start: 250-500mg BID/TID• Loading: 20-30mg/ kg/day possible• Titrate based on response and levels	<ul style="list-style-type: none">• Immediate release• Extended release (Depakote ER)• Sprinkle capsules• IV formulation
 Lamotrigine (Betchel & Saadabadi, 2023)	<ul style="list-style-type: none">• 25-30 hours• Reduced to 15h with enzyme inducers• Doubled with valproate	<ul style="list-style-type: none">• Not established• Clinical response guides dosing• Monitoring not routinely recommended	<ul style="list-style-type: none">• Start: 25mg daily• Slow titration essential• With valproate: start 25mg every other day• Target: 100-200mg/day	<ul style="list-style-type: none">• Immediate release• Chewable/ dispersible• Extended release (XR)
 Carbamazepine	<ul style="list-style-type: none">• Initial: 25-65 hours• Chronic: 12-17 hours (autoinduction)• Active epoxide metabolite	<ul style="list-style-type: none">• 4-12 µg/mL• Lower end often sufficient• Free fraction varies with albumin	<ul style="list-style-type: none">• Start: 100-200mg BID• Slow titration reduces side effects• Autoinduction requires dose increases• Target: 800-1200mg/day	<ul style="list-style-type: none">• Immediate release• Extended release (Tegretol XR)• Suspension• Chewable tablets
 Oxcarbazepine (Preuss et al., 2021)	<ul style="list-style-type: none">• 8-10 hours• Active metabolite: 9-11 hours	<ul style="list-style-type: none">• Not routinely monitored• MHD metabolite can be measured• Clinical response guides dosing	<ul style="list-style-type: none">• Start: 150-300mg BID• Titrate every 3-7 days• Target: 1200-2400mg/day• Reduce in renal impairment	<ul style="list-style-type: none">• Immediate release• Extended release (Oxtellar XR)• Oral suspension

Laboratory Monitoring and Safety Considerations

AGENT	BASELINE ASSESSMENTS	FOLLOW-UP MONITORING	SPECIAL POPULATIONS	DISCONTINUATION
 Lithium	<ul style="list-style-type: none"> • CBC, electrolytes, BUN/Cr • TSH, calcium • ECG (>40y or cardiac history) • Pregnancy test • Weight/BMI 	<ul style="list-style-type: none"> • Lithium level: 5-7 days after changes, then q3-6mo • Renal/thyroid function: q6-12mo • Calcium/PTH: annually • Weight: each visit 	<ul style="list-style-type: none"> • Pregnancy: ↑ risk of Ebstein's anomaly (0.05-0.1%) • Elderly: ↓ starting dose, ↑ monitoring • Renal impairment: ↓ dose, ↑ monitoring • Avoid NSAIDs, ACEIs, ARBs, diuretics 	<ul style="list-style-type: none"> • Gradual taper (2-4 weeks) • High relapse risk with rapid discontinuation • Monitor closely during taper • Consider cross titration
 Valproate	<ul style="list-style-type: none"> • LFTs, CBC with platelets • Weight/BMI • Pregnancy test • Consider ammonia level 	<ul style="list-style-type: none"> • LFTs: baseline, 1mo, then q3-6mo • CBC: baseline, 1mo, then annually • Weight: each visit • Valproate level: as clinically indicated 	<ul style="list-style-type: none"> • Pregnancy: contraindicated (neural tube defects, autism risk) • Women of childbearing age: use effective contraception • Elderly: ↑ risk of sedation, tremor • Hepatic impairment: ↓ dose, ↑ monitoring 	<ul style="list-style-type: none"> • Gradual taper (2-4 weeks) • Seizure risk with abrupt discontinuation • Monitor for withdrawal symptoms • Consider cross titration
 Lamotrigine (Betchel & Saadabadi, 2023)	<ul style="list-style-type: none"> • CBC, LFTs • Pregnancy test • Detailed skin examination 	<ul style="list-style-type: none"> • No routine lab monitoring required • Monitor closely for rash in first 8 weeks • Reassess if new medications added 	<ul style="list-style-type: none"> • Pregnancy: relatively safe, registry data reassuring • Hepatic impairment: ↓ dose • With valproate: ↓ starting dose and titration rate by 50% • With enzyme inducers: ↑ dose may be needed 	<ul style="list-style-type: none"> • Gradual taper (2-4 weeks) • Seizure risk with abrupt discontinuation • Restart titration if stopped >5 days • Consider cross titration



Carbamazepine
 • CBC, LFTs, electrolytes
 • Pregnancy test •
 Consider HLA B*1502 in Asian patients
 • Consider HLA A*3101 in

q3-6mo
 • Sodium: baseline, 1mo, then q3-6mo •
 Carbamazepine level: 2-4wks after


neural tube defects (0.5-1%) •
 Asian ancestry: screen for HLA B*1502 (SJS/TEN risk)
 • Significant drug •
 Gradual taper (2-4 weeks)
 • Seizure risk with abrupt discontinuation •

Monitor for withdrawal symptoms

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• Pregnancy: ↑ risk of

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AGENT	BASELINE ASSESSMENTS	FOLLOW-UP MONITORING	SPECIAL POPULATIONS	DISCONTINUATION
	European ancestry	changes, then q6-12mo	interactions due to CYP induction • Autoinduction requires dose adjustments	• Consider cross titration
 Oxcarbazepine (Preuss et al., 2021)	• Sodium, BUN/Cr • CBC, LFTs • Pregnancy test • Consider HLA B*1502 in Asian patients	• Sodium: baseline, 1mo, then periodically • No routine drug level monitoring • LFTs: baseline, then as needed	• Pregnancy: limited data, possibly safer than carbamazepine • Asian ancestry: consider HLA B*1502 screening • Fewer drug interactions than carbamazepine • Hyponatremia risk, especially in elderly	• Gradual taper (2-4 weeks) • Seizure risk with abrupt discontinuation • Monitor for withdrawal symptoms • Consider cross titration



Adverse Effects and Management Strategies

AGENT	COMMON ADVERSE EFFECTS	SERIOUS ADVERSE EFFECTS	MANAGEMENT STRATEGIES	CONTRAINDICATIONS

<div> <div></div> <div>Lithium</div> <div>(Chokhwal et al., 2024)</div> </div>	<ul style="list-style-type: none"> • Tremor • Polyuria/polydipsia • Nausea/diarrhea • Weight gain • Cognitive dulling 	<ul style="list-style-type: none"> • Toxicity: ataxia, confusion, seizures • Nephrogenic diabetes insipidus • Hypothyroidism • Hyperparathyroidism • Sinus node dysfunction 	<ul style="list-style-type: none"> • Tremor: β blockers, dose reduction • GI effects: take with food, ER formulation • Polyuria: amiloride, dose reduction • Cognitive: lower dose, divided dosing • Hypothyroidism: levothyroxine 	<ul style="list-style-type: none"> • Significant renal impairment • Severe cardiovascular disease • Severe dehydration • Sodium-depleting diuretics • First trimester pregnancy (relative)
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<div> <div></div> <div>Valproate</div> </div> <ul style="list-style-type: none"> • Sedation • Tremor • Weight gain • Hair loss • GI distress 	<ul style="list-style-type: none"> • Hyperammonemia • Thrombocytopenia • Teratogenicity 	<div>5</div> <div>dosing, ER formulation</div>	<ul style="list-style-type: none"> • Sedation: divided • Tremor: dose reduction, β blockers • Hair loss: biotin, zinc supplementation • Pregnancy/women of childbearing potential without contraception • Hepatic disease 	<ul style="list-style-type: none"> • Urea cycle disorders • Mitochondrial disorders • Pancreatitis
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AGENT	COMMON ADVERSE EFFECTS	SERIOUS ADVERSE EFFECTS	MANAGEMENT STRATEGIES	CONTRAINDICATIONS
			<ul style="list-style-type: none"> • Weight gain: dietary counseling, exercise • GI effects: take with food 	

<div> <div></div> Lamotrigine (Betchel & Saadabadi, 2023) </div>	<ul style="list-style-type: none"> • Headache • Nausea • Dizziness • Diplopia • Insomnia 	<ul style="list-style-type: none"> • Serious rash (SJS/TEN) • Drug reaction with eosinophilia and systemic symptoms (DRESS) • Aseptic meningitis (rare) 	<ul style="list-style-type: none"> • Rash: immediate discontinuation if serious • Headache: divided dosing, analgesics • GI effects: take with food • Insomnia: morning dosing • Proper titration prevents many side effects 	<ul style="list-style-type: none"> • Previous hypersensitivity reaction • Concurrent valproate without dose adjustment • Abrupt discontinuation in epilepsy patients
<div> <div></div> Carbamazepine (Maan & Saadabadi, 2023) </div>	<ul style="list-style-type: none"> • Dizziness • Ataxia • Nausea • Diplopia • Hyponatremia 	<ul style="list-style-type: none"> • Agranulocytosis • Aplastic anemia • SJS/TEN • DRESS • Hepatotoxicity 	<ul style="list-style-type: none"> • Neurological effects: divided dosing, ER formulation • Hyponatremia: fluid restriction, salt supplementation • GI effects: take with food • Proper titration prevents many side effects • HLA testing prevents serious reactions 	<ul style="list-style-type: none"> • Bone marrow suppression • HLA-B*1502 positive (Asian ancestry) • History of serious hematological reaction • MAOIs within 14 days • AV block

Oxcarbazepine

- Dizziness
- Somnolence • Headache • Nausea
- Hyponatremia
- Serious rash (less



common than carbamazepine)


- Hyponatremia (more common than carbamazepine)
- Cross-sensitivity with carbamazepine



- Neurological dosing, ER formulation
- Severe hyponatremia • Hyponatremia: fluid HLA-B*1502 positive restriction, salt (Asian ancestry) supplementation • GI effects: take with food
- Hypersensitivity to oxcarbazepine or carbamazepine

AGENT	COMMON ADVERSE EFFECTS	SERIOUS ADVERSE EFFECTS	MANAGEMENT STRATEGIES	CONTRAINDICATIONS
			<ul style="list-style-type: none"> • Proper titration prevents many side effects 	

Drug Interactions and Combination Strategies

AGENT	SIGNIFICANT INTERACTIONS	EFFECT ON OTHER DRUGS	COMBINATION STRATEGIES	PREGNANCY CONSIDERATIONS
 Lithium (Chokhwal a et al., 2024)	<ul style="list-style-type: none"> • NSAIDs: ↑ lithium levels • Diuretics: ↑ lithium levels • ACEIs/ARBs: ↑ lithium levels • Antipsychotics: ↑ risk of neurotoxicity • Methylodopa: ↑ risk of toxicity 	<ul style="list-style-type: none"> • Minimal effect on other drugs • No significant CYP interactions • No protein binding displacement 	<ul style="list-style-type: none"> • With antipsychotics: monitor for neurotoxicity • With antidepressants: monitor for serotonin syndrome • With anticonvulsants: generally safe combinations 	<ul style="list-style-type: none"> • Category D • First trimester: ↑ risk of Ebstein's anomaly (0.05-0.1%) • Third trimester: "floppy baby syndrome" • Breastfeeding: generally compatible with monitoring • Pregnancy registry available
 Valproate (Rahman et al., 2023)	<ul style="list-style-type: none"> • Lamotrigine: ↑ lamotrigine levels • Carbamazepine: ↓ valproate levels • Aspirin: ↑ free valproate • Warfarin: ↑ bleeding risk • Topiramate: ↑ ammonia levels 	<ul style="list-style-type: none"> • Inhibits multiple UGTs • Weak CYP2C9 inhibitor • Displaces protein-bound drugs 	<ul style="list-style-type: none"> • With lamotrigine: reduce lamotrigine dose by 50% • With antipsychotics: monitor for sedation • With carbamazepine: monitor levels of both 	<ul style="list-style-type: none"> • Category X • Neural tube defects (1-2%) • Autism spectrum disorders • Decreased IQ • Contraindicated in women of childbearing potential unless essential

 Lamotrigine	<ul style="list-style-type: none"> • Valproate: ↑ lamotrigine levels • Carbamazepine: ↓ lamotrigine levels • Oral contraceptives: ↓ lamotrigine levels • Folate: possible ↓ efficacy 	<ul style="list-style-type: none"> • Minimal effect on other drugs • No significant CYP interactions • Weak UGT inducer 	<ul style="list-style-type: none"> • With valproate: reduce lamotrigine dose by 50% • With enzyme inducers: may need ↑ lamotrigine dose • With antidepressants: generally safe combinations 	<ul style="list-style-type: none"> • Category C • Registry data reassuring • No clear association with major malformations • Dose may need to be increased during pregnancy • Breastfeeding generally considered safe
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AGENT	SIGNIFICANT INTERACTIONS	EFFECT ON OTHER DRUGS	COMBINATION STRATEGIES	PREGNANCY CONSIDERATIONS
 Carbamazepine (Maan & Saadabadi, 2023)	<ul style="list-style-type: none"> • Oral contraceptives: ↓ efficacy • Warfarin: ↓ efficacy • Antipsychotics: ↓ levels • Antidepressants: ↓ levels • HIV medications: multiple interactions 	<ul style="list-style-type: none"> • Strong CYP3A4 inducer • Induces CYP1A2, 2B6, 2C9, 2C19 • Induces UGTs • Induces P glycoprotein 	<ul style="list-style-type: none"> • With other anticonvulsants: monitor levels • With antipsychotics: may need ↑ antipsychotic dose • Alternative contraception recommended 	<ul style="list-style-type: none"> • Category D • Neural tube defects (0.5-1%) • Craniofacial defects • Developmental delay • Vitamin K supplementation recommended at birth • Pregnancy registry available
 Oxcarbazepine (Preuss et al., 2021)	<ul style="list-style-type: none"> • Oral contraceptives: ↓ efficacy • Phenytoin: ↑ phenytoin levels • Fewer interactions than carbamazepine 	<ul style="list-style-type: none"> • Moderate CYP3A4/5 inducer • Inhibits CYP2C19 • Less enzyme induction than carbamazepine 	<ul style="list-style-type: none"> • With other anticonvulsants: fewer interactions than carbamazepine • Alternative contraception recommended • Generally better tolerated in combinations 	<ul style="list-style-type: none"> • Category C • Limited data, possibly safer than carbamazepine • Monitor closely if used during pregnancy • Vitamin K supplementation recommended at birth • Pregnancy registry available

Switching and Cross-Titration Protocols

FROM	TO LITHIUM	TO VALPROATE	TO LAMOTRIGINE	TO CARBAMAZEPINE
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Lithium	—	<ul style="list-style-type: none"> • Start valproate 250-500mg BID • Titrate to 1000-1500mg/day over 1-2 weeks • Begin lithium taper when valproate therapeutic • Reduce lithium by 300mg every 3-4 days 	<ul style="list-style-type: none"> • Start lamotrigine 25mg daily • Follow standard titration schedule • Begin lithium taper when lamotrigine reaches 100mg/day • Reduce lithium by 300mg every week 	<ul style="list-style-type: none"> • Start carbamazepine 100-200mg BID • Titrate to 400-800mg/day over 2 weeks • Begin lithium taper when carbamazepine therapeutic • Reduce lithium by 300mg every 3-4 days
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Valproate

- Start lithium 300mg BID
- Titrate based on levels over 1-2 weeks
- Begin valproate

- 25mg every other day
- Follow slow titration schedule (valproate inhibits carbamazepine)
- Start carbamazepine 100-200mg BID
- Titrate to 400-800mg/day over 2 weeks
- Begin valproate

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- Start lamotrigine

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FROM	TO LITHIUM	TO VALPROATE	TO LAMOTRIGINE	TO CARBAMAZEPINE
	taper when lithium therapeutic <ul style="list-style-type: none"> • Reduce valproate by 250mg every 3-4 days 		lamotrigine metabolism) <ul style="list-style-type: none"> • Begin valproate taper when lamotrigine reaches 100mg/day • Reduce valproate by 250mg every week 	taper when carbamazepine therapeutic <ul style="list-style-type: none"> • Reduce valproate by 250mg every 3-4 days

Lamotrigine (Betchel & Saadabadi, 2023)	<ul style="list-style-type: none"> • Start lithium 300mg BID • Titrate based on levels over 1-2 weeks • Begin lamotrigine taper when lithium therapeutic • Reduce lamotrigine by 25-50mg every week 	<ul style="list-style-type: none"> • Start valproate 250-500mg BID • Titrate to 1000-1500mg/day over 1-2 weeks • Begin lamotrigine taper when valproate therapeutic • Reduce lamotrigine by 25-50mg every week • Note: valproate doubles lamotrigine levels 	—	<ul style="list-style-type: none"> • Start carbamazepine 100-200mg BID • Titrate to 400-800mg/day over 2 weeks • Begin lamotrigine taper when carbamazepine therapeutic • Reduce lamotrigine by 25-50mg every week • Note: carbamazepine may reduce lamotrigine levels
Carbamazepine (Maan & Saadabadi, 2023)	<ul style="list-style-type: none"> • Start lithium 300mg BID • Titrate based on levels over 1-2 weeks • Begin carbamazepine taper when lithium therapeutic • Reduce carbamazepine by 200mg every week 	<ul style="list-style-type: none"> • Start valproate 250-500mg BID • Titrate to 1000-1500mg/day over 1-2 weeks • Begin carbamazepine taper when valproate therapeutic • Reduce carbamazepine by 200mg every week 	<ul style="list-style-type: none"> • Start lamotrigine 50mg daily (higher initial dose due to enzyme induction) • Titrate to 200-400mg/day over 6-8 weeks • Begin carbamazepine taper when lamotrigine reaches 200mg/day • Reduce carbamazepine by 200mg every 1-2 weeks 	—



Neuropsychiatric Effects and Cognitive Impact


AGENT	COGNITIVE EFFECTS	NEUROPSYCHIATRIC EFFECTS	LONG-TERM CONSIDERATIONS	PATIENT SELECTION FACTORS

<div></div> Lithium	<ul style="list-style-type: none"> • Mild word-finding difficulties • Subjective cognitive dulling • Mental slowing • Memory effects at higher levels 	<ul style="list-style-type: none"> • Tremor (dose-related) • Rare EPS/movement disorders • Rare encephalopathy at toxic levels • Possible neuroprotective effects 	<ul style="list-style-type: none"> • Stable cognitive profile over time • Possible neuroprotection against dementia • Minimal tolerance development • Possible neurotrophic effects 	<ul style="list-style-type: none"> • Better for "classic" bipolar I • Preferred in patients with suicide risk • Caution in cognitive complaints • Avoid in significant renal disease
<div></div> Valproate (Rahman et al., 2023)	<ul style="list-style-type: none"> • Minimal cognitive effects at therapeutic doses • Dose-related sedation • Rare encephalopathy (with hyperammonemia) 	<ul style="list-style-type: none"> • Tremor (dose-related) • Parkinsonism (rare) • Sedation/somnolence • Rare encephalopathy 	<ul style="list-style-type: none"> • Generally stable cognitive profile • Tolerance to sedation develops • Weight gain may worsen over time • Possible hair thinning over time 	<ul style="list-style-type: none"> • Better for mixed states, rapid cycling • Good for agitated/manic presentations • Caution in women of childbearing potential • Avoid in liver disease
<div></div> Lamotrigine	<ul style="list-style-type: none"> • Minimal cognitive effects • May improve cognitive function in some patients • Rare word-finding difficulties 	<ul style="list-style-type: none"> • Minimal neuropsychiatric effects • Possible activation/insomnia • Rare aseptic meningitis • Possible mood elevation 	<ul style="list-style-type: none"> • Excellent cognitive profile long-term • Minimal tolerance development • Weight neutral • Sustained antidepressant effects 	<ul style="list-style-type: none"> • Better for bipolar depression • Good for patients with cognitive concerns • Preferred in women of childbearing potential • Good for patients with medication sensitivity

<div></div> Carbamazepine	Psychomotor slowing • Moderate cognitive dulling • Memory effects	• Rare paradoxical agitation 10 • Cognitive effects	may persist • Autoinduction may require dose adjustments • Consider for	treatment resistant cases • Good for patients with
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AGENT	COGNITIVE EFFECTS	NEUROPSYCHIATRIC EFFECTS	LONG-TERM CONSIDERATIONS	PATIENT SELECTION FACTORS
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	<ul style="list-style-type: none"> • Word-finding difficulties 		<ul style="list-style-type: none"> • Drug interactions increase over time • Tolerance to sedation develops 	comorbid pain <ul style="list-style-type: none"> • Caution with polypharmacy due to interactions • Avoid in Asian patients without HLA testing
 Oxcarbazepine (Preuss et al., 2021)	<ul style="list-style-type: none"> • Milder cognitive effects than carbamazepine • Less psychomotor slowing • Dose-related cognitive effects 	<ul style="list-style-type: none"> • Dizziness/ataxia • Diplopia • Sedation • Hyponatremia (more common than carbamazepine) 	<ul style="list-style-type: none"> • Better cognitive profile than carbamazepine • Less autoinduction than carbamazepine • Fewer long-term drug interactions • Hyponatremia risk persists 	<ul style="list-style-type: none"> • Alternative when carbamazepine effective but not tolerated • Better tolerated than carbamazepine • Caution in elderly (hyponatremia risk) • Fewer drug interactions than carbamazepine






Clinical Pearls and Practical Considerations

AGENT	UNIQUE ADVANTAGES	CLINICAL PEARLS	COMMON PITFALLS	PATIENT EDUCATION POINTS
 Lithium (Chokhawa et al., 2024)	<ul style="list-style-type: none"> • Only agent with proven anti-suicide effects • Neuroprotective properties • Long history of use and safety data • May prevent dementia 	<ul style="list-style-type: none"> • Therapeutic window narrows with age • Dehydration can precipitate toxicity • Once-daily dosing possible for maintenance • Slow onset (1-2 weeks) for antimanic effects 	<ul style="list-style-type: none"> • Failure to adjust dose with age/renal function • Inadequate monitoring during acute illness • Concurrent medications affecting levels • Inadequate hydration 	<ul style="list-style-type: none"> • Maintain consistent salt/fluid intake • Take with food to reduce GI effects • Report vomiting/diarrhea promptly • Avoid NSAIDs, limit alcohol • Regular monitoring is essential



- Rapid onset of action
- Loading dose
- Failure to warn
- Absolute contraindication in pregnancy
- possible for acute about teratogenic risk
- Valproate
- Multiple mania

AGENT	UNIQUE ADVANTAGES	CLINICAL PEARLS	COMMON PITFALLS	PATIENT EDUCATION POINTS
	formulations available <ul style="list-style-type: none">• Effective for mixed states• Less monitoring than lithium	<ul style="list-style-type: none">• Extended-release reduces side effects• Protein binding varies with dose• Hyperammonemia can occur without LFT elevation	<ul style="list-style-type: none">• Missing rare but serious adverse effects• Inadequate monitoring in combination therapy• Overlooking drug interactions	<ul style="list-style-type: none">• Take with food to reduce GI effects• Report unusual bruising/bleeding• Hair loss often improves with time• Multiple formulations available
 Lamotrigine	<ul style="list-style-type: none">• Excellent tolerability• Weight neutral• Minimal cognitive effects• Effective for bipolar depression	<ul style="list-style-type: none">• Slow titration prevents rash• Restart titration if missed >5 days• Oral contraceptives may reduce levels• Pregnancy may reduce levels	<ul style="list-style-type: none">• Titrating too rapidly• Failure to adjust dose with valproate• Missing early signs of serious rash• Inadequate dosing for maintenance	<ul style="list-style-type: none">• Report any rash immediately• Follow exact titration schedule• May need dose adjustments with other medications• Relatively safe in pregnancy• Minimal weight/cognitive effects

 Carbamazepine (Maan & Saadabadi, 2023)	<ul style="list-style-type: none"> • Effective for treatment-resistant cases • Helpful for comorbid pain • Multiple formulations available • Long history of use 	<ul style="list-style-type: none"> • Autoinduction requires dose adjustments • Extended-release improves tolerability • HLA testing prevents serious reactions • Significant drug interaction potential 	<ul style="list-style-type: none"> • Overlooking serious drug interactions • Failure to adjust for autoinduction • Inadequate monitoring of sodium levels • Missing early signs of blood dyscrasias 	<ul style="list-style-type: none"> • Take with food to reduce GI effects • Report unusual bruising/bleeding • Many drug interactions, including contraceptives • Avoid abrupt discontinuation • Report visual changes/dizziness
 Oxcarbazepine (Preuss et al., 2021)	<ul style="list-style-type: none"> • Better tolerated than carbamazepine • Fewer drug interactions • Less enzyme autoinduction • Less cognitive impact 	<ul style="list-style-type: none"> • Monitor sodium levels, especially in elderly • Extended-release improves tolerability • Cross-reactivity with carbamazepine possible • Less evidence for bipolar disorder 	<ul style="list-style-type: none"> • Overlooking hyponatremia risk • Assuming complete cross efficacy with carbamazepine • Inadequate monitoring in elderly • Overlooking contraceptive interactions 	<ul style="list-style-type: none"> • Monitor for symptoms of low sodium • Take with food to reduce GI effects • May affect hormonal contraceptives • Better tolerated than carbamazepine • Report visual changes/dizziness