Pharmacotherapy for BPSD

- Polypharmacy is **NOT RECOMMENDED** for the management of BPSD.
- For more information on antipsychotic use to manage BPSD, see <u>Considerations for antipsychotic use in BPSD</u>.
- For more information on deprescribing, see <u>Deprescribing antipsychotics</u>.

A note on antipsychotic use:

When prescribing an antipsychotic agent for managing BPSD, prescriptions should be for the **LOWEST** dose that is clinically effective, and for the **SHORTEST** amount of time.

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
	be beneficial in the early stages o	f dementia by addressing anxiety and behavio	 ural symptoms, as cognitive impairment often pre BPSD before symptoms escalate, especially in Lev Initial: 5 mg, once daily (AM or PM). Dose should be maintained for 4-6 weeks before considering a dose increase. Usual: 5-10 mg, once daily. Based on clinical judgement, the 10 mg daily dose may be considered following 4-6 weeks of treatment at 5 mg/day. Max: 10 mg, once daily. In older adult women of low body weight, daily dose should not exceed 5 mg. Administration: Tablets can be crushed or dispersed in 10-20mL of water. Hepatic: Dose + should be individualized in 	(1-month) sents as
			mild to moderate hepatic impairment due to potential increased drug exposure, but there is no data for severe hepatic impairment. Renal: No dose adjustment is required for renal impairment.	



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Rivastigmine (Exelon®) 1.5 mg, 3 mg, 4.5 mg, 6 mg capsule (Exelon® Oral Solution) 2 mg/mL oral solution (Exelon® Patch 5, 10, 15) 4.6 mg/24hr, 9.5 mg/24hr, 13.3 mg/24hr transdermal patch Patches and oral form are not interchangeable on a 1:1 basis due to differences in drug absorption and bioavailability.	Health Canada indication for symptomatic treatment of mild to moderate dementia of the Alzheimer's type. Unlikely to have clinically significant changes to agitation in AD. Best evidence in class for delusions and hallucinations.	 Side effects: CNS: dizziness, tremor, anorexia GI: nausea, vomiting, diarrhea Other: abdominal pain, weight loss of >7% of baseline weight Marnings: Caution in low body weight, + seizure risk, may worsen EPS symptoms and asthma/COPD, may cause bradycardia, QT-prolongation, syncope, + gastric ulcer risk, bladder outflow obstruction. Contraindications: Contraindicated in patients with history of QT prolongation and/or torsade de pointes, including congenital long QT syndromes, history of cardiac arrhythmias. 	 Initial: 1.5 mg BID (3 mg/day). If initial dose is well tolerated, dosage may be increased to 3 mg BID (6 mg/day) after a minimum of 2 weeks. Usual: 6-12 mg/day. Dose increases above 6 mg/day should proceed cautiously. Increases to 4.5 mg BID (9 mg/day) and then 6 mg BID (12 mg/day) should also be based on good tolerability of the current dose and should only be considered after a minimum of 2 weeks at that dose level. Max: 6 mg BID (12 mg/day). Administration: Capsules can be opened and sprinkled; bad taste. Switch from Oral to Patch: < 3 mg BID (6 mg/day) → Exelon® Patch 5 3-6 mg BID (6 mg/day) → Exelon® Patch 10 Apply the first patch on the day following the last oral dose. Patients >85 years old with low body weight or serious comorbid disease should start treatment as less frequent dosing (1.5 mg/day) and increase at a slower rate. Thepatic: Contraindicated in severe hepatic impairment, and dose escalation in mild to moderate impairment should proceed with caution due to potential increased adverse effects. Renal: Limited data available in renally impaired patients, dose escalation should be done cautiously with close monitoring for adverse effects. 	Brand: Cap: \$223 Solution: \$100-362 Patch: \$193- 200 Generic: Cap: \$56 Solution: N/A Patch: \$144 (except Patch 15) ODB: ✓ (LU 347.348) (except solution and patch) NIHB: ✓ (LU) (except solution and patch)



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Galantamine (Reminyl ER™) 8, 16, 24 mg ER capsule	Health Canada indication for symptomatic treatment of mild to moderate dementia of the Alzheimer's type. Unlikely to have clinically significant changes to agitation in AD.	 ♥ Side effects: CNS: Dizziness, syncope, anorexia G: Nausea, vomiting: ▲ Warnings: Caution in low body weight, + seizure risk, may worsen EPS symptoms and asthma/COPD, may cause bradycardia and AV block, QT-prolongation, syncope, + gastric ulcer risk, bladder outflow obstruction. 	 Initial: 4 mg BID (AM/PM, preferably with meals) (8 mg/day) for 4 weeks. Dosage should be increased to initial maintenance dose of 8 mg BID (16 mg/day) after 4 weeks. Usual: 8-12 mg BID (16-24 mg/day). If the initial maintenance dose is well tolerated, a further increase to 12 mg BID (24 mg/day) may be considered only after a minimum of 4 weeks at 8 mg BID (16 mg/day). Max: 12 mg BID (24 mg/day). Dosage escalation for patients >85 years old with low body weight should be undertaken with particular caution. Administration: Capsules can be opened and sprinkled. ✓ Hepatic: Dose + should be done cautiously in mild to moderate impairment (Child-Pugh A or B) with a reduced initial dose of 8 mg every other day, + gradually; contraindicated in severe hepatic impairment (Child-Pugh C) due to increased drug exposure. ✓ Renal: Dose escalation should proceed cautiously in patients with moderate impairment (CrCl 9-60 mL/min), with a maximum recommended dose of 16 mg/day; contraindicated in severe renal impairment (CrCl -9 mL/min) due to lack of data. 	Brand: D/C Generic: \$54 ODB: √ (LU <u>347.348</u>) NIHB: √ (LU)





Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
not tolerated or contraindica	· <u> f or others).</u> risperidone is first-l ted. Second-line options includ	ine for short-term use; aripiprazole or brexpi	prazole may be used as first-line alternatives if trapyramidal symptoms are a concern. Third-lister agents are ineffective or not tolerated. Initial: 2.5 mg/day (PM). Usual: 5-10 mg/day (PM). Max: 12.5 mg/day (PM). Administration: Tablets can be crushed. May be an occupational hazard to person preparing medication and protective measures may be required. Prepatic: No dosage adjustment is required in patients with hepatic impairment. Prepatients with renal impairment.	(1-month) risperidone is

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Brexpiprazole (Rexulti®) 0.25 mg, 0.5 mg, 1 mg, 2 mg, 3 mg, 4 mg, tablets	Health Canada indication for the symptomatic management of agitation associated with dementia of the Alzheimer's type in patients with aggressive behaviours, unresponsive to non-pharmacological approaches.	 Side effects: CNS: Akathisia, myalgia, tremor, dizziness, sedation, restlessness CV: Orthostatic hypotension, increased blood pressure GI: constipation, dyspepsia, nausea, vomiting Metabolic: Weight gain, increased appetite, Other: Muscle stiffness Marnings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk, aspiration pneumonia). Caution in QT prolongation, orthostatic hypotension, bradycardia, syncope, + seizure risk, weight gain, diabetes, dyslipidemia, EPS & tardive dyskinesia, NMS, venous thromboembolism, skin reactions (SJS, DRESS), suicidal ideation, and impulse-control disorders (pathological gambling, compulsive eating/spending/sexual urges). 	 Initial: 0.5 mg/day for one week. Titrate to 1 mg/day for one week. Titrate to usual dose of 2 mg/day. Usual: 2 mg/day. After at least two weeks at 2 mg/day, the dose can be increased to the maximum 3 mg/day if clinically warranted. Max: 3 mg/day. To minimize the risk of adverse events, the lowest effective dose should be used. Administration: Manufacturer recommends swallowing tablets whole. Hepatic: For moderate to severe hepatic impairment (Child-Pugh B or C), the maximum recommended dose for AD is reduced to 2mg/day. Renal: For moderate, severe, or end-stage renal impairment (CrCl <60 mL/min), the maximum recommended dose for AD is reduced to 2 mg/day. 	(1-month) Brand: \$129 Generic: N/A ODB: √ NIHB: √

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Risperidone (Risperdal®) 0.25 mg, 0.5 mg, 1 mg, 2 mg, 3 mg, 4 mg, tablet (Risperidone ODT) 0.5, 1, 2, 3, 4mg orally disintegrating tablet (Risperidone Oral Solution) 1mg/mL	Health Canada indication for the short-term symptomatic management of aggression or psychotic symptoms in patients with severe dementia of the Alzheimer type that is unresponsive to non- pharmacological approaches or other treatments and when there is a risk of harm to self or others. Most likely to cause EPS of antipsychotic agents, especially at higher doses.	 ♥ Side effects: CNS: Somnolence, extrapyramidal symptoms (EPS), agitation, lethargy, falls CV: Orthostatic hypotension, tachycardia GI: Nausea, vomiting, constipation, dry mouth, increased salivation, dyspepsia Metabolic: Weight gain, increased appetite, hyperprolactinemia Other: Peripheral edema, muscle stiffness, urinary tract infection, cough Marnings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk). Caution in QT prolongation, orthostatic hypotension, bradycardia, syncope, + seizure risk, weight gain, diabetes, dyslipidemia, EPS & tardive dyskinesia, neuroleptic malignant syndrome (NMS), skin reactions (SJS, DRESS venous thromboembolism. 	Initial: 0.25 mg BID. Dosage should be adjusted by increments of 0.25 mg per day, approximately every 2 to 4 days. Usual: 0.5 mg BID (1.0 mg/day). Max: 1.0 mg BID (2.0 mg/day). Administration: Tablets can be dispersed in 10- 20mL of water. May be an occupational hazard to person preparing medication and protective measures may be required (<u>change to ODT or</u> oral solution formulations if appropriate). Perpeatic: • plasma concentration of risperidone in hepatic impairment, which may lead to an enhanced pharmacological effect. Lower starting doses and slower titration are recommended. Penal: Reduced clearance in moderate to severe impairment, leading to increased drug exposure. Lower starting doses and slower titration are recommended.	Brand: D/C Generic: Tab: \$16-40 ODT: \$30-80 Solution: \$36- 60 ODB: √ NIHB: √ (except ODT)



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Quetiapine (Seroquel®) 25 mg, 100 mg, 200 mg, 300 mg, immediate-release tablets (Seroquel® XR) 50, 150, 200, 300, 400 mg extended-release tablets	Not indicated by Health Canada for use in older adult patients with dementia. Caution should be used when treating older adult patients. Very sedating. Least likely to cause extrapyramidal side-effects, should be used first in Parkinson's and Lewy body dementia.	 Side effects: CNS: Somnolence, dizziness, extrapyramidal symptoms CY: Orthostatic hypotension, tachycardia, palpitations GI: Dry mouth, constipation, dyspepsia, nausea, vomiting, dysphagia Metabolic: Weight gain, increased appetite, hyperglycemia, hypercholesterolemia, hypertriglyceridemia Other: Withdrawal symptoms, decreased haemoglobin Mathematical and the symptomic of the symptomy of the symptomic of	Initial: 25 mg/day BID. Usual: 100-200 mg/day BID. Max: 300 mg/day BID. Administration: IR tablets can be crushed; bad taste. If taking XR formulation, change to equivalent IR dose and crush. Switching from IR to XR formulations: If treated with divided doses of quetiapine IR, may be switched to XR formulation at the equivalent total daily dose taken once daily. Dosing for frail patients should be reduced by half. Hepatic: + clearance in hepatic impairment; patients with mild impairment should start at 25 mg/day, with slow titration in 25-50 mg increments based on tolerance; use with caution in moderate to severe impairment due to lack of pharmacokinetic data. Renal: Limited clinical data, caution is advised, especially during the initial dosing period.	Brand: IR: \$30-148 XR: \$49-206 Generic: IR: \$15-26 XR: \$21-57 ODB: √ NIHB: √



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Olanzapine (Zyprexa®) 2.5 mg, 5 mg, 7.5 mg, 10 mg, 15 mg, 20 mg, tablets (Zyprexa® Zydis®) 5 mg, 10 mg, 15 mg, 20 mg, orally disintegrating tablets (Zyprexa® Intramuscular) 10 mg, vial	Not indicated by Health Canada for use in older adult patients with dementia. Potential use as short-term emergency treatment of severe agitation. Most likely to cause weight gain and metabolic side effects. Fewer extrapyramidal side effects than risperidone. More sedating than risperidone.	 ♥ Side effects: CNS: Drowsiness, dizziness, restlessness, amnesia, confusion CV: Orthostatic hypotension GI: Increased appetite, constipation, dry mouth, abdominal distention Other: Weight gain, fluid retention, increased salivation, abnormal LFTs, pyrexia, arthralgia, epistaxis ▲ Warnings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk, aspiration pneumonia). Caution in + weight gain, diabetes, dyslipidemia, hyperprolactinemia, QT prolongation, orthostatic hypotension, bradycardia, syncope, + seizure risk, EPS & tardive dyskinesia, neuroleptic malignant syndrome (NMS), venous thromboembolism, skin reactions (SJS, DRESS), and suicidal ideation. 	 Initial: 5 mg/day. Usual: 5-20 mg/day. When indicated, dose escalation should be performed with caution. Max: 20 mg/day. Administration: If crushing is required, switch to the ODT formulation, as the regular tablet can irritate the skin and pose an occupational hazard to the person preparing it. P Hepatic: Olanzapine is associated with aminotransferase (ALT/AST) elevations and rare cases of hepatitis and hepatic failure; thus, caution should be exercised in patients with hepatic impairment, with liver function monitoring recommended. Renal: No major alterations in pharmacokinetics were observed in renal impairment, but caution is advised due to limited clinical experience. 	Brand: Tab: \$89-637 ODT: \$161-643 Vial: \$1195- 2376 Generic: Tab: \$19-60 ODT: \$25-264 Vial: N/A ODB: √ NIHB: √



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Antidepressant Citalopram (Celexa®) 10 mg, 20 mg, 40 mg, tablet Escitalopram (Cipralex®) is not recommended due to negative effects on BPSD symptoms. (expert opinion)	Health Canada indication for the symptomatic relief of depressive illness in adults. Consider for moderate severity agitation.	 Side effects: CNS: Drowsiness, insomnia, dizziness, headache, fatigue, CY: Palpitations, tachycardia, orthostatic hypotension Gi: Nausea, dry mouth, diarrhea, constipation, vomiting, dyspepsia Other: Increased sweating, fatigue, tremor, rhinitis, weight changes, abnormal dreams, sexual dysfunction Varnings: Caution in QT prolongation, suicidal ideation, serotonin syndrome, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, elderly (fall risk, SIADH), poor CYP2C19 metabolizers, and drug interactions (MAOIs, serotonergic drugs, QT-prolonging agents). Contraindicated with monoamine oxidase inhibitors (MAOIs) and pimozide. Contraindicated in patients with known QT interval prolongation or with congenital long QT syndrome. 	 Initial: 10 mg/day. Titrate upwards depending on patient response and tolerability. Usual: 20 mg/day. Max: 30 mg/day. Efficacy may take up to 9 weeks. Administration: Tablets can be crushed or dispersed in 10-20mL of water. ✓ Hepatic: Dose should be + in mild to moderate impairment (Child-Pugh A or B), with a maximum recommended dose of 20 mg/day; use with caution in severe impairment (Child-Pugh C) due to prolonged drug half-life. ✓ Renal: No dosage adjustment is required for mild to moderate impairment, but caution is advised in severe impairment (CrCl <30 mL/min) due to lack of data. 	Brand: \$77-101 Generic: \$17- 28 ODB: √ (except 10mg) NIHB: √



(Tegretol®) use as an anticonvulsant drug. Testo genetics: approached with caution. Tables 200 mg, tablet Teratogenic in nature and not to be handled by women of cle, diploping age. May be anocupational hazard to person preparing medication and protective measures may be required. Test set in the set in t	Medication	Cost and coverage (1-month)
[Tegretol®] use as an anticonvulsant drug. Test sea anticonvulsant drug. Test sea an anticonvulsant drug. Test sea anticonvulsant.	Anticonvulsant/Antimanic	
recent (≤14 days) MAOI use, concurrent itraconazole or voriconazole use.	Carbamazepine (Tegretol®) 100, 200mg chewable tablet 200 mg, tablet (Tegretol® CR) 200 mg, 400 mg, controlled release tablets (Tegretol® Suspension) 100 mg/5 mL, oral use	Brand: Tab: \$23-54 Chew: N/A CR: \$33-55 Susp: \$32-88 Generic: Tab: \$17-28 Chew: \$19-35 CR: \$26-38 Solution: \$23-

Medication	Considerations	Adverse reactions, warnings, and	Dosing	Cost and
		contraindications		coverage (1-month)
Synthetic cannabinoids				(Pinonal)
Nabilone (Cesamet®) 0.25 mg, 0.5 mg, 1 mg capsules	Not indicated by Health Canada for use in older adult patients with dementia. Could be considered for severe agitation unresponsive to other agents.	 Side effects: CNS: Drowsiness, vertigo, psychological high, depression, ataxia, blurred vision, sensation disturbance, headache, euphoria, hallucinations, nightmares, distortion in the perception of time, confusion, dissociation, dysphoria, psychotic reactions, seizures, tremors CV: Orthostatic hypotension, tachycardia, syncope GI: Dry mouth, anorexia Other: Asthenia Warnings: May impair mental and physical abilities, effects can persist 48–72 hours after stopping, may cause hallucinations, euphoria, confusion, dissociation, depression, psychosis, and emotional ability. Contraindications: Known sensitivity to cannabis or other cannabinoid agents or a history of psychotic reactions. 	Initial: 0.5 mg once daily (PM). Usual: 0.5 mg BID or 1 mg BID. Max: 4 mg/day (mean therapeutic dose in studies ~1.6 mg/day in elderly patients). Hepatic: Use with extreme caution in patients with severe liver dysfunction due to risk of prolonged effects or accumulation. Renal: No specific renal dosing guidance is provided but use with caution in renal impairment.	Brand: \$162-1116 Generic: \$110- 497 ODB: √ (except 0.25 mg) NIHB: √



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Typical antipsychotic agents Haloperidol (Haldol®) 0.5 mg, 1 mg, 2 mg, 5 mg, 10 mg tablets 5 mg/mL IM injection	Not indicated by Health Canada for use in older adult patients with dementia. Low-dose, short-term use may be considered under close monitoring. Avoid in Lewy Body Dementia or Parkinson's Disease Dementia due to high EPS risk. Generally, should be avoided in elderly due to + risk of mortality as the dose is increased compared to atypical antipsychotics: + risk of EPS and neuroleptic malignant syndrome, + risk of anticholinergic effects, + sedation, minimal weight gain.	 Side effects: CNS: High risk of EPS, tardive dyskinesia, lethargy, and neuroleptic malignant syndrome; may also lower seizure threshold CV: Risk of QT prolongation, torsade de pointes, and sudden death GI: Nausea, vomiting, diarrhea, heartburn, dry mouth, decreased appetite, dehydration Metabolic: Weight changes, hyperprolactinemia Other: Urinary incontinence, sweating, skin changes, nasal congestion Marnings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk, aspiration pneumonia). Contraindications: Intravenous administration, comatose states, CNS depression, Parkinson's syndrome (except for levodopa-induced dyskinesias). 	 Initial: 0.25-0.5 mg PO once or twice daily 0.5-1 mg IM as needed every hour until desired result achieved (do not exceed max dose). Usual: Lowest effective dose, shortest duration. Max: 2 mg/day in elderly patients. Hepatic: Use cautiously in patients with liver dysfunction due to risk of hepatotoxicity and prolonged drug clearance. Renal: Use cautiously in severe renal dysfunction; monitor closely. 	Brand: N/A Generic: Tab: \$18-24 IM: \$28-76 ODB: √ NIHB: √

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Benzodiazepine Lorazepam (Ativan®) 0.5 mg, 1 mg, 2 mg, oral tablets (Ativan® SL) 0.5 mg, 1 mg, 2 mg, sublingual tablets Lorazepam Intramuscular 4 mg/mL	Health Canada indication for short-term relief of manifestations of excessive anxiety in patients with anxiety neurosis. Indicated as an adjunct for the relief of excessive anxiety. Potential use as short-term emergency treatment of severe agitation.	 Side effects: CNS: Drowsiness, confusion, dizziness, fatigue, amnesia, sedation, ataxia, depression, tremor, headache, visual disturbances (diplopia, blurred vision) GI: Nausea, constipation, change in appetite Other: Respiratory depression, worsening COPD, apnea, muscle weakness, hypersensitivity reactions, SIADH, hypothermia Warnings: Caution in CNS depression, addiction, abuse, and dependence, withdrawal risk (seizures, delirium, psychosis, rebound anxiety/insomnia, autonomic dysfunction), paradoxical reactions (aggression, agitation, hallucinations, behavioral changes), • fall and fracture risk in elderly, cognitive impairment, myasthenia gravis exacerbation, respiratory depression, suicidal ideation, and concomitant CNS depressant use (opioids, alcohol, sedatives, antidepressants, antipsychotics, anticonvulsants, muscle relaxants, anesthetics). Contraindications: Myasthenia gravis, acute narrow-angle glaucoma, severe respiratory insufficiency, sleep apnea syndrome, concurrent use with opioids or other CNS depressants in high-risk patients. 	 Initial: 0.5 mg/day. Dosage should be very carefully and gradually adjusted by 0.5 mg, depending upon tolerance and response. Usual: 2-3 mg/day, taken in divided doses. Max: 6 mg/day, taken in divided doses. Administration: Tablets can be dispersed in 10-20mL of water (<i>change to SL if appropriate</i>). Hepatic: Use lowest effective dose in mild to moderate hepatic impairment; caution in severe hepatic impairment due to risk of hepatic encephalopathy. Renal: Use lowest effective dose in mild to moderate renal impairment; effect may be prolonged. 	Brand: Tab: \$15-22 SL: \$18-41 Generic: Tab: \$15-22 SL: \$17-35 IM: \$713 ODB: √ (except SL) NIHB: √

Medication	Considerations	Adverse reactions, warnings, and	Dosing	Cost and
		contraindications		coverage
Psychosis (moderate-severe)		e for <u>moderate symptoms</u> , while risperidone or ar 3 Side effects: CNS : Drowsiness, insomnia, dizziness, headache, fatigue, CV : Palpitations, tachycardia, orthostatic hypotension GI: Nausea, dry mouth, diarrhea, constipation, vomiting, dyspepsia Other : Increased sweating, fatigue, tremor,	 ipiprazole are second-line choices for more severe or Initial: 10 mg/day. Titrate upwards depending on patient response and tolerability. Usual: 20 mg/day. Max: 30 mg/day. Administration: Tablets can be crushed or 	coverage treatment- Brand: \$77-101 Generic: \$17-28 ODB: √ (except 10mg)
		rhinitis, weight changes, abnormal dreams, sexual dysfunction ▲ Warnings: Caution in QT prolongation, suicidal ideation, serotonin syndrome, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, elderly (fall risk, SIADH), poor CYP2CI9 metabolizers, and drug interactions (MAOIs, serotonergic drugs, QT-prolonging agents). Contraindicated with MAOIs and pimozide. Contraindicated in patients with known QT interval prolongation or with congenital long QT syndrome.	 dispersed in 10-20mL of water. Efficacy may take up to 9 weeks. Phepatic: Dose should be ↓ in mild to moderate impairment (Child-Pugh A or B), with a maximum recommended dose of 20 mg/day; use with caution in severe impairment (Child-Pugh C) due to prolonged drug half-life. Renal: No dosage adjustment is required for mild to moderate impairment, but caution is advised in severe impairment (CrCl <30 mL/min) due to lack of data. 	NIHB: √



Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Atypical antipsychotic agents Aripiprazole (Abilify®) 2 mg, 5 mg, 10 mg, 15 mg, 20 mg, 25 mg, 30 mg, tablets	Not indicated by Health Canada for use in older adult patients with dementia. Caution should be used when treating older adult patients. Consider for severe symptoms or if citalopram is ineffective or not tolerated.	 Side effects: CNS: Akathisia, sedation, restlessness, extrapyramidal disorder, fatigue, blurred vision CV: Hypertension, hypotension, syncope GI: Nausea, constipation, dyspepsia, vomiting, stomach discomfort, GERD, dysphagia, dry mouth Other: Weight loss, toothache, hyperglycemia, elevated LFTs, musculoskeletal stiffness, dyspnea, hyperhidrosis Manings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk, aspiration pneumonia). Caution in QT prolongation, orthostatic hypotension, bradycardia, syncope, + seizure risk, weight gain, diabetes, dyslipidemia, EPS & tardive dyskinesia, NMS, venous thromboembolism, skin reactions (SJS, DRESS), suicidal ideation, and impulse-control disorders (pathological gambling, compulsive eating/spending/sexual urges). 	 Initial: 2.5 mg/day (PM). Usual: 5-10 mg/day (PM). Max: 12.5 mg/day (PM). Administration: Tablets can be crushed. May be an occupational hazard to person preparing medication and protective measures may be required. P Hepatic: No dosage adjustment is required in patients with hepatic impairment. Renal: No dosage adjustment is required in patients with renal impairment. 	Brand: \$122- 185 Ceneric: \$40- 55 ODB: √ NIHB: √

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Risperidone (Risperdal®) 0.25 mg, 0.5 mg, 1 mg, 2 mg, 3 mg, 4 mg, tablet (Risperidone ODT) 0.5, 1, 2, 3, 4 mg orally disintegrating tablet (Risperidone Oral Solution) 1 mg/mL	Health Canada indication for the short-term symptomatic management of aggression or psychotic symptoms in patients with severe dementia of the Alzheimer type that is unresponsive to non- pharmacological approaches or other treatments and when there is a risk of harm to self or others. Consider for severe symptoms or if citalopram is ineffective or not tolerated. Most likely to cause EPS of antipsychotic agents, especially at higher doses.	 Side effects: CNS: Somnolence, extrapyramidal symptoms (EPS), agitation, lethargy, falls CY: Orthostatic hypotension, tachycardia GI: Nausea, vomiting, constipation, dry mouth, increased salivation, dyspepsia Metabolic: Weight gain, increased appetite, hyperprolactinemia Other: Peripheral edema, muscle stiffness, urinary tract infection, cough Marnings: Older adult patients with dementia treated with antipsychotic drugs are at an increased risk of death (+ stroke risk). Caution in QT prolongation, orthostatic hypotension, bradycardia, syncope, + seizure risk, weight gain, diabetes, dyslipidemia, EPS & tardive dyskinesia, neuroleptic malignant syndrome (NMS), skin reactions (SJS, DRESS venous thromboembolism. 	 Initial: 0.25 mg BID. Dosage should be adjusted by increments of 0.25 mg per day, approximately every 2 to 4 days. Usual: 0.5 mg BID (1.0 mg/day). Max: 1.0 mg BID (2.0 mg/day). Administration: Tablets can be dispersed in 10-20mL of water. May be an occupational hazard to person preparing medication and protective measures may be required (<u>change to ODT or oral solution formulations if appropriate</u>). P Hepatic: + plasma concentration of risperidone in hepatic impairment, which may lead to an enhanced pharmacological effect. Lower starting doses and slower titration are recommended. Renal: Reduced clearance in moderate to severe impairment, leading to increased drug exposure. Lower starting doses and slower titration are retimpairment, and slower titration are recommended. 	Brand: D/C Generic: Tab: \$16-40 ODT: \$30-80 Solution: \$36- 60 ODB: √ NIHB: √ (except ODT)

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Depression (moderate-to-sev	vere)			
Pharmacological treatment fo	or depressive symptoms in deme	ntia should <u>only</u> be considered if symptoms are u	unresponsive to psychosocial interventions.	
profiles and coexisting sympt Antidepressants Sertraline	Health Canada indication for	Ø Side effects:	ence in older adults, with the choice guided by sid Initial: 50 mg/day.	Brand: \$46-
(Zoloft®) 25 mg, 50 mg, 100 mg, capsules	symptomatic relief of depressive illness.	 CNS: Insomnia, somnolence, tremor, dizziness G: Nausea, loose stools/diarrhea, dyspepsia Other: Male sexual dysfunction, dry mouth, increased sweating ✓ Warnings: Caution in QT prolongation, syncope, serotonin syndrome, suicidal ideation, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, bone fracture risk, discontinuation symptoms, angle-closure glaucoma, cognitive and motor impairment, impulse-control disorders (gambling, compulsive behavior), diabetes (loss of glycemic control, new-onset diabetes risk). ✓ Contraindications: Contraindicated with MAOIs and pimozide. 	 Usual: 50-200 mg/day. A gradual increase in dosage may be considered if no clinical improvement is observed. Dosage changes should be made at intervals of at least one week. Max: 200 mg/day. Administration: Capsules can be opened and sprinkled or opened and dispersed in 10-20mL of water. Pepatic: Use with caution in mild hepatic impairment; dose reduction or less frequent dosing should be considered due to prolonged half-life. Renal: No specific renal dose adjustment required but use with caution in severe impairment. 	155 Generic: \$18- 33 ODB: √ NIHB: √

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Duloxetine 30 mg, 60 mg, delayed-release capsules	Health Canada indication for symptomatic relief of major depressive disorder (MDD).	 Side effects: CNS: Fatigue, somnolence, dizziness, tremor, anxiety, insomnia, blurred vision G: Dry mouth, nausea, constipation, diarrhea, vomiting Other: Decreased appetite, decreased libido, anorgasmia, erectile dysfunction, hot flushes, increased sweating Contraindications: Concurrent or recent (\$14 days) MAOI use, concomitant use with thioridazine, uncontrolled narrow-angle glaucoma, concurrent use with potent CYP1A2 inhibitors (e.g., fluvoxamine, ciprofloxacin). 	Initial: 60 mg/day. A starting lower dose of 30 mg/day may be considered for tolerability reasons in some patients. If starting at the lower dose, target dose of 60 mg/day should be achieved within 1-2 weeks. Usual: 60 mg/day. Max: 60 mg/day. Administration: Capsules can be opened and sprinkled to apple juice, apple sauce, but <u>NOT</u> chocolate pudding (pH affects enteric coating). Hepatic: Contraindicated in any hepatic impairment; 5-fold increase in drug exposure in moderate liver impairment (Child-Pugh Class B). Renal: Not recommended in severe renal impairment (CrCl <30 mL/min) or end-stage renal disease (ESRD) due to 2-fold increase in drug exposure.	Brand: \$99- 187 Generic: \$25- 37 ODB: √ NIHB: √

Medication	Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
Citalopram (Celexa®) 10 mg, 20 mg, 40 mg, tablet Escitalopram (Cipralex®) is not recommended due to negative effects on BPSD symptoms.	Health Canada indication for the symptomatic relief of depressive illness in adults. May be used if comorbid anxiety and/or agitation present.	 Side effects: CNS: Drowsiness, insomnia, dizziness, headache, fatigue, CV: Palpitations, tachycardia, orthostatic hypotension GI: Nausea, dry mouth, diarrhea, constipation, vomiting, dyspepsia Other: Increased sweating, fatigue, tremor, rhinitis, weight changes, abnormal dreams, sexual dysfunction Warnings: Caution in QT prolongation, suicidal ideation, serotonin syndrome, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, elderly (fall risk, SIADH), poor CYP2C19 metabolizers, and drug interactions (MAOIs, serotonergic drugs, QT-prolonging agents). Contraindicated with MAOI and pimozide. Contraindicated in patients with known QT interval prolongation or with congenital long QT syndrome. 	 Initial: 10 mg/day. Titrate upwards depending on patient response and tolerability. Usual: 20 mg/day. Max: 30 mg/day. Administration: Tablets can be crushed or dispersed in 10-20mL of water. Efficacy may take up to 9 weeks. Hepatic: Dose should be + in mild to moderate impairment (Child-Pugh A or B), with a maximum recommended dose of 20 mg/day; use with caution in severe impairment (Child-Pugh C) due to prolonged drug half-life. Renal: No dosage adjustment is required for mild to moderate impairment, but caution is advised in severe impairment (CrCl <30 mL/min) due to lack of data. 	Brand: \$77-101 Ceneric: \$17- 28 ODB: √ (except 10mg) NIHB: √



Considerations	Adverse reactions, warnings, and contraindications	Dosing	Cost and coverage (1-month)
sidered first-line for moderate t	to severe symptoms.		
Health Canada indication for the symptomatic relief of depressive illness in adults.	 Side effects: CNS: Drowsiness, insomnia, dizziness, headache, fatigue, CY: Palpitations, tachycardia, orthostatic hypotension GI: Nausea, dry mouth, diarrhea, constipation, vomiting, dyspepsia Other: Increased sweating, fatigue, tremor, rhinitis, weight changes, abnormal dreams, sexual dysfunction Warnings: Caution in QT prolongation, suicidal ideation, serotonin syndrome, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, elderly (fall risk, SIADH), poor CYP2C19 metabolizers, and drug interactions (MAOIs, serotonergic drugs, QT-prolonging agents). Contraindications: Contraindicated with MAOIs and pimozide. 	 Initial: 10 mg/day. Titrate upwards depending on patient response and tolerability. Usual: 20 mg/day. Max: 30 mg/day. Administration: Tablets can be crushed or dispersed in 10-20mL of water. Efficacy may take up to 9 weeks. P Hepatic: Dose should be ↓ in mild to moderate impairment, with a maximum recommended dose of 20 mg/day; use with caution in severe hepatic impairment due to prolonged drug half-life. Renal: No dosage adjustment is required for mild to moderate impairment, but caution is advised in severe impairment (CrCl <30 mL/min) due to lack of data. 	Brand: \$77-10 Generic: \$17- 28 ODB: √ (except 10mg NIHB: √
	Health Canada indication for the symptomatic relief of	idered first-line for moderate to severe symptoms. Health Canada indication for the symptomatic relief of depressive illness in adults.	 idered first-line for moderate to severe symptoms. Health Canada indication for the symptomatic relief of depressive illness in adults. Side effects: CNS: Drowsiness, insomnia, dizziness, headache, fatigue, CV: Palpitations, tachycardia, orthostatic hypotension GI: Nausea, dry mouth, diarrhea, constipation, vomiting, dyspepsia Other: Increased sweating, fatigue, tremor, rhinitis, weight changes, abnormal dreams, sexual dysfunction Marings: Caution in QT prolongation, suicidal ideation, serotonin syndrome, abnormal bleeding, hyponatremia, seizure risk, mania/hypomania activation, elderly (fall risk, SIADH), poor CYP2C19 metabolizers, and drug interactions (MAOIs, serotonergic drugs, QT-prolonging agents). Contraindicated with MAOIs and pimozide. Contraindicated in patients with known QT interval prolongation or with congenital long

Legend

AD = Alzheimer's disease; BID = twice a day; BP = Blood Pressure; CNS = Central Nervous System; CR = Complete response; CrCl = Creatinine Clearance; CV = Cardiovascular; D/C = Discontinued; DRESS = Drug reaction with eosinophilia and systemic symptoms; EPS = Extrapyramidal symptoms (e.g., dystonia, akathisia, parkinsonism, tardive dyskinesia); GERD = Gastroesophageal Reflux Disease; GI = Gastrointestinal; HR = Heart Rate; IM = Intramuscular; IR= Immediate-release; LFT = Liver function test; MAOI = Monoamine oxidase inhibitor; N/A = Not Applicable; NMDA = Nmethyl-D-aspartate receptor; NMS = Neuroleptic malignant syndrome; ODT = Orally disintegrating tablets; PO = by mouth; SIADH = Syndrome of Inappropriate Antidiuretic Hormone Secretion; SJS = Stevens-Johnson Syndrome; SL= sublingual; TEN = Toxic Epidermal Necrolysis; XR = Extended-release

Cost: an approximate range for a 1-month supply using the initial and max dose (includes a markup of 10% and a dispensing fee of \$12.99)



21