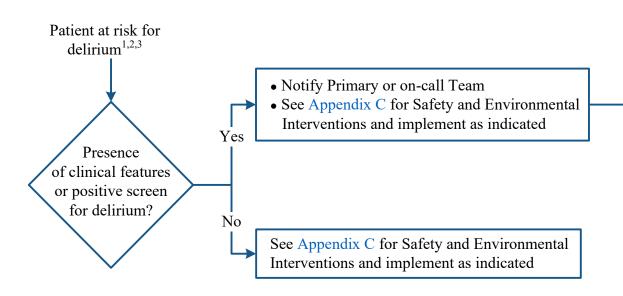
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Note: This algorithm is not intended for patients with alcohol withdrawal related delirium.

### INITIAL PRESENTATION/ASSESSMENT



### **CLINICAL EVALUATION**

- History and Physical and chart review
  - o Confirm history with family/caregivers
  - o Physical examination with attention to neurological status
  - o Review current and home medications
    - Confirm home medication use with family/caregivers
    - Consider drug overdose versus withdrawal, serotonin syndrome and/or neuroleptic malignant syndrome
    - Review for correct dosing based on age and clinical condition
  - Avoid abrupt discontinuation of medications with potential for dependence and/or withdrawal syndrome
  - Consider ongoing need for medications that may contribute to delirium (see Appendix B)
  - Review history for alcohol and substance use/misuse
  - o Clinical interview and mental status exam
  - o Consider evaluation using standardized tools (CAM and/or MDAS)
- Consider the following as clinically indicated:
  - CBC with differential, basic metabolic panel with calcium, liver function tests, oxygen saturation/arterial blood gas, troponin T, albumin, thyroid function tests, ammonia, cortisol
  - o Urinalysis, urine culture, blood cultures, cerebral spinal fluid studies
  - o Serum/urine drug screen
  - o Chest x-ray and EKG
  - o EEG, CT head, MRI brain
- Consultations as appropriate
- Treat acute severe causes such as pain, sepsis, hypoxia, electrolyte disturbances, and medication toxicities

→ See Page 2

<sup>&</sup>lt;sup>1</sup> See Appendix A for clinical features of delirium

<sup>&</sup>lt;sup>2</sup> See Appendix B for risk factors and contributing factors

<sup>&</sup>lt;sup>3</sup> Routine screening in the Critical Care Unit performed with the Intensive Care Delirium Screening Checklist (ICDSC) and screening for Supportive Care patients performed with the Memorial Delirium Assessment Scale (MDAS)



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Note: This algorithm is not intended for patients with alcohol withdrawal related delirium. **EVALUATION AND INTERVENTIONS** INTERVENTIONS • Correct contributing factors Continue to assess and monitor as appropriate (see Appendix B) Yes • Continue with safety and • Continue interventions and • Consider medications as environmental interventions monitor as appropriate ⊢ Hypoactive<sup>2,3</sup> → Response<sup>5</sup>? appropriate for short-term (see Appendix C) • Reduce pharmacologic management of severe • Monitor airway, breathing, and treatment as indicated? No agitation and/or patient risk of aspiration Response<sup>5</sup>? discomfort<sup>6</sup> Patient with • Consider specialty consultation<sup>4</sup> confirmed (see Appendix D and E) No diagnosis of • Consider specialty Consider specialty consultation<sup>4</sup> delirium<sup>1,2</sup> consultation<sup>4</sup> Potential medical emergency • Ensure safety for patient, family, and staff • Continue interventions and monitor as appropriate • Correct contributing factors • Reduce pharmacologic treatment as indicated<sup>7</sup> (see Appendix B) Yes • Continue with safety and Hyperactive<sup>2,8</sup> Response<sup>5</sup>? environmental interventions or Mixed<sup>2,9</sup> (see Appendix C) No • Consider medications as appropriate Consider specialty consultation for short-term management of severe agitation and/or patient discomfort<sup>6</sup> (see Appendix D and E) • Consider specialty consultation<sup>4</sup>

<sup>&</sup>lt;sup>1</sup> Consider Social Work consult to determine Legal Next of Kin and/or Medical Power of Attorney status

<sup>&</sup>lt;sup>2</sup> Follow algorithm based on delirium type at time of evaluation

<sup>&</sup>lt;sup>3</sup> Hypoactive clinical features include withdrawal, flat affect, lethargy, and/or diminished responsiveness

<sup>&</sup>lt;sup>4</sup>Consider specialty consultation with Pharmacy, Psychiatry, Neurology, Supportive Care and/or Anesthesiology as indicated

<sup>&</sup>lt;sup>5</sup>Response to interventions should be based on continuous evaluation over a period of time and not on a single evaluation

<sup>&</sup>lt;sup>6</sup> Specialty specific management of delirium may include dexmedetomidine (ICU setting), combination of haloperidol and lorazepam (palliative care setting or patients with severe agitation) or combination of other psychotropics as deemed appropriate by consultants

<sup>&</sup>lt;sup>7</sup>Chronic use of antipsychotic therapy may not be indicated in the absence of underlying psychiatric conditions (*e.g.*, schizophrenia)

<sup>&</sup>lt;sup>8</sup> Hyperactive clinical features include hallucinations, agitation, restlessness, combativeness, pulling at catheters and/or tubes

<sup>&</sup>lt;sup>9</sup> Mixed clinical features include fluctuations between hyperactive and hypoactive delirium



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## **APPENDIX A: Clinical Features of Delirium**

- Acute onset
- Confusion, disorientation, impaired reality testing
- Inability to pay attention (distractibility)
- Psychomotor agitation or retardation
- Illusions (misperceptions) and hallucinations (usually visual)
- Diurnal variation (worse at night, early AM)

- Sleep-wake cycle disruption
- Fluctuating course, lucid intervals
- Autonomic dysfunction
- Fear and anxiety
- Delusions, especially with paranoid themes



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## **APPENDIX B: Risk Factors and Contributing Factors for Delirium**

ciprofloxacin), and voriconazole

<b>Patient Characteristics</b>	<ul><li>Age &gt; 64 years</li><li>Sensory impairment (visual and/or hearing)</li></ul>	Pain Management	Unrelieved pain
Metabolic Disturbance	<ul> <li>Hypoxia</li> <li>Hypercapnia</li> <li>Hypo or Hyperglycemia</li> <li>Hypo or Hypernatremia</li> <li>Hypercalcemia</li> <li>Impaired liver function and/or kidney function</li> <li>Thyroid disorders</li> </ul>	Cancer Therapies	<ul> <li>Chemotherapy agents (e.g., ifosfamide, methotrexate, cytosine arabinoside)</li> <li>Biotherapy agents [e.g., interleukin-2 (IL-2), interferon-alpha, blinatumomab]</li> <li>Chimeric antigen receptor (CAR) T-cell therapy</li> <li>Supportive therapy agents (e.g., opioids, benzodiazepines, corticosteroids)</li> </ul>
Drugs <sup>1</sup>	<ul> <li>Polypharmacy</li> <li>Medications with anticholinergic effects<sup>2,3</sup> (e.g., scopolamine, promethazine, prochlorperazine, diphenhydramine, hydroxyzine, oxybutynin, hycoscyamine, tricyclic antidepressants)</li> <li>Opioids</li> <li>Benzodiazepines</li> <li>Zolpidem, eszopiclone, zaleplon</li> <li>Cyclobenzaprine, baclofen</li> <li>Anticonvulsants (e.g., phenytoin, phenobarbital, levetiracetam)</li> <li>Corticosteroids (e.g., methylprednisolone, prednisone)</li> <li>Histamine-type 2 receptor antagonist (e.g., famotidine)</li> <li>Digoxin (particularly with elevated blood levels)</li> <li>Anti-Parkinson agents</li> </ul>	Disease/condition Related	<ul> <li>History of cognitive impairment including dementia</li> <li>Direct and indirect effects of primary brain tumors</li> <li>Central nervous system conditions (e.g., metastasis, stroke, seizures)</li> <li>Paraneoplastic syndromes (rarely)</li> <li>Terminal stages of disease/end of life</li> <li>Alcohol or drug (e.g., opioids, benzodiazepines) intoxication or withdrawal</li> <li>History of alcohol or substance misuse</li> <li>Hypertensive crisis</li> <li>Posterior reversible encephalopathy syndrome (PRES)</li> <li>Urinary retention and/or fecal impaction</li> <li>Depression</li> <li>Frailty</li> <li>Infection</li> </ul>
	<ul> <li>Anticholinergics<sup>3</sup> (e.g., cogentin)</li> <li>Adjunctive agents (e.g., amantadine, selegiline)</li> <li>Dopamine agonists (e.g., bromocriptine, ropinirole)</li> <li>Carbidopa/levodopa</li> <li>Sympathomimetics (e.g., methylphenidate, amphetamine, dextroamphetamine)</li> </ul>	Other	<ul> <li>Use of restraints</li> <li>Use of indwelling urinary catheters</li> <li>Recent discharge from acute hospital</li> <li>Patient with recent history or undergoing anesthesia/surgery</li> <li>Immobility</li> <li>Lack of sleep</li> </ul>
	• Sympathomimetics (e.g., methylphenidate, amphetamine,	<sup>1</sup> Consider Pharmacy consult for me	anesthesia/surgery • Immobility • Lack of sleep



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## **APPENDIX C: Safety and Environmental Interventions**

Category	Interventions				
Prevent accidental self harm	<ul> <li>Implement Comprehensive Managed Fall Protection Program as per policy</li> <li>Implement strategies to prevent self removal of lines, tubes, and drains. See interventions for close observation and physical environment.</li> <li>Avoid catheterizations</li> <li>Remove lines, tubes, and drains as soon as indicated</li> <li>Physical restraints if other measures are unsuccessful</li> </ul>				
Close observation	• Nurse • Sitter				
Physical agitation and physiological instability	Reassess for consideration of transfer to next level of care				
Physical environment	<ul> <li>Adequate, but not excessive, sensory stimulation</li> <li>Sleep promotion strategies <ul> <li>Minimize disruption of sleep-wake cycle</li> <li>Avoid long periods of daytime sleep</li> </ul> </li> <li>Lights on during day <ul> <li>Maximize mobility</li> </ul> </li> <li>Frequent reorientation (use of clocks, calendars, and updates on whiteboard)</li> <li>Address sensory deficits (e.g., eyeglasses, other vision aids such as magnifiers and special lighting, hearing aids, amplifying devices)</li> <li>Address language barriers as indicated through the use of Language Assistance program and provision of language specific patient education materials</li> <li>Night: low level background light and sound (music or television) maintained</li> <li>Family presence</li> </ul>				
Provide reassurance and education to patient and caregivers	Communicate and educate about delirium and delirium management     Encourage family members to take breaks				



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## APPENDIX D: Medications for Management of Delirium For All Inpatient Care Areas

Note: Oral formulations should be avoided in patients who cannot safely swallow or who are at risk for aspiration

Therapeutic Class	Medication	Typical Initial Dose	Recommended Maximum Dose	Onset of Action	Comments/Cautions/Adverse Reactions
Typical Antipsychotics	Haloperidol (Haldol <sup>®</sup> )	<ul> <li>IV: Age ≥ 65 years: 0.5-2 mg every 6 hours PRN Age &lt; 65 years: 2-5 mg every 6 hours PRN</li> <li>PO: Age ≥ 65 years: 0.5-2 mg every 12 hours PRN Age &lt; 65 years: 2-5 mg every 12 hours PRN</li> <li>Loading regimen for hyperactive delirium: Age ≥ 65 years: 0.5 mg IV Age &lt; 65 years: 2 mg IV</li> <li>Repeat dose every 20-30 minutes until patient is calm, then schedule 25% of total loading dose IV every 6 hours</li> </ul>	IV: 30 mg/day PO: 30 mg/day		<ul> <li>Likely of greatest utility in acute management of hyperactive delirium (<i>i.e.</i>, establishing initial control and PRN for breakthrough agitation)</li> <li>QTc prolongation (dose dependent)/risk of torsades de pointes: <ul> <li>Obtain 12-lead EKG at baseline and consider repeating every 48-72 hours</li> <li>Caution with QTc &gt; 450 ms or increase by 25% or more from baseline</li> <li>Not recommended if QTc &gt; 500 ms</li> </ul> </li> <li>Extrapyramidal reactions (acute dystonia, akathisia, parkinsonism, tardive dyskinesia) – higher incidence relative to atypical antipsychotics</li> <li>Hypotension, particularly with IV administration</li> <li>Neuroleptic malignant syndrome has been reported with antipsychotic administration (manifests as hyperpyrexia, muscle rigidity, autonomic instability)</li> <li>May lower seizure threshold</li> </ul>
Atypical Antipsychotics	Quetiapine (Seroquel®)	PO: 25-50 mg every 12 hours Hepatic impairment: 12.5 mg every 12 hours Age > 60 years: 12.5-25 mg every 12 hours	400 mg/day	1.5 hours	<ul> <li>Likely of greatest benefit as maintenance therapy for hyperactive/mixed delirium; can be considered for hypoactive delirium unresponsive to non-pharmacologic management</li> <li>May cause hyperglycemia; cases of diabetic ketoacidosis and hyperosmolar coma have been reported</li> <li>Orthostatic hypotension, especially upon initiation and titration of therapy</li> <li>QTc prolongation (dose dependent)/risk of torsades de pointes;</li> <li>Obtain 12-lead EKG at baseline and consider repeating every 48-72 hours</li> <li>Caution with QTc &gt; 450 ms or increases by 25% or more from baseline</li> <li>Not recommended if QTc &gt; 500 ms</li> <li>Neuroleptic malignant syndrome has been reported with antipsychotic administration (manifests as hyperpyrexia, muscle rigidity, autonomic instability)</li> <li>May lower seizure threshold</li> <li>Extrapyramidal reactions may occur, but are less common than with typical antipsychotics</li> <li>Metabolized by CYP450 enzyme system; caution with concomitant use of CYP450 inhibitors and inducers</li> <li>IM administration contraindicated in patients with thrombocytopenia</li> </ul>
	Olanzapine (Zyprexa <sup>®</sup> ; Zyprexa Zydis <sup>®</sup> )	PO/ODT: 2.5-5 mg nightly Age > 60 years: 2.5 mg nightly  Parenteral formulation non-formulary	20 mg/day	6 hours	
		PO: 20 mg every 12 hours  IM: 10 mg every 2 hours PRN or 20 mg every 4 hours PRN	PO: 160 mg/day IM: 40 mg/day	<b>PO:</b> 6-8 hours <b>IM:</b> ≤ 60 minutes	



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## **APPENDIX E: Medications for Management of Delirium in Critical Care Unit Only**

Therapeuti Class	Medication	Typical Initial Infusion Rate	Recommended Maximum Infusion Rate	Onset of Action	Comments/Cautions/Adverse Reactions
Alpha Agon	st Dexmedetomidine (Precedex®)	IV infusion: 0.2 mcg/kg/hour	1.4 mcg/kg/hour	Immediate	<ul> <li>Refer to Critical Care Sedation for Mechanically Ventilated Adult Patients order set for treatment of delirium in mechanically ventilated patients</li> <li>Refer to ICU Dexmedetomidine for Non-Mechanically Ventilated Patients order panel for treatment of delirium in non-mechanically ventilated patients</li> <li>Caution with use of &gt; 0.7 mcg/kg/hour in non-mechanically ventilated patients</li> <li>Bradycardia, hypotension</li> <li>Do not use if heart rate &lt; 60 bpm or MAP &lt; 65 mmHg</li> </ul>



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