

# MANAGEMENT OF ACUTE DELIRIUM (on floors)

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## DELIRIUM PROPHYLAXIS FOR HIGH-RISK PATIENTS

Patient is high risk if ≥2 of these criteria present: -

- Age ≥65 years
- Cognitive Impairment (current admission or prior history)
- History of Delirium
- History of Parkinson's disease
- Current admission for Stroke or Traumatic Brain Injury
- Patient receiving any of these medications this admission: Opiates, Benzodiazepines (eg. Valium/Ativan), Corticosteroids, or Diphenhydramine (Benadryl)
- Hearing or Vision Impairment
- Language Barrier
- History of Bipolar disorder, Depression, PTSD, or Schizophrenia
- History of substance abuse



- \* Minimize deliriogenic medications
- \* Non-pharmacological measures (By Nursing)
- \* Melatonin 3 mg PO qHS (2000 hrs)

## HYPOACTIVE DELIRIUM

If ≥4 of these criteria present & based on clinical assessment: -

- Unawareness
- Lethargy
- Sparse/Slow speech
- Staring
- Decreased alertness
- Lack of enthusiasm
- Decreased/Slow movement



- \* Minimize deliriogenic medications
- \* Non-pharmacological measures (By Nursing)
- \* Work up precipitating cause of delirium
- \* May consult Neurology/Psychiatry for diagnosis/management

Inouye SK, et al. JAMA 1996;275(11):852-7.

Moerman S, et al. Gen Hosp Psych 2012;34:153-9.

Oldenbeuving AW, et al. Neurol Neurosurg Psychiatry 2013;00:1-4.

Liptzin B, et al. Br J Psych 1992;161:843-5.

# HYPERACTIVE DELIRIUM\*

For all patients with Hyperactive Delirium:

- \* Minimize deliriogenic medications
- \* Melatonin 6 mg PO qPM (at 2000 hrs)
- \* Non-pharmacological measures (By Nursing)
- \* Sleep aid qHS PRN (such as Trazodone 25 mg or Zolpidem 2.5 mg) IF no results with Melatonin and non-pharmacological measures
- \* Work up precipitating cause of delirium

Lewy Body Dementia

- a) No antipsychotics
- b) Continue non-pharmacological measures
- c) Consult Neurology

QTc > 500 ms

Yes

Parkinson's disease / Traumatic Brain Injury

No

- a) Non-pharmacological measures
- b) Remove other QT prolonging medications if possible and reassess QTc
- c) If persistent agitation  
→ Consult Neurology/Psychiatry

Patient able to swallow

Yes

**PO QUETIAPINE**  
**(Sedating)**

No

Consult Psychiatry/  
Neurology

**PO QUETIAPINE**  
**(1<sup>st</sup> choice)**  
**(Sedating)**

**IV HALOPERIDOL**  
**(2<sup>nd</sup> choice)**

*	IV HALOPERIDOL (Haldol)	PO QUETIAPINE (Seroquel)
<b>MILD AGITATION</b> (easily redirectable patient: RASS +1)	0.25-0.5 mg q6h PRN	12.5 mg q6h PRN
<b>MODERATE AGITATION</b> (Restless, impulsive, poor safety awareness, not easily redirectable: RASS +2)	1-2 mg qHS + 0.5-1 mg q6h PRN	25 mg qHS + 12.5 mg q6h PRN
<b>SEVERE AGITATION</b> (Pulling lines/tubes, combative: RASS +3, +4)	2-4 mg qHS + 1-3 mg q4h PRN	25 mg qHS (May repeat x1) + 25 mg q6h PRN

	<b>IV/PO Haloperidol</b>	<b>PO Quetiapine</b>	<b>Melatonin</b>	<b>Trazodone</b>	<b>Zolpidem (Ambien)</b>
<b>Pharmacology</b>	Dopamine D2 > Serotonin 5HT <sub>2A</sub> receptor blockade.  <u>IV Haloperidol:</u> 100% Bioavailability Onset: seconds, Duration: 4-6 hrs Elimination half-life: 26-56 hrs  <u>PO Haloperidol:</u> 60-70% Bioavailability (lower if ileus, diarrhea) Peak action: 1.5-6 hrs Elimination half-life: 14-30 hrs	Serotonin 5HT <sub>2A</sub> > Dopamine D2 receptor blockade, Adrenergic ( $\alpha$ 1/2), Histaminergic H1 receptor, and Muscarinic M1 receptor blockade  Peak: 1-2 hrs, Half-life: 6 hrs (Half-life of metabolite: 12 hrs)	Hormone from pineal gland. Acts on melatonin receptors. Improves sleep quality.  Peak: 1 hr, Half-life 0.5-2 hrs	Unclear mechanism. Improved sleep quality (reduced awakenings), no improvement in sleep onset latency or total sleep time.  Onset of action: 1-3 hrs, Half life 11 hrs in age >65 yrs	Non-BZD: Agonist at one subtype of GABA-A receptor (BZD are agonists at all subtypes). May reduce sleep onset latency, total sleep time, no effect on sleep quality (awakenings).  Onset of action: 30 min, Half life <3 hrs, no active metabolite, effect may lasts up to 8 hrs
<b>Safety Profile</b>	1) <u>Extrapyramidal side effects (EPS) less with IV than PO Haloperidol</u> (7.2% for IV versus 22.6% for PO Haloperidol) - Akathisia (30%, Dose/Duration dependent) - Tardive dyskinesia (4-5%) - Acute dystonia (14%) - Medication induced Parkinsonism (50%, Dose/Duration dependent) - Less EPS if dose <4.5 mg/d  2) <u>QT prolongation more with IV than PO</u> Haloperidol: Ventricular arrhythmias/Torsades de pointes (<0.3%), especially if other QT prolonging drugs being given and/or electrolyte abnormalities 3) Minimal sedative/hypotensive effects 4) Prolactin elevation (reversible)	1) <u>SEDATION</u> (18-35%, H1 receptor) 2) Orthostatic hypotension (7% - $\alpha$ 1/2 blockade) 3) Minimal to no EPS 4) Ok to use in Parkinson's disease 5) Rare QT prolongation 6) Minimal to no effect on Prolactin 7) Moderate anticholinergic activity	At high doses: Daytime fatigue, Dizziness.	Anticholinergic. Hypotension (10%), Next day sedation & Dizziness (21%), N/V (15%), Headache (10%), Priapism, Constipation (13%), Rebound insomnia after discontinuation on 2nd withdrawal night, worsen RLS	At doses >20 mg/d → Drowsiness, Falls, N/V. May have some rebound insomnia. Reports of confusion/delirium/falls (?dose of zolpidem), ?fugue state (sleep-walk/eat/drive).
<b>Drug interactions</b>	Carbamazepine and Phenytoin may <i>increase</i> clearance of Haloperidol. Prozac, Wellbutrin, Paxil, Phenergan, Effexor may <i>decrease</i> clearance of Haloperidol.	Carbamazepine and Phenytoin may <i>increase</i> clearance of Quetiapine.			

Antipsychotics: Wang EHZ, et al. Neurocrit Care 2012;16:170-83. Maldonado JR, et al. J Psychosom Res 2003;55:140-1. Lonergan E, et al. Cochrane Database Syst Rev 2007; (2). Sleep aids: Ramakrishnan K, et al. AAFP 2007;76:517-26. Bellon, A. J Psych Practice 2006;12:229-43. McCall WV. Prim Care Companion J Clin Psych 2004;6:9-20. Al-Aama T, et al. Int J Geriatr Psych 2011;26:687-694. Bellapart J, et al. Br J Anaesth 2012;108(4):572-80. Kolla BP, et al. JHM 2013;8:1-6. Hill KP, et al. Psychosomatics 2004;45(1):88-9.