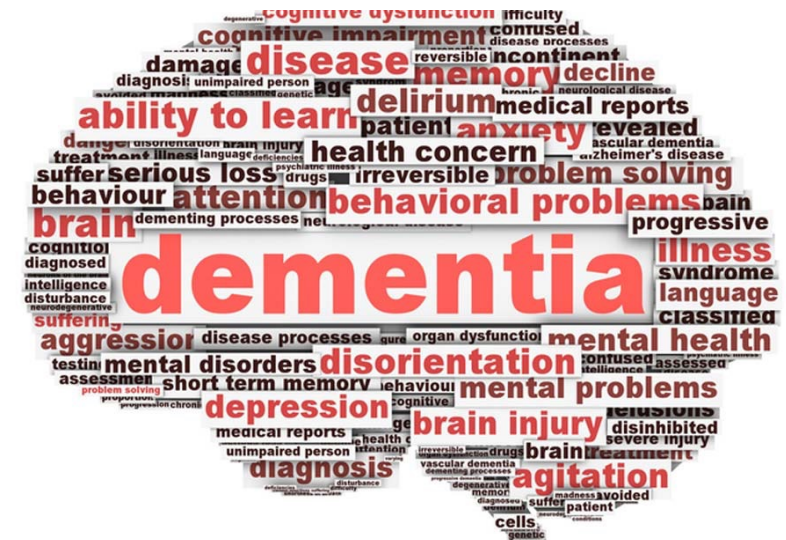




Nancy Isenberg MD MPH



## Objectives:

- Implement 5M person-centered care in Dementia
- Understand Delirium risks, prevention, treatment pathways

## How does AFHS (5M) improve dementia care?

- Dementia care is whole person care, at the core of which is the person with dementia and their caregiver (the dyad)
- Dementia care is complex, but it's the simple interventions that matter most.
- Incorporating 5Ms to guide your care of dementia patients (and all older adults, and all people!) will help you provide consistent and comprehensive person-centered care.
- <https://www.youtube.com/watch?v=QoODRr4m0kA>





# Choosing what matters, doing what works

## Patient Priorities & Care

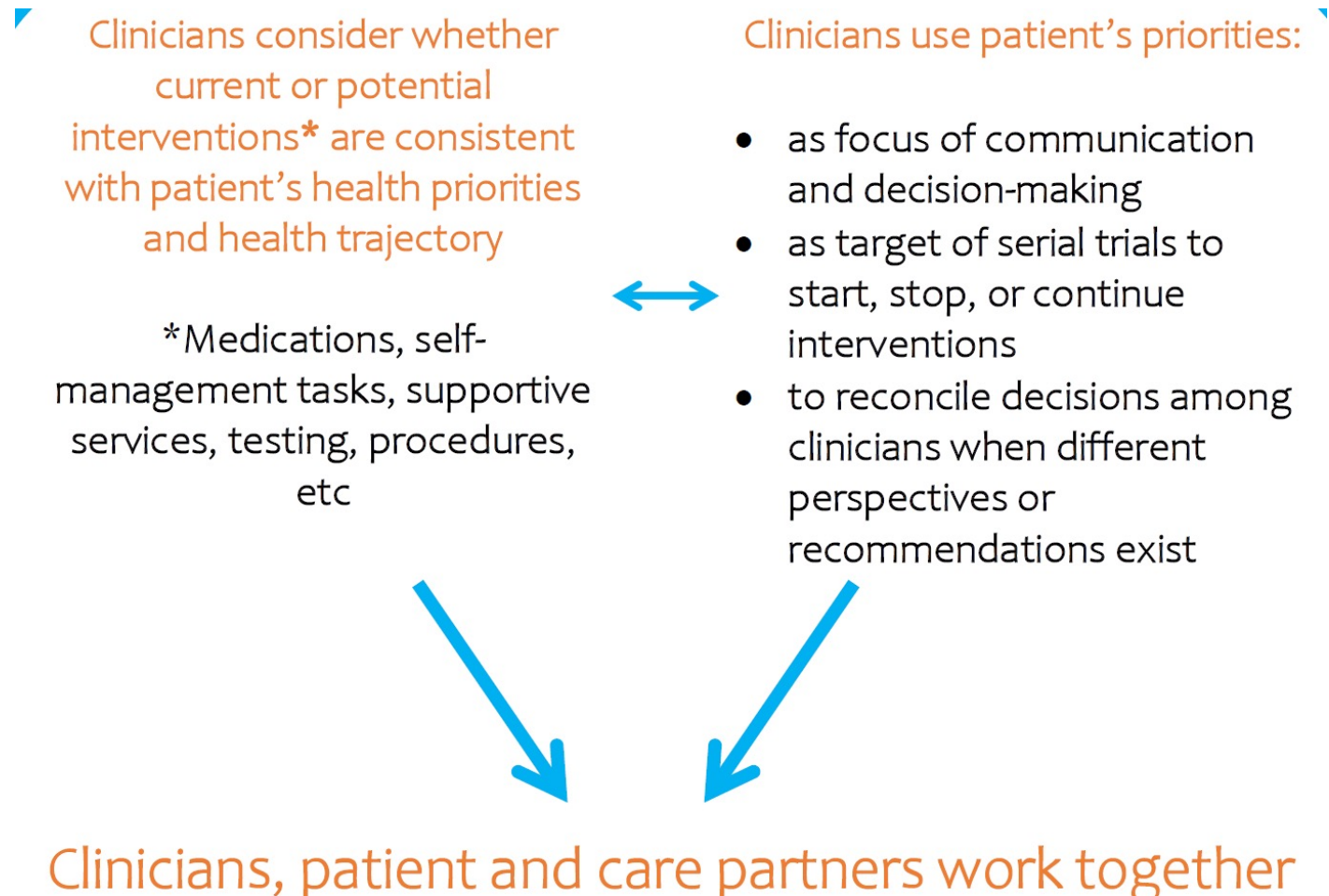
M Tinetti MD

- <https://patientprioritiescare.org/patient-facing-materials/>
- <https://patientprioritiescare.org/what-is-patient-priorities-care-and-why-is-it-important/>

### Patient's Health Priorities are identified

- Values (What Matters Most)
- Actionable, specific and realistic health outcome goals
- Healthcare preferences (care that is helpful or burdensome) and tradeoffs
- "One Thing" patient most wants to address

# Aligning care with patient's priorities



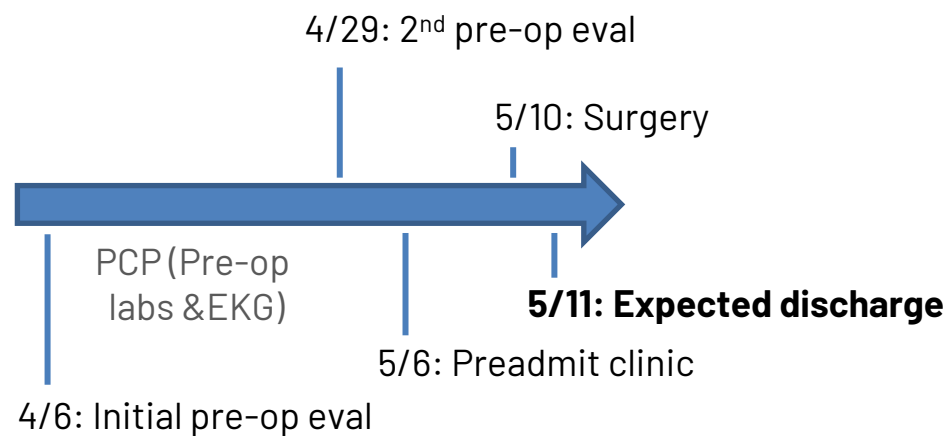
# Hospitalization Rates in Dementia

- Each year **40%** of community-dwelling People with dementia (PwD) will visit ED and **30%** will be hospitalized at least once.
- Hospital care is **3 times as costly** compared to older people w/o dementia
- Acute hospitalization in PwD is associated with increased **risk of delirium**, falls, **cognitive and functional decline**, **30 day readmission**, longer LOS, long-term care admission and death
- ***What are causes of delirium you commonly see in your practice?***

- Shepherd et al, BMC Medicine 2019

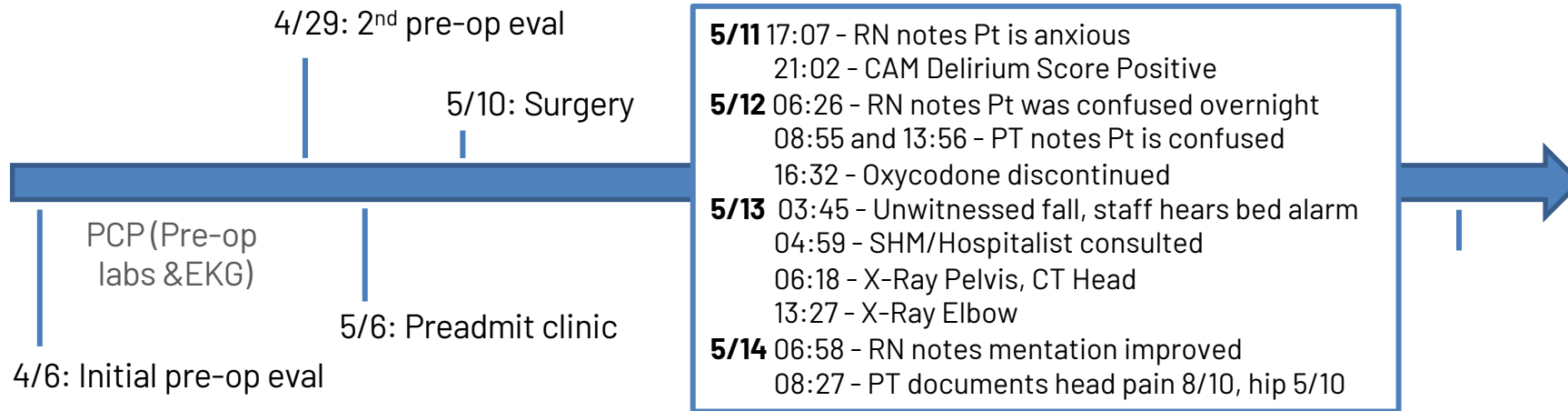
# Delirium related safety event

**72yo F sustains a radial head fracture and head trauma after an unwitnessed fall on POD#3 from her THA, in the setting of post-operative delirium**



# Safety Event

**72yo F sustains a radial head fracture and head trauma after an unwitnessed fall on POD#3 from her THA, in the setting of post-operative delirium**



**LOS 3 days > expected | Hospital acquired radial head fracture | DOH reportable fall**



## Safety Event – MAR

**5/10**

13:17 Scopolamine patch  
 13:17 Famotidine 20mg BID started  
 21:27 Tylenol 1000mg  
 21:28 Xanax 1mg  
 22:37 Ambien 10mg

**5/11**

08:39 Oxycodone 5mg  
 12:22 Oxycodone 5mg  
 12:54 Xanax 1mg  
 14:01 Tylenol 1000mg  
 15:54 Oxycodone 5mg  
 19:59 Oxycodone 5mg  
**21:01 CAM Positive**  
 21:52 Xanax 1mg  
 21:52 Oxycodone 5mg  
 22:00 Ambien 10mg

Narcotic

Anticholinergic/Antihistamine

**5/12**

01:41 Oxycodone 5mg  
 06:22 Tramadol 50mg  
 06:22 Tylenol 1000mg  
 09:37 Xanax 1mg (initially held at 08:22  
 as Pt was delirious)

14:44 Tylenol 1000mg  
 21:06 Tylenol 1000mg  
 22:45 Ambien 10mg

**5/13**

**03:45 Unwitnessed fall**  
 08:47 Lidocaine patches added  
 13:59 Scopolamine patch removed  
 22:29 Melatonin 3mg added  
*No additional Ambien, Oxycodone, or  
 Xanax administered*

Benzodiazepine / Benzodiazepine receptor agonists

Unchanged Medications	Details
acetaminophen 650 MG CR tablet	Take 1 tablet by mouth every 8 hours. aka: TYLENOL 8 HOUR
albuterol 90 mcg/puff inhaler	INHALE 2 PUFFS INTO THE LUNGS EVERY 8 HOURS AS NEEDED FOR WHEEZING
ALPRAZolam 0.5 mg tablet	TAKE 1 TO 2 TABLETS BY MOUTH EVERY DAY AS NEEDED. aka: XANAX
aspirin 325 MG EC tablet	Take 1 tablet by mouth 2 times daily.
BENADRYL ALLERGY PO	Take by mouth nightly as needed.
CLARITIN-D 24 HOUR PO	Take by mouth Daily as needed
cyclobenzaprine 10 mg tablet	TAKE 1 TABLET BY MOUTH EVERY NIGHT aka: FLEXERIL
EPINEPHrine auto-injector 0.3 mg/0.3 mL injection	Inject 0.3 mLs into the muscle as needed for Anaphylaxis.
FLONASE 50 mcg/nasal spray Generic drug: fluticasone	1 spray by Nasal route as needed .
fluconazole 200 MG tablet	TAKE 1 TABLET BY MOUTH EVERY DAY FOR 10 DAYS, REPEATING AS NECESSARY. aka: DIFLUCAN
gabapentin 300 mg capsule	TAKE 3 CAPSULES BY MOUTH EVERY NIGHT aka: NEURONTIN
ixekizumab 80 mg/mL injection (syringe)	Inject 80 mg under the skin Every 30 days. aka: TALIZ
meloxicam 7.5 mg tablet	Take 1 tablet by mouth Daily. aka: MOBIC
MULTIVITAMIN ADULT PO	Take by mouth Daily.
ondansetron 4 mg disintegrating tablet	Take 1 tablet by mouth every 6 hours as needed for Nausea or Vomiting. aka: ZOFRAN ODT
polyethylene glycol 17 g/capful powder	Take 17 g by mouth Daily as needed Mix into 4-8 oz of juice or water and take by mouth once daily as needed for constipation. aka: MIRALAX
valACYclovir 1 g tablet	TAKE 1 TABLET BY MOUTH THREE TIMES DAILY AS NEEDED. aka: VALTREX
zolpidem 10 mg tablet	Take 1 tablet by mouth nightly. aka: AMBIEN

- resources for deprescribing
- <https://deprescribing.org/resources/deprescribing-guidelines-algorithms/>



AARP

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Renew

Help

Member Benefits



AARP Rewards

Register | Login



likelihood of long-term consequences from this potential complication.



Health care professionals think **patients** should be informed about delirium, but are they?

Health care professionals who agree that delirium should be discussed with patients prior to surgery or hospitalization

84%

Patients who said they were informed about delirium prior to surgery or hospitalization

4%

## 6 D's of Delirium

- Distress
- Death
- Duration
- Dollars
- Dementia
- Loss of Dignity

+ a b | e a u



# Delirium

Acute & **under-recognized** disorder of attention, consciousness and cognition

Types:

Hyperactive with agitation

Hypoactive with lethargy- assc. w/ worse outcomes, higher mortality, lower QOL, longer LOS, increased falls, institutionalization.

Mixed BOTH

In ~20% cases delirium may persist for weeks to months

Associated with high rates of rapid readmission, increased mortality, cost, LOS (twice length and average \$2500/pt) & post-op/post-hospital cognitive decline, institutionalization

# Prevalence

- 2020 meta-analysis of 33 studies of medical inpatients found ~1/4
- Increased in major surgery
- Meta-analysis after acute stroke 32 studies ~1/4 post-stroke
- 2019 systematic review 42 studies with pooled prevalence 35% across community to inpatient units receiving palliative care
- Nursing homes 4-38%
- Ventilated patients in ICU 50-70%

Wilson J. et al *Nature*, 2020

Mart, M et al, *Semin Respir Crit Care Med* 2021

# Delirium

- 48% hospital days are patients >65 yr
- 42% medical patients and 80% of critical care patients experience *undiagnosed* delirium, likely higher in Covid
- 73.5% incidence of delirium during perioperative period
- Delirium is WELL ESTABLISHED RISK FACTOR for incident dementia and worsening of existing dementia.



# Delirium: a missing piece in the COVID-19 pandemic puzzle

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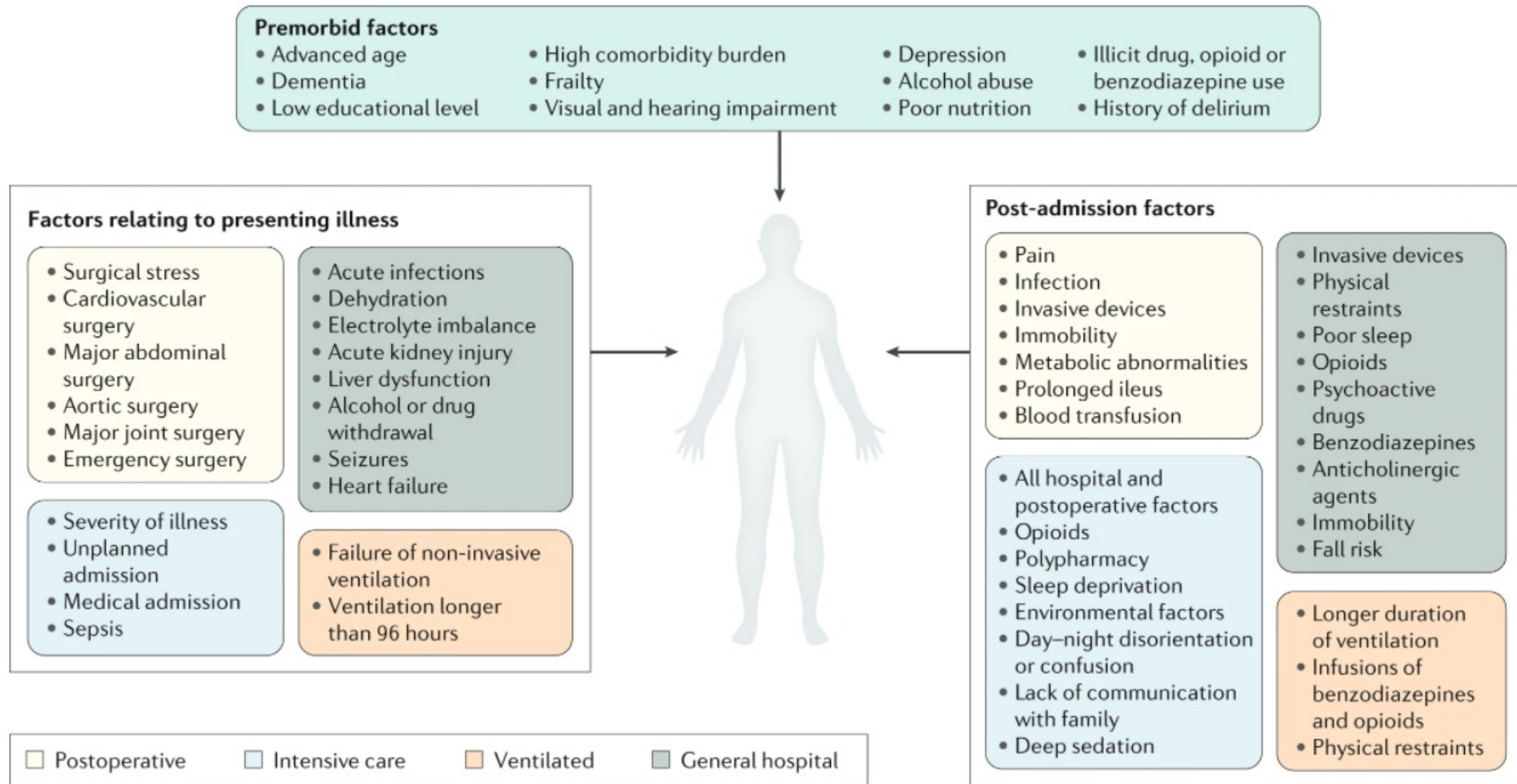
## Key points

- Older people are most vulnerable to severe COVID-19 infections and mortality.
- Current guidance for diagnosis does not routinely include delirium, which may lead to under-detection of COVID-19.
- The care home population is particularly at risk, as failure to promptly detect COVID-19 may lead to outbreaks.
- Non-pharmacological approaches to management of delirium may be more difficult to implement but remain the priority.

Mnemonic Clinical causes and interventions **Dr. DRE Diseases** Evaluate the patient for new or worsening disease, such as congestive heart failure or sepsis, as well as other disease findings such as metabolic abnormalities

**Drug Removal** Look for and stop deliriogenic medications including benzodiazepines, antihistamines, and inappropriate opioids

**Environment** Encourage daytime mobilization and remove restraints, provide frequent reorientation as well as cues such as clocks and windows to the outside, reduce night-time interventions to promote restful sleep



Risk factors for delirium relate to premorbid or predisposing factors (that is, a patient's characteristics) and to precipitating factors, which are factors relating to the presenting illness or that occur after hospital or intensive care unit admission.

# Across Care Continuum

## Box 2 | Delirium prevention in different health-care settings

Consensus guidelines<sup>241,262</sup> make a number of recommendations for delirium prevention in various health-care settings.

### General settings

#### Multicomponent interventions

- Early recognition of high-risk factors (age >65 years, dementia, hip surgery and high acuity)
- Daily screening for delirium
- Environmental orientation (sensory, auditory, dentures, time, events, family visits and music)
- Maintain normal hydration
- Regulation of bladder and bowel function
- Early establishment of normal diet
- Correction of metabolic disorders
- Cardiorespiratory optimization (with provision of oxygen if appropriate)
- Early identification of infection
- Effective treatment of pain
- Daily mobilization
- Avoidance of antipsychotic drugs
- Avoidance of benzodiazepines
- Reduced nocturnal disturbances to promote sleep

- Early removal of devices (intravascular and airway devices)
- Avoidance of physical restraints
- Sleep promotion (eye mask and earplugs)

#### Pharmacological interventions

- None with high-level evidence

### Intensive care and high acuity units, intubated and non-intubated patients

#### Non-pharmacological interventions (as above, consider also)

- Early recognition of high-risk patients (age >65 years, high acuity, sepsis, shock, dementia and ventilation)
- Light sedation
- No benzodiazepines
- Early mobilization
- Promotion of day-night routine
- Environmental awareness and orientation
- Removal of devices (intravascular and airway devices)

#### Pharmacological interventions

- Some suggestive evidence but not recommended by consensus guidelines

## Table 1 Primary delirium prevention principles

### Primary delirium prevention principles

- Repeated reorientation
- Provisions of cognitively stimulating activities multiple times a day
- A sleep protocol
- Early mobilization
- Timely removal of catheters and physical restraints
- Use of eye glasses, magnifying lenses, hearing aids, and earwax disimpaction
- Correction of dehydration
- Use of a scheduled pain management protocol
- Minimization of unnecessary noise and tactile stimuli

<https://www.cms.gov/Outreach-and-Education/Medicare-Learning-Network-MLN/MLNProducts/downloads/Transitional-Care-Management-Services-Fact-Sheet-ICN908628.pdf>

# Delirium Prevention

## **30-40% cases are preventable**

- Avoid anticholinergics, benzodiazepines, opioids, H2 blocker, TCA, steroids, 3<sup>rd</sup> gen cephalosporins
- Fluid management- avoid dehydration
- Early mobilization
- Avoid sleep disturbances
- Minimize perceptual deficits/glasses/aids
- Environmental awareness, nutrition, oxygenation.
- HELP, ABCDE bundle and ACE program multicomponent program
- Pharmacist led medication review in institutional long-term care

# Assessment

- Confusion Assessment Method (CAM) has sensitivity of 94-100%, specificity 90-95%
1. Acute onset/fluctuating course
  2. Inattention
  3. Disorganized thinking
  4. Altered level of consciousness
- (see resources in appendix)



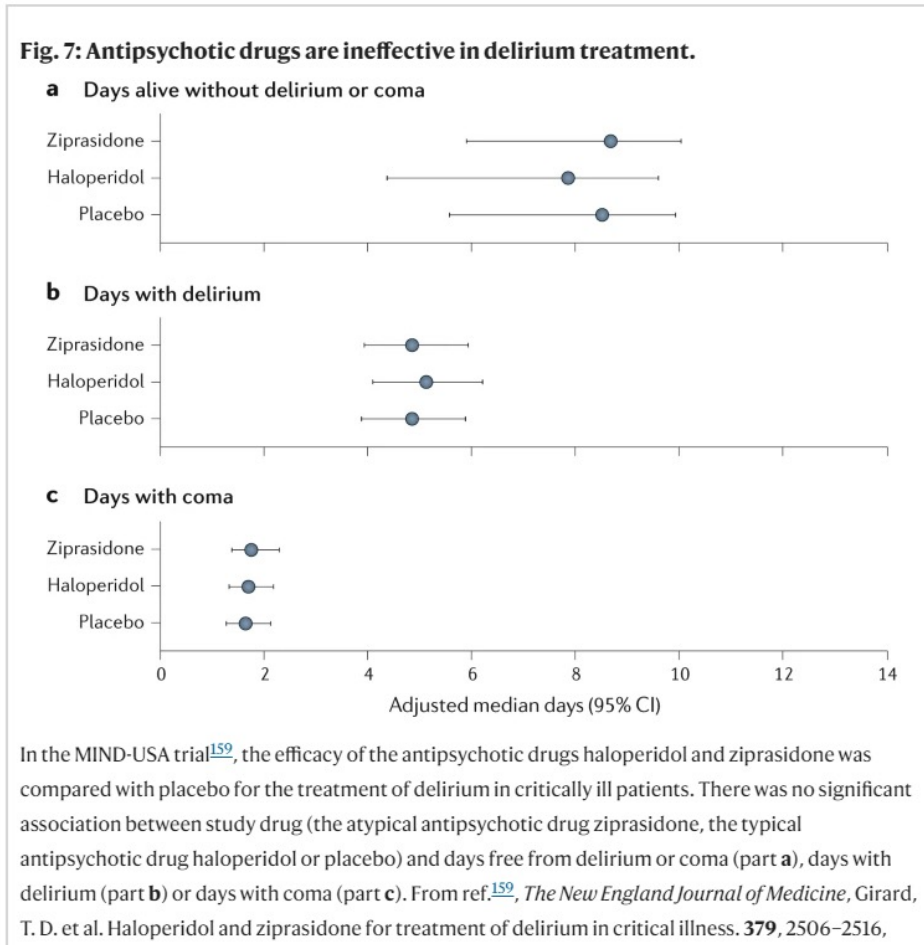
# Delirium Management

- 2019 Cochrane Review of 14 trials with 1844 patients:
  - Precedex (dexmedetomidine) is most preferred as compared to placebo for shortening length of ICU stay
  - No antipsychotic has been shown to treat length/severity of delirium, pre-op administration does not prevent occurrence of delirium
- Melatonin has only weak evidence for benefit, newer agents ramelteon and orexin-antagonist suvorexant being studied
- Anticholinesterase (e.g Rivastigmine) show no benefit, may increase LOS
- Avoid benzodiazepines, can worsen delirium

# Antipsychotics in delirium treatment

- 2018, 2019, 2020 Cochrane reviews no effect on delirium severity, symptom resolution or mortality. No effect on duration, severity or LOS. Efficacy unclear
- Cholinesterase inhibitors unclear benefit, but abruptly stopping can worsen delirium
- If distressing psychosis/intractable distress there is limited role in patients with delirium when other measures have been ineffective  
(SIGN 2019)
- Staffing shortages, safety issues eg falls, self or other harm, and avoiding physical restraints

# MIND-US largest randomized study examining antipsychotics for delirium in 566 ICU patients with delirium.



# ABCDEF Bundle

## 6 PROVEN\* STRATEGIES TO PREVENT DELIRIUM IN OLDER ADULTS

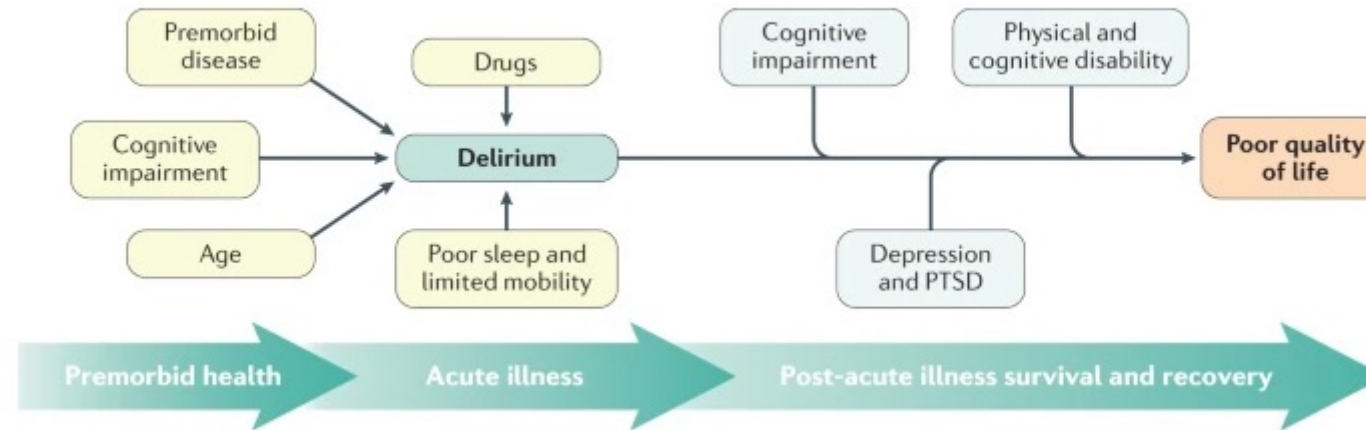


ABCDEF bundle component <sup>267</sup>	Approach	Delirium risk factor
A	Assess, prevent and manage pain	Undertreated pain; over-sedation caused by analgesics
B	Both spontaneous awakening trials and spontaneous breathing trials	Over-sedation; mechanical ventilation
C	Choice of sedation and analgesia	Over-sedation caused by analgesics; over-sedation caused by sedatives; exposure to deliriogenic sedatives (for example, benzodiazepines)
D	Delirium: assess, prevent and manage	Disordered sleep-wake cycle; vision and hearing impairment; other unrecognized delirium risk factors
E	Early mobility and exercise	Immobility
F	Family engagement and empowerment	Disorientation

Information compiled from ref.<sup>267</sup>.

# Delirium and QOL

**Fig. 9: Relationship between delirium and post-ICU quality of life.**





## Mild cognitive impairment and perioperative delirium

1. Delirium is common in the elderly following surgery, especially in those with preexisting reduction in neurocognitive function. Other RF include frailty, ICU admission, foley use
2. Delirium is associated with risk of cognitive decline following surgery.
3. 60% of geriatric patients with delirium at hospital admission developed dementia over next 3 years compared to 18.5% of those without delirium.
4. 23% of patients with perioperative delirium developed MCI at first follow up compared to 7% w/o delirium.
5. **MCI increases risk of perioperative delirium.**

[Mayo Clin Proc. 2016 Feb; 91\(2\): 273–274.](#)

doi: [10.1016/j.mayocp.2015.12.008](#)

# Medication overview

## Key messages for patients about medication management

- A complete medication review (including over-the-counter and herbal remedies) should be performed frequently, and especially during care transitions, such as post-surgery or hospital discharge.
- Over-the-counter medications (such as antihistamines, sedatives, and other medications that have strong anticholinergic activity), may have significant cognitive side effects, so their use should be carefully assessed.

### Online resources for safe medication use in older adults:

- American Geriatrics Society Beers Criteria (including public education resources)
- American Geriatrics Society—What To Do and What to Ask Your Healthcare Provider If A Medication You Take is Listed in the Beers Criteria for Potentially Inappropriate Medications to Use in Older Adults
- National Institute on Aging—Safe Use of Medicines
- National Institutes of Health Senior Health—Taking Medications Safely
- Centers for Disease Control and Prevention—Adults and Older Adult Adverse Drug Events
- Food and Drug Administration—Medicines and You: A Guide for Older Adults
- Institute of Medicine—Preventing Medication Errors: Quality Chasm Series

[www.nas.edu/cognitiveaging](http://www.nas.edu/cognitiveaging)  
<https://www.youtube.com/watch?v=E1h5jOWdX30>

# Early detection of MCI/dementia

- Growing evidence regarding the importance of early detection and accurate diagnosis –
  - Cardiometabolic risk reduction/improving management of comorbid conditions/reducing polypharmacy/iatrogenic harm
  - Connecting with community resources, programs/services
  - Reducing preventable hospitalizations, inappropriate surgeries, delirium, PHCD/POCD and emergency room visits
  - Identifying goals around end-of-life care and improving advance care planning

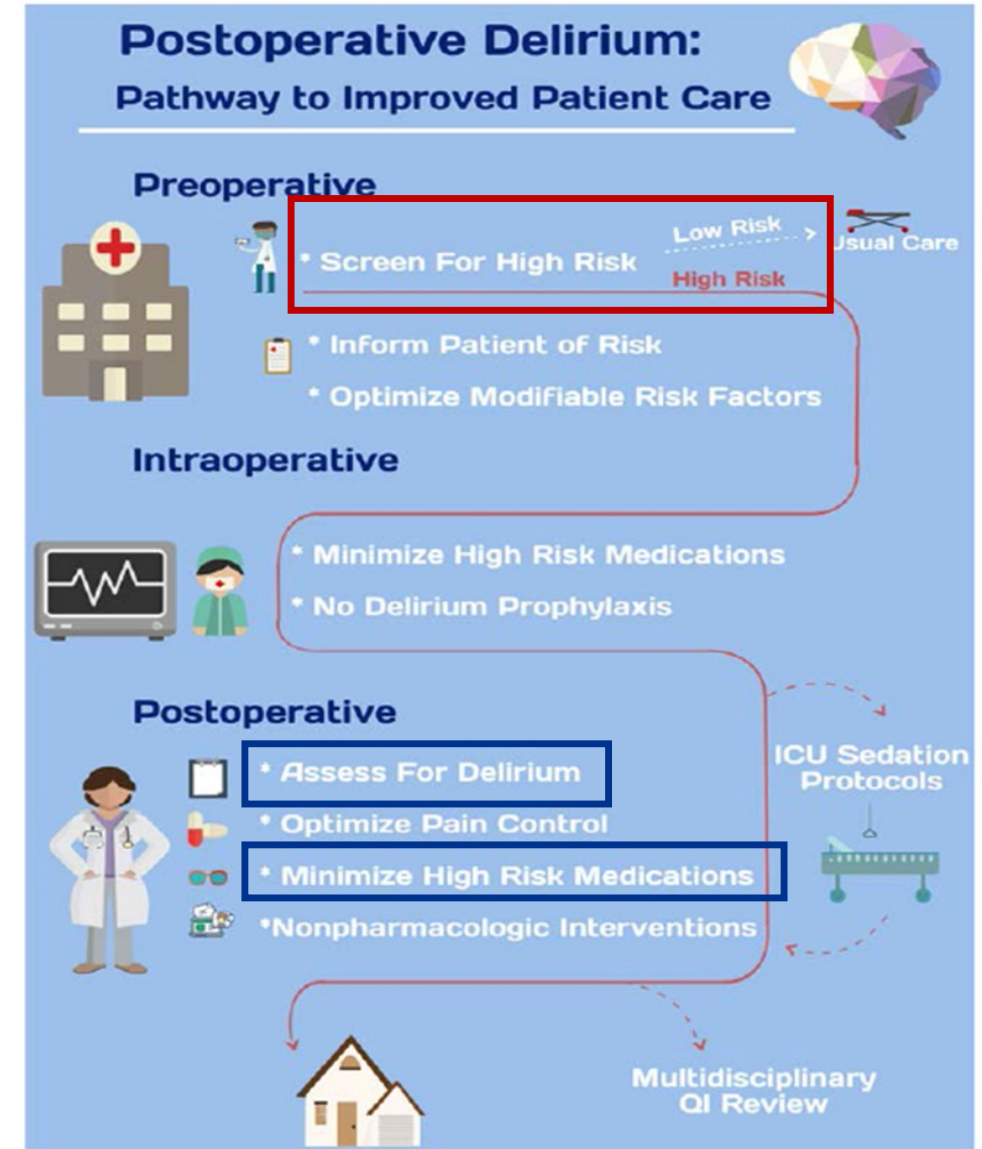
# Post-Operative Delirium

Occurs in up to 50% of patients after major surgery<sup>1</sup>

Associated with:

- Increased length of stay
- Higher rates of institutionalization after discharge
- Higher rates of readmission
- Functional decline and cognitive impairment after surgery

**30-40%** of delirium cases are preventable<sup>2</sup>



American Society for Enhanced Recovery and Perioperative Quality Initiative Joint Consensus Statement on Postoperative Delirium Prevention  
June 2020 doi: 10.1213/ANE.00000000000004641

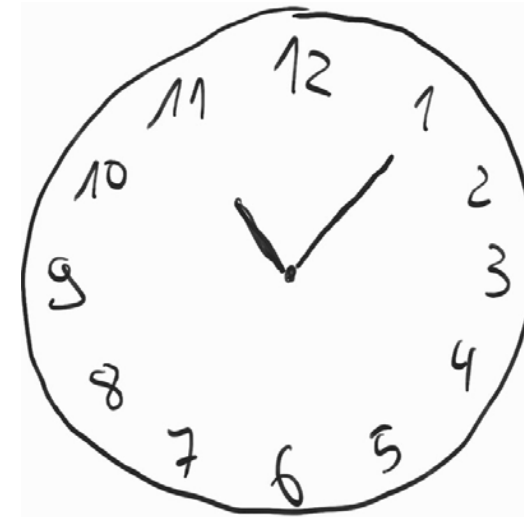
## Delirium Risk Screening | Brief Cognitive Screen

Patients with **Mild Cognitive Impairment (MCI)** are at higher risk for delirium (16/126 (13%) vs 12/562 (2%), **OR (95% C.I.) = 6.6**,  $P < 0.01$  in a 2017 Mayo Clinic retrospective analysis<sup>1)</sup>)

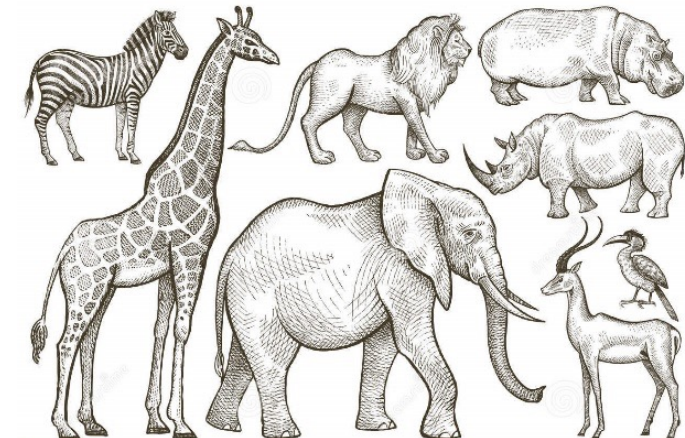
**Dementia** patients have superimposed delirium during hospitalizations at rates ranging from **22% to 89%**<sup>2</sup>

- In a 2001 prospective cohort of Swedish hip fracture patients age >65 (n=101), 86% of the patients with dementia developed delirium (18/21)

**MCI has an estimated prevalence of 20% in patients > 65, and dementia 6.5%. Both are underrecognized and underdiagnosed.**



Train  
Egg  
Hat  
Chair  
Blue



1. Perioperative Delirium and Mild Cognitive Impairment <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4961244/>

2. Delirium Superimposed on Dementia [https://agsjournals.onlinelibrary.wiley.com/doi/full/10.1046/j.1532-](https://agsjournals.onlinelibrary.wiley.com/doi/full/10.1046/j.1532-5415.2002.50468.x?casa_token=kzbmSjY5ZEcAAAAA%3AFulk62k9gzVSHiztp-COaSu-segbHuOUaMjfr3D2BRA93Fcp4vNBq_5dsPABoeP5_I50b_ul9iL-m7M)

5415.2002.50468.x?casa\_token=kzbmSjY5ZEcAAAAA%3AFulk62k9gzVSHiztp-COaSu-segbHuOUaMjfr3D2BRA93Fcp4vNBq\_5dsPABoeP5\_I50b\_ul9iL-m7M



# Brief Cognitive Screen | Epic Integration (In progress)

**Flowsheets**

File Add Rows LDA Avatar Add Col Insert Col Data Validate Hide Device Data Last Filed Reg Doc Graph Gg to Date Responsible Refresh Chart Correction

Respiratory Assessment Vital Signs Complex RT Treatments Mechanical Vent NIV/CPAP Procedures/Monitoring Rapid Response Team Vital Signs **Medicare AWW** Medicare AWW

Search (...) Medic...

Hide All Show All

General... ☐ Hearin... ☐ Activiti... ☐ Home s... ☐ Fall Ris... ☐ Inconti... ☐ Advanc... ☐ Nutrition ☐ GET UP... ☐ Cogniti... ☐ **Brie... ☒** PHQ-2... ☐

1m 5m 10m 15m 30m **1h** 2h 4h 8h 24h Interval Start: 0700 Reset Now

Documentation from 10/14/2021 in Provid...

	10/14/21	1300
<b>Brief Cognitive Screen</b>		
Clock Drawing	1-Circle Dra...	
Verbal Fluency	2 - Twelve A...	
Delayed Recall	1- Hat	
<b>Total Score</b>		4

**Group Information**

[Patient Handout Resource](#)  
[Provider Guidelines](#)

**Cognitive Screening Patient Information**

Annual wellness visit - Cognitive screening is an important part of your annual wellness visit. This screening takes only a few minutes, but gives us good information about your risk for dementia. We strongly encourage our patients to do this screening annually, because if we are able to identify problems early we can intervene.

Pre-operative Consults - Cognitive screening is an important part of your preoperative visit. This screening takes only a few minutes, but gives us information about your risk for dementia. We also know that if we identify any problems, we may be able to lower your risk for getting a condition called "delirium" during your hospital stay. We will make sure to have Dr. \_\_\_\_\_ discuss this today as part of your visit."

**Screening Instructions**

**Memory:** Read the list of 5 words (Train, Egg, Hat, Chair, Blue) at a rate of one per second, giving the following instructions:

- "This is a memory test. I am going to read a list of words that you will have to remember now and later on. Listen carefully. When I am through, tell me as many words as you can remember. It doesn't matter in what order you say them.
- "I am going to read the same list for a second time. Try to remember and tell me as many words as you can, including

**Clock Drawing**

Select multiple options (F5)

1-Circle Drawn  
1-Correct Numbers  
1-Correct Time

Comments (Alt+M)

10/14/21 1200

**Verbal Fluency**

Select multiple options (F5)

2 - Twelve Animals Named  
0 - Zero to Eleven Animals Named

Comments (Alt+M)

10/14/21 1200

**Delayed Recall**

Select multiple options (F5)

1 - Train  
1 - Egg  
1 - Hat  
1 - Chair  
1 - Blue

Comments (Alt+M)



# Delirium Risk Screening – High Risk Medications

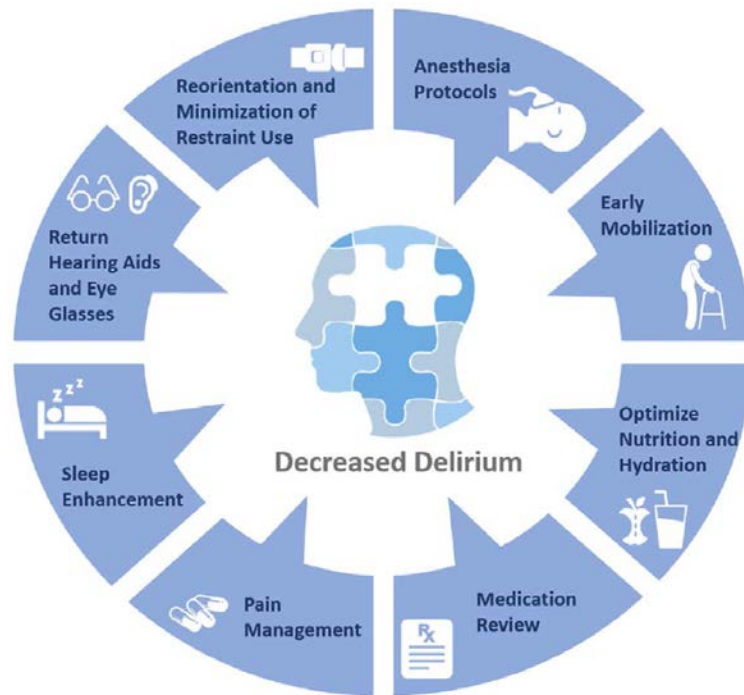


A long list of common offenders

- Benzodiazepines
- Antihistamines, especially 1<sup>st</sup> generation (diphenhydramine, promethazine, hydroxyzine)
- Opioids (note low-dose opioids for pain may reduce delirium, but excessive use or use as a sedative will exacerbate)
- Anticholinergics (tricyclic antidepressants, scopolamine, atropine)
- H2 blockers (cimetidine, famotidine, ranitidine)
- Zolpidem and related benzodiazepine-receptor agonist sedatives
- Antispasmodics (baclofen, cyclobenzaprine)
- Steroids
- Metoclopramide (reglan)

To be balanced against the risk of withdrawal effects for longstanding home medications

# Delirium Risk Screening and Prevention | Improved Outcomes



## Risk stratify patients pre-operatively

- Brief Cognitive Screen as a high yield tool
- Flag patients with multiple high risk medications
- Cognitive Vital sign visible on Dashboard

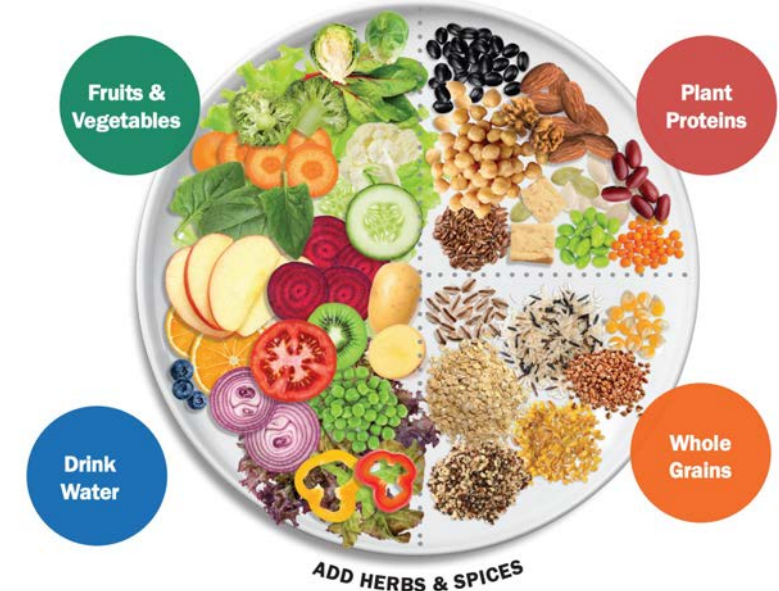
## Prioritize delirium prevention strategies for high risk patients once admitted

- Delirium and fall risk nursing protocols
- Earlier pharmacist and/or hospitalist assistance once delirium detected
- Consider order set adjustments (e.g. PPI instead of famotidine for stress ulcer prophylaxis)

# Brain Health Rx



- ❑ Alcohol (and drugs): Limiting 0-1 drinks
- ❑ Medications: deprescribe/avoid sedating and anticholinergic
- ❑ Contributing Conditions: Sleep apnea, hearing loss, cataracts.
- ❑ Exercise: >150 min/week aerobic & strength training 2-3 times week.
- ❑ Cognitive Stimulation/Mindfulness
- ❑ Socialization (generally more useful than puzzles) try HealthyMinds
  - <https://hminnovations.org/meditation-app>
- ❑ Nutrition Mediterranean/ MIND/ WFPD
  - ❑ SBP <120
  - ❑ LDL <70 if TIA or stroke



<https://www.aarp.org/health/brain-health/global-council-on-brain-health/resource-library/>

Thank you for your attention!



Questions?



# Contact Information

- Center for Healthy Aging

Swedish Neuroscience Institute. Moving in May to Cherry Hill

7320 216<sup>th</sup> Street SW

Edmonds, WA 98026

Ph. 206-320-7200 fax 425 673 3803

[Nancy.Isenberg@Swedish.org](mailto:Nancy.Isenberg@Swedish.org)

<https://www.swedish.org/locations/center-for-healthy-aging>



- Wilson, J.E., Mart, M.F., Cunningham, C. *et al.* Delirium. *Nat Rev Dis Primers* **6**, 90 (2020). <https://www.nature.com/articles/s41572-020-00223-4>
- Oldham MA, Flanagan NM, Khan A, Boukrina O, Marcantonio ER. Responding to Ten Common Delirium Misconceptions With Best Evidence: An Educational Review for Clinicians. *J Neuropsychiatry Clin Neurosci*. 2018 Winter;30(1):51-57. doi: 10.1176/appi.neuropsych.17030065. Epub 2017 Sep 6. PMID: 28876970.



# Age-Friendly Health Systems (AFHS)

Providence's 5 Ms for Age-Friendly Health

WHAT **MATTERS**



Know your care preferences and set goals for your health. Establish Advance Directives and Trusted Decision Makers.

**MEDICATION**



Manage your medications and understand how they may impact your mobility and cognition.

**MENTATION**



Get the emotional and cognitive support you need. Understand, prevent, and seek treatment for dementia, delirium, and depression.

**MOBILITY**



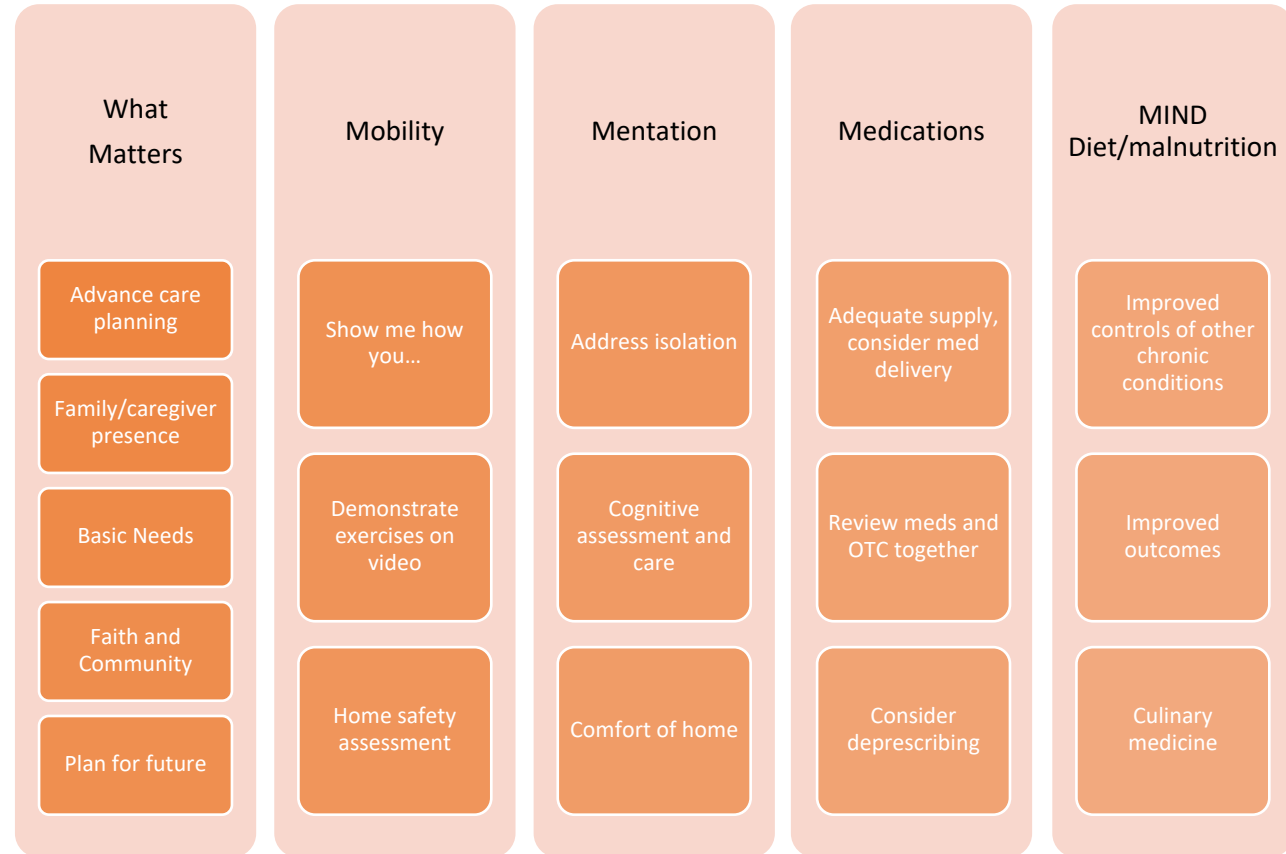
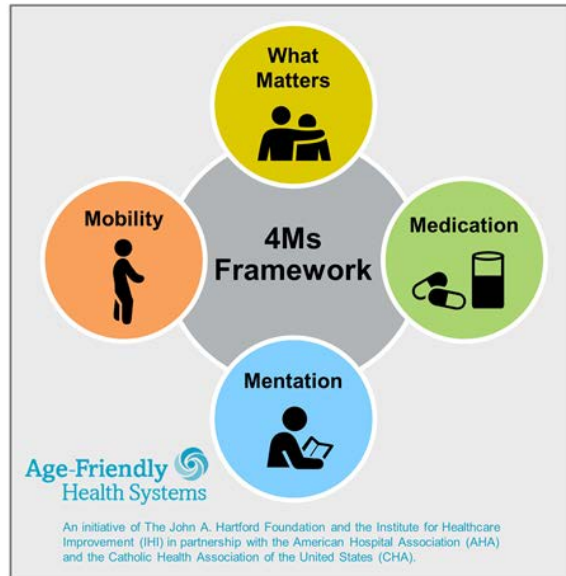
Keep active and mobile, preventing injuries and falls. Learn how to safely mobilize as you age.

**MALNUTRITION**



Commit to proper nutrition and assess malnutrition risk regularly.

# You can use the 5Ms in any setting: AFHS and Telehealth



ence  
ce  
Gr  
an

From: [Delirium](#)

### Microglia

Primed by prior degenerative pathology

Secrete IL-1 $\beta$ , TNF, NO and ROS

**Neuronal dysfunction and injury**

### Drugs

- GABAergic sedatives
- Anticholinergic drugs
- Antihistamine drugs

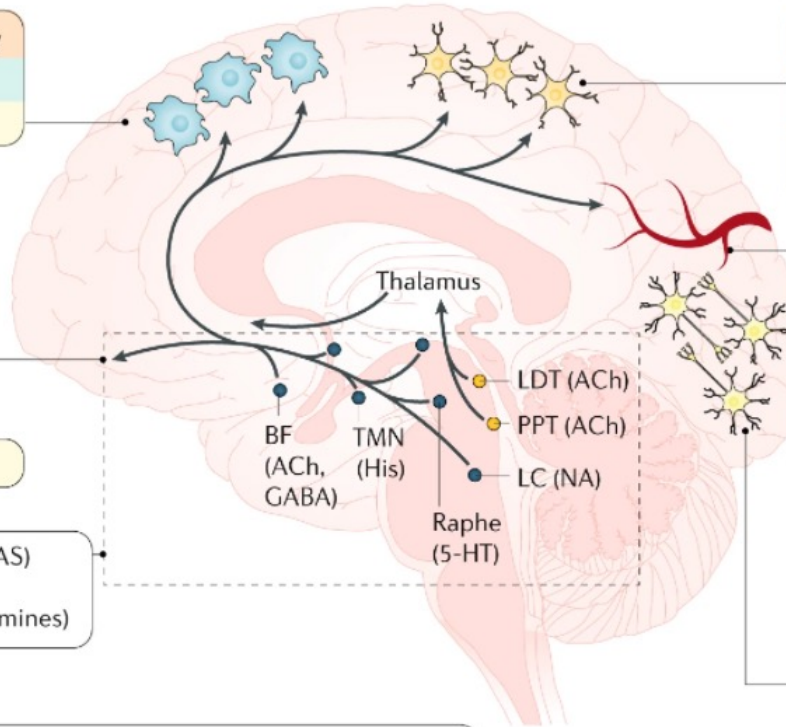
**Neurotransmitter disturbance**

- Reticular ascending arousal system (RAS)
- Thalamocortical activation (ACh)
- Cortical integration (NA, other monoamines)

### Systemic triggers

- Acute systemic inflammation
- Hypoxaemia ( $\downarrow$  O<sub>2</sub>)
- Blood flow (shock, impaired perfusion)
- Metabolic derangement (Na<sup>+</sup>, hypoglycaemia)

- Vulnerabilities
- Acute cellular changes
- Functional consequence



### Astrocytes

Primed by prior degenerative pathology

- Secrete chemokines
- Switched phenotype
- $\uparrow$  Immune cell infiltration
- $\downarrow$  Metabolic support

### Vasculature dysfunction

Endothelial and BBB dysfunction

- Further endothelial and BBB injury
- Impaired neurovascular coupling
- Microvascular dysfunction

**Metabolic insufficiency**

### Neuronal networks

Neurodegenerative pathology

- Synaptic changes
- Impaired neurovascular coupling

**Reduced integration of brain networks**

**Delirium**

# Confusion assessment method (CAM) for the diagnosis of delirium\*

\*The diagnosis of delirium requires the presence of features **1 AND 2 plus either 3 OR 4.**

## 1. Acute onset and fluctuating course

"Is there evidence of an acute change in mental status from the patient's baseline?";

"Did the abnormal behavior fluctuate during the day, that is, tend to come and go, or increase and decrease in severity?"

## 2. Inattention

"Did the patient have difficulty focusing attention, for example, being easily distractible or having difficulty keeping track of what was being said?"

YES if Letters attention test with >2 errors

Say C-A-S-A-B-L-A-N-C-A. Patient should squeeze your hand when the letter A is spoken. Error is missing an A or squeezing without an A.

## 3. Disorganized thinking

"Was the patient's thinking disorganized or incoherent, such as rambling or irrelevant conversation, unclear or illogical flow of ideas, or unpredictable switching from subject to subject?"

YES if the combined number of errors to questions and commands >1

Ask the patient the following yes/no questions and count errors: 1. Will a stone float on water?; 2. Are there fish in the sea?;

3. Does 1 pound weigh more than 2 pounds?; 4. Can you use a hammer to pound a nail?

Next, ask the patient to follow your commands: a) "Hold up this many fingers" (hold up 2 fingers) ;

b) "Now do the same thing with the other hand" (do not demonstrate the number of fingers).

If unable to move both arms, for part "b" ask patient to hold up one more finger. Count errors if patient is unable to complete the entire command.

## 4. Altered level of consciousness

YES if RASS is not 0 (calm and alert) - either sedated or agitated

Patient is either **CAM Positive** or **CAM Negative** after performing the above, with a sensitivity of 94-100% and specificity of 90-95%.

**Fig. 5: Common tools to screen for delirium in different settings.**

