

Family Medicine 29: 72-year-old male with dementia

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Learning Objectives

The student should be able to:

- Differentiate among the presentations of delirium, dementia, and depression in an older adult patient.
- Interpret at least one standardized instrument (e.g., MiniCog or Folstein Mini-Mental State Examination (MMSE)) to screen for cognitive loss in an older adult patient for whom there are concerns regarding memory or function.
- Describe baseline functional abilities (instrumental activities of daily living, activities of daily living) in an older adult patient with altered mental status.
- Formulate the differential diagnosis for acute change in mental status for a patient with dementia.
- Recognize management options, including pharmacological, non-pharmacological, complementary therapies, and caregiver support for an older adult patient with dementia.
- Recognize caregiver stress and its impact on the care of a patient with dementia.
- Describe therapeutic interventions to prevent or treat delirium in the hospital setting.
- Describe the role of social service, healthcare agencies, hospice, and other community organizations to provide care and assistance to older adult patients with dementia and their families.

Knowledge

Major Neurocognitive Disorder - DSM Criteria

DSM V Criteria for Major Neurocognitive Disorder (previously termed "dementia")

A. Evidence of significant cognitive decline from a previous level of performance in one or more cognitive domains:

- Learning and memory
- Language
- Executive function
- Complex attention
- · Perceptual-motor
- · Social cognition

B. The cognitive deficits interfere with independence in everyday activities. At a minimum, assistance should be required with complex Instrumental Activities of Daily Living (IADL), such as paying bills or managing medications.

- C. The cognitive deficits do not occur exclusively in the context of a delirium
- D. The cognitive deficits are not better explained by another mental disorder (eg, major depressive disorder, schizophrenia)

Causes of Dementia

Most Common Causes of Dementia

Alzheimer's accounts for approximately 60% of all cases of dementia and is therefore the most likely etiology in most cases. The differential diagnosis of dementia is fairly long, however; and other causes need to be considered. New criteria and guidelines identify three stages of Alzheimer's disease:

- 1. preclinical Alzheimer's disease
- 2. mild cognitive impairment (MCI) due to Alzheimer's disease (MCI)
- 3. dementia due to Alzheimer's disease.

These reflect current thinking that changes in the brain begin years before symptoms are apparent.

Vascular dementia (e.g. due to multiple infarcts) accounts for about 15-20% of causes. Patients with this diagnosis usually have cardiovascular risk factors such as hypertension and tobacco use. Approximately 10% of patients with Alzheimer's dementia also have pathologic findings of vascular dementia and may be considered to have mixed dementia.

Dementia with Lewy bodies (DLB) has recently been recognized as a more common cause of dementia than previously thought, accounting for 10% to 15% of cases at autopsy. In addition to typical symptoms of dementia, patients with DLB manifest fluctuations in alertness and attention (delirium), visual hallucinations, and Parkinsonian symptoms.

Less Common Causes of Dementia

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- Parkinson's disease is a cause of dementia late in the course of the disease, with the prominent motor symptoms starting years before. This sequence distinguishes dementia from Parkinson's from DLB, in which the Parkinsonian symptoms and dementia begin simultaneously. Patients presenting with dementia without the classic findings of Parkinson's are unlikely to have Parkinson's disease.
- Frontal temporal dementia, such as Pick's disease, differs from Alzheimer's because, as the name implies, this type of dementia presents with more frontal lobe symptoms (such as changes in personality, demeanor, and behavior).
- Other rare forms include Huntington's disease and Creutzfeldt-Jakob disease. Huntington's disease follows an autosomal dominant pattern and therefore typically has a strong family history. Its initial presentation is characterized by random, jerky and uncontrollable movements (choreiform movements). Creutzfeldt-Jakob disease is an extremely rare, rapidly progressive infectious condition transmitted by a prion.

Ankle Clonus

A series of abnormal alternating contractions and relaxations of the foot induced by sudden dorsiflexion of the foot. Its presence is suggestive of upper motor neuron pathology.

Caregiver Stress

- Caregiver stress has been increasingly recognized as a risk factor for morbidity and mortality. For example, a prospective
 study demonstrated that older spousal caregivers who report emotional strain have a relative risk of mortality of 1.63 over
 four years, compared to non-caregivers. Depression, anxiety, and use of psychotropic medications have also all been
 demonstrated to be more common among caregivers.
- Formal tools to identify caregiver stress have not been developed for primary care settings. As such, direct questioning of the caregiver is recommended to identify the level of perceived burden. This should include screening for anxiety and major depression. Additionally, physicians should identify the level and types of external support the caregiver is receiving (e.g. family member or home health aid visits).
- Respite care may be extremely helpful. This can involve either a family member or a paid professional taking care of the disabled patient for a block of time, thereby allowing the caregiver a break for recreation or for the completion of tasks that are difficult with the disabled patient. Enrolling the patient in an adult day program is the most structured form of respite.
- Caregiver support groups may also be helpful. Such groups allow caregivers to share experiences, challenges and solutions with other people in similar circumstances.
- Knowing how to find resources for disabled patients and their caregivers is an essential tool for the family physician. One
 such resource is the <u>Family Caregiver Alliance</u>, which maintains an online listing of respite, home care, palliative care, and
 caregiver support services by state. The US Department of Health and Human Services also maintains a listing at
 www.eldercare.gov.

Financing Home and Long-Term Care Services

Skilled home services

Medicare and most private insurances will cover a short stint of skilled services at home. Skilled services are those provided by licensed providers such as nurses, physical therapists, speech therapists, etc...

Other home services

Home care agencies also provide home health aides, who help with ADLs such as bathing and dressing; homemakers, who assist with some IADLs such as laundry, light housework, and errands; and companions, who provide sitting and respite services. These types of services are not covered by most insurance plans, and generally cost between 15 and 30 dollars an hour. To pay for them, some patients who own their own houses enter into a reverse mortgage, in which the home care is paid for gradually from the value of the house, all the while allowing them to stay at home.

Long-term care such as nursing home or adult day program

If there is a need for skilled services, Medicare will cover 100% of the cost for the first 20 days of a nursing home stay after a hospitalization.

Long-term care is not covered at all, however. Adult day programs cost anywhere from 50 to 100 dollars a day, while nursing homes cost between 150 and 250 dollars a day. Most states have payment assistance programs to help families that cannot afford these rates, but still the costs can be prohibitive. It is not uncommon for a family to work through their entire life savings paying for long-term care. Eventually, they qualify for Medicaid, which picks up the remaining cost. Nowadays, many families are considering purchasing separate long-term care insurance policies that cover these types of services.

Primitive Reflexes

Patients with advanced dementia or other neurodegenerative diseases may develop primitive reflexes with loss of frontal lobe control. Primitive reflexes are those that may be seen in infants, including grasp, suck, and the "glabellar tap" (patient continues to blink each time he or she is tapped on the forehead; healthy patients stop blinking after the second or third tap).

Management

UTI Treatment

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UTI Treatment

Treat UTI with IV ceftriaxone after making certain that the urine sample gets sent for gram stain and culture.

Delirium Treatment

Delirium Treatment

Prescribe a short course of oral haloperidol, which is helpful in aiding sleep, diminishing agitation, and clearing hallucinations.

Some providers prefer newer atypical antipsychotics such as olanzapine and quetiapine, which have fewer extrapyramidal side effects. There is a recent systematic review of the efficacy and safety of the off-label use of atypical antipsychotic medications for various conditions. For symptoms such as psychosis, mood alterations, and aggression associated with dementia in elderly patients, small but statistically significant benefits were observed for aripiprazole, olanzapine, and risperidone. However, these agents can lead to a prolonged QT interval and don't have the long track record of haloperidol.

Side effects of haloperidol:

- Somewhat sedating and can cause constipation.
- Tardive dyskinesia involuntary spasms of the neck, tongue and lips. Generally seen in patients on higher doses of this medication for a long time (such as over a year).
- QT prolongation rare.

Hospital Interventions That Can Prevent or Minimize Delirium

Providing a safe environment for agitated and combative patients is a challenge for physicians and nurses. Frequent reorientation and redirection by a provider who is familiar with the patient can be very effective in reducing agitation. Nurses on a regular hospital ward typically have responsibilities for too many patients to redirect effectively a delirious patient. A one-to-one sitter provides a safe alternative that can minimize the use of physical and chemical restraints. Repeated reorientation, by a sitter, family member, nurse, or doctor can help reduce confusion. Review of a multifaceted hospital-unit-wide intervention (compared with two "usual care" units) showed a reduction in the number and duration of episodes of delirium. One element of the intervention was an orientation protocol that included frequent cognitive stimulation and active orientation.	
As you recall from earlier in the care of this patient, a wide variety of medications can lead to delirium. Sedati medications and those with anticholinergic side effects (e.g. narcotics, benzodiazepines, and tricyclic antidepressants) should generally be avoided in patients at risk of delirium, unless they are absolutely necess (e.g., narcotics for post-operative pain). The updated American Geriatrics Society Beers Criteria is an evidence based review of potentially inappropriate medications for older adults and providers should consider avoiding drugs on this list.	
Providing early mobility and range of motion exercises to bedridden patients can help prevent delirium. Such exercises provide physical touch, cognitive and sensory stimulation, and can prevent decubitus ulcers.	
Foley catheters, while preventing urinary incontinence in bed, can be uncomfortable and provide noxious stimuli that can augment delirium. Additionally, these lines and wires can lead to preventable falls. The same may be said for oxygen tubing, IV lines, telemetry monitors, and, particularly, physical restraints. All of these may be used when there is no alternative, but they should be discontinued as soon as safely possible.	
Placing a patient at the end of the hall where it is dark and quiet at night would not be helpful. Mounting evidence suggests that sensory deprivation is a contributing factor to delirium in the hospital. Interventions that increase, rather than minimize, stimulation seem effective at preventing or minimizing delirium. As such, keeping patients in rooms with adequate lighting, windows, large clocks, and calendars helps keep the patient calm and more oriented. Keeping the patient close to the activity of the nursing station is also helpful. Visual and hearing disturbances can contribute to delirium. Dim lighting, ambient noise, and the removal of hearing and visual aids therefore should be avoided.	

Treatment of Alzheimer's Dementia & Symptoms

Four cholinesterase inhibitors (donepezil, rivastigmine, tacrine and galantamine) show statistically significant, though clinically small benefits for patients with mild to moderate dementia. Studies have demonstrated improvements in MMSE, ADL, and IADL scores, but the improvements are small enough that most families and clinicians will be unable to detect them. A recent meta-analysis demonstrated that a physician would need to treat 12 patients with mild to moderate Alzheimer's dementia for one to achieve minimal improvement or better,

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and for one to have a treatment-related side effect. A systematic review in 2008 concluded that cholinesterase inhibitors lead to statistically significant but "clinically marginal" effects. The use of vitamin E in the treatment of dementia is controversial. One large trial (the Alzheimer's Disease Cooperative Study) of 2000 IU daily demonstrated a delay in the onset of death, institutionalization, or progression to severe dementia of 670 days compared to placebo among patients with moderate to severe dementia. Due to this study, the American Academy of Neurology recommends the use of Vitamin E in the Vitamin E management of dementia. Concern has arisen from a recent meta-analysis of vitamin E supplementation that demonstrated a rise in all-cause mortality. A 2012 Cochrane Collaboration review concluded that there was no convincing evidence that Vitamin E is of benefit in the treatment of Alzheimer's dementia. Clinicians should discuss the potential risks and benefits with patients and their families on an individual basis. Memantine (Namenda) is an N-methyl d-aspartate (NMDA) receptor antagonist that is FDA-approved for use in moderate to severe dementia. Randomized trials suggest that, like the cholinesterase inhibitors, memantine leads to small but statistically significant improvements in cognition. The clinical relevance of the improvements Memantine is not clear, but the medication is generally well tolerated and there are few other options. It may be used in combination with cholinesterase inhibitors in patients with severe dementia. A 2011 systematic review found no evidence that memantine is effective in mild Alzheimer's and only meager evidence of its efficacy in moderate Alzheimer's. For community dwelling patients with dementia, providing support and respite to their caregivers is essential to their care. Respite care may take a variety of forms. For a family with ample local support, simply having family members take turns replacing the primary caregiver is inexpensive and effective. Patients may pay to have a home health aid come at various intervals. Such aids can provide assistance with specific non-medical tasks Respite such as bathing, shopping, and laundry, or they may simply allow the primary caregiver to leave for a while. The most structured form of respite is an adult day program. Such programs provide all the services available in a nursing home (e.g., nutrition, nursing, medication management, recreation) but only during work hours on weekdays. Such programs may allow a patient with advanced dementia to delay long-term institutionalization. Behavioral disturbance (such as aggression or psychosis) is a common and difficult problem as dementia progresses. Careful identification of exacerbating factors can lead to specific approaches to alleviate the problem (e.g. recognition and treatment of constipation leading to agitation). However, medications often have a role in the management of behavior. The atypical antipsychotics olanzapine (Zyprexa) and risperidone (Risperdal) appear to have a modest benefit in managing such neuropsychiatric symptoms and are commonly used. Two recent studies have suggested that patients given atypical antipsychotics have a slightly higher risk Atypical of death. In 2006 the FDA placed "black box" warnings on the labeling for these medications and the antipsychotics conventional antipsychotics (such as haloperidol) indicating the increased risk of death among elderly patients with dementia. Because of this, it is important to attempt behavioral treatments first and to openly discuss the goals and risks of treatment with patients' families prior to starting these medications. For some families, managing severe behavioral symptoms using a medication with a known mortality risk is preferable to the alternative of living with the symptoms. Additionally, providers should understand that the increased mortality concerns come from data about the long-term use of antipsychotics in chronic behavioral disturbance. The safety of these medications in the acute management of delirium should be considered separately.

Cognitive rehabilitation therapy has not been demonstrated to be helpful in slowing the progression of Alzheimer's dementia. A recent Cochrane Collaboration systematic review published in 2003 concluded that there was no evidence to support its use, though it pointed out that the number of existing randomized trials is limited.

A 2007 Cochrane Collaboration systematic review concluded that the evidence for gingkgo biloba's role in the management of dementia is "inconsistent and unconvincing."

Asymptomatic Bacteriuria

Asymptomatic bacteriuria is very common in older patients. There is no evidence that treating it helps. As many as 50% of older men will have a positive urine culture within six weeks of treatment for a UTI. As such, there is no need for a test of cure. If an older patient has a UTI, and then has similar symptoms again, he will need a urine culture at that point.

Studies

Dementia - Diagnostic Testing & Neuroimaging

Role of diagnostic testing

The role of diagnostic testing in the setting of a patient with dementia is primarily to rule out treatable causes, which occur in less than 1% of cases of dementia.

- Depression is commonly comorbid with dementia symptoms and is widely recognized as a cause of "pseudo-dementia." The <u>Geriatric Depression Scale</u> is a well-validated screening test for depression in older patients that may be used to rule out depression.
- Other tests recommended by leadership groups include:

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- 1. complete blood counts
- 2. basic chemistries
- 3. calcium
- 4. thyroid
- 5. vitamin B12 deficiency testing
- 6. Folate deficiency can cause dementia, but is now extremely rare in North America due to the widespread fortification of grain products with folate. Testing for folate deficiency is therefore no longer recommended.
- 7. Syphilis testing may be performed if there is a clinical suspicion or if the local prevalence is high.

Role of neuroimaging

Neuroimaging with either noncontrast computed tomography (CT) or magnetic resonance imaging (MRI) has not been clearly demonstrated to be beneficial in the diagnosis of dementia, but can rule out structural defects (e.g., tumors or normal pressure hydrocephalus) and vascular causes. The American Academy of Neurology therefore recommends the use of either one of these tests in patients for whom these diagnoses are a concern. Currently, no guidelines recommend the use of functional MRI, positron emission tomography (PET) scans, or single photon emission computed tomography (SPECT) scans in the diagnosis of dementia.

Non-Specific White Matter Changes on MRI

Non-specific white matter changes are a common finding on MRIs of the brain. They represent microvascular disease, which may be caused by hypertension, smoking, obesity or simply by aging. They are commonly found in asymptomatic individuals and are often felt to be clinically irrelevant.

The Clinical Dementia Rating

The Clinical Dementia Rating or CDR was developed for the evaluation of the severity of dementia, primarily for use in persons with dementia of the Alzheimer type (the equivalent of probable Alzheimer's disease). It can also be used to stage dementia in other illnesses as well. The protocol is based on interviews of the patient and caregiver. The areas tested are memory, orientation, judgment, community affairs, home and hobbies, and personal care. The CDR is a five-point scale:

	0	No cognitive impairment
	0.5	Very mild dementia
	1	Mild dementia
	2	Moderate dementia
	3	Severe dementia

Diagnosing Delirium

The **Confusion Assessment Method (CAM)** has been widely used over two decades and is the predominant tool used in clinical research regarding delirium.

A recent systematic review of assessment tools in diagnosing the presence of delirium demonstrated that the best evidence supports use of the CAM, which takes five minutes to administer. The CAM has a summary-positive likelihood ratio of 9.6 (95% CI, 5.8 - 16.0) as a bedside delirium instrument.

The MMSE, which has been widely used clinically during the same time period, is a multidimensional tool to assess cognitive function, and was not developed specifically for the assessment of delirium. In the same study mentioned above, the MMSE was the least useful in identifying a patient with delirium (likelihood ratio, 1.6; 95% CI, 1.2 - 2.0).

A newer test, the Montreal Cognitive Assessment (MoCA), has been proposed to be a better alternative than the MMSE to assess cognitive function.

Functional Status Assessment

Functional status assessment is traditionally divided into evaluation of Activities of Daily Living (ADL, or skills required for basic living) and Instrumental Activities of Daily Living (IADL, or skills required for living independently).

ADLs	IADLs
• bathing	• shopping
• dressing	preparing meal
transferring	using the telephone

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• continence	managing transportation needs
• toileting	managing medications
• feeding	managing finances
, ,	These don't have to be done on a daily basis, are not as severe a threat to independent living, and other people can be hired to do them. They typically require a cognitive component in addition to physical ability.
They are usually acquired by the first time one leaves home (about 5 or 6 years old, or kindergarten age).	They are usually acquired by the second time one leaves home (about 16-17 years old, beginning college or career).
	Over 50% of people over the age of 75 have at least one IADL deficit.

Screening for both ADLs and IADLs includes asking the patient if he/she has any problems with each activity; and, if so, who helps? Ask: "Do you have any problems with...? If you have problems, then who does it for you?" Some providers use a standardized form to assess a patient's functional status. (See example.)

MMSE Calculation

Scoring the MMSE should be familiar to most clinicians due to its wide use. The U.S. Preventive Services Task Force (USPSTF) concludes that the evidence is insufficient to recommend for or against routine screening for dementia in older adults (Grade "I" recommendation). It should be noted that in the case of Mr. Marshall, you are not screening. He has symptoms of a change in mental status, so you are doing the MMSE for diagnosis.

Providers commonly perform the attention and calculation part of the MMSE incorrectly. Providers are intended to ask *all* patients to perform the serial 7s. Those who do not correctly complete this entire sequence (93, 86, 79, 72, 65) should be asked to spell "WORLD" backwards. The scorer then uses the better of the two scores. Using the better performance accommodates for the fact that some people are naturally more gifted with literacy than numeracy, and vice versa.

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Pneumonia on Chest X-ray

Pneumonia typically appears on chest x-ray as a density in one of the lung fields.

When a consolidated pneumonia rests up against solid tissue, such as the diaphragm, you lose the ability to distinguish the border of that tissue from the lung tissue.

Sometimes air-bronchograms are visible in the infiltrate. Air bronchograms are I inear lucencies within an infiltrate that represent the air within a bronchus that is surrounded by consolidated lung tissue. These are pathognomonic for an airspace illness such as pneumonia.

Particularly in older patients, patients with atypical pneumonia such as Legionella, and patients early in their illness, pneumonia may not be seen on an x-ray. If there is a high clinical suspicion of pneumonia, current guidelines would suggest starting pneumonia treatment and repeating a chest x-ray in 24-48 hours.

Clinical Reasoning

Distinguishing Between Dementia, Delirium, and Depression

Dementia	 Dementia is an acquired syndrome of gradual progressive deterioration in global intellectual ability that interferes with the ability to function in social and occupational roles. Important elements of the definition include: acquired (i.e., not congenital), progressive (i.e., worse over time), global (not isolated to memory), and interfering with function.
Delirium	 Delirium also causes a decline in cognition, but is particularly notable for acute disturbances in attention (e.g., an inability to shift focus), alertness (e.g., impaired consciousness and sleep cycles) and perception (e.g., hallucinations). In delirium, the symptoms fluctuate over short periods of time, whereas in dementia the symptoms are slowly
	 Depression is an alteration in mood, which can be confused with dementia in older patients. It frequently causes a decline in the ability to concentrate, which may worsen memory. Depression also leads to a lack of

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Depression

interest and energy, which may appear similar to the symptoms of dementia. Since depression is readily treatable, it is essential to diagnose this condition in patients who present with memory loss.

• In addition to representing the primary diagnosis, depression can be comorbid with dementia, and should be considered both because it may worsen symptoms related to dementia, and because it can be reversible.

Differential Diagnosis of Change in Mental Status in a Patient with Dementia

Most Likely Diagnoses

Causes of delirium:

Causes of delirium:		
Urinary tract infection (UTI)	UTIs are among the most common causes of delirium in older patients. Such patients are seldom able to identify the common symptoms of UTI's such as dysuria or frequency, so the absence of these symptoms does not rule out this diagnosis. UTIs also may cause or exacerbate urinary incontinence.	
Respiratiory infection	Respiratory infections (e.g. pneumonia) frequently complicate dementia and often cause delirium in such patients. The mortality rate of patients with advanced dementia is high and is primarily attributable to infections such as pneumonia and urosepsis. Aspiration pneumonia may also cause delirium.	
Electrolyte disturbance	A wide range of electrolyte disturbances may cause delirium (e.g. hypo or hypernatremia, hypercalcemia, hypokalemia, and metabolic acidosis). These conditions are easily detected and highly treatable, so they should always be considered in the work-up of acute change in mental status and delirium.	
Urinary retention	Any condition that leads to discomfort may cause delirium in a patient with dementia. Acute urinary retention leads to the complete inability to pass urine. Chronic urinary retention (most commonly due to benign prostatic hypertrophy [BPH]), however, leads to incomplete emptying and distention of the bladder, which further may lead to UTI and overflow incontinence. Risk factors for urinary retention include male sex, age over 70 and BPH. When this is a consideration, a post void residual should be measured (either by catheterization or ultrasound assessment of bladder volume after voiding). Catheterization allows the collection of a urine culture that is unlikely to be contaminated by skin flora. When urinary retention is severe, the bladder may be palpated in the suprapubic area.	
Pain	Pain from any source may cause delirium among patients with dementia. This presents a clinical challenge, since the treatment of pain with opiates also can cause delirium. Since patients in this condition are frequently unable to localize and report their symptoms, it is important to do a full physical exam looking for hidden sources of pain (abdominal tenderness, joint or bony injury).	
Depression	Depression and other psychiatric conditions can present with altered mental status in older patients; however, psychiatric conditions usually present dif ferently from delirium.	
Withdrawal	Withdrawal from medications or other substances is a common cause of delirium. This reaction to withdrawal occurs most frequently from alcohol and benzodiazepines. In these cases, the delirium may progress to a more severe form known as delirium tremens, characterized by tachycardia, hypertension, delirium with agitation, visual hallucinations and formication (the sensation of insects crawling on the patient).	
Acute cerebrovascular events	Acute cerebrovascular events such as ischemic stroke or intracranial bleeding may present with an acute change in mental status. This is most likely in a patient with vascular dementia who experiences a new ischemic event. In this case, the vascular event could be small enough to not cause new neurologic symptoms. In combination with the other pre-existing insults, however, the new event may cause an acute alteration in mental status. Large brain insults (e.g. those due to larger ischemic events or acute hemorrhage) typically have associated neurological findings on physical exam (e.g. hemiplegia or upper motor neuron findings - such as hyperreflexia, clonus, or positive Babinski's).	
Adverse drug effects	As a group, adverse drug effects represents one of the most common causes of delirium in older patients. Patients with dementia, in particular, are sensitive to medications with sedating effects. Medications with anticholinergic effects (opiates, benzodiazepines, sedating antihistamines, tricyclic antidepressants, antipsychotics, and some anti-nausea medications) are the drugs most likely to cause delirium. Other potential offenders include certain antibiotics (e.g. fluoroquinolones), beta-blockers, and H2-blockers.	

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