Medicines and Falls in Hospital

All patients should have their drug burden reviewed with respect to its propensity to cause falls. The history should establish the reason the drug was given, when it started, whether it is effective and what its side effects have been.

An attempt should be made to reduce the number and dosage of medications and ensure they are appropriate, and not causing undue side effects.

Falls can be caused by almost any drug that acts on the brain or on the circulation. Usually the mechanism leading to a fall is one or more of:

- sedation, with slowing of reaction times and impaired balance,
- hypotension, including the 3 syndromes of paroxysmal hypotension OH, VVS and VD-CSH
- bradycardia, tachycardia or periods of asystole

Falls may be the consequence of recent medication changes, but are usually caused by medicines that have been given for some time.

Red: High risk: can commonly cause falls alone or in combination Amber: Moderate risk: can cause falls, especially in combination Yellow: Possibly causes falls, particularly in combination

Drugs acting on the brain (aka psychotropic drugs)

There is good evidence that stopping these drugs can reduce falls (1).

Taking such a medicine roughly doubles the risk of falling. There is no data on the effect of taking two or more such tablets at the same time. (2)

Sedatives, antipsychotics, sedating antidepressants cause drowsiness and slow reaction times. Some antidepressants and antipsychotics also cause orthostatic hypotension.

| Sedatives: | Temazepam, Nitrazepam | Drowsiness, slow reactions, |
|--------------------------|-------------------------------|------------------------------------|
| Benzodiazepines | Diazepam, Lortemazepam | impaired balance. |
| | Chlordiazepoxide, Flurazepam, | Caution in patients who have |
| | Lorazepam, Oxazepam, | been taking them long term |
| | Clonazepam | |
| Sedatives: "Zs" | Zopiclone, Zolpidem | Drowsiness, slow reactions, |
| | | impaired balance. |
| Sedating antidepressants | Amitriptyline, Dosulepin | All have some alpha blocking |
| (tricyclics and related | Imipramine, Doxepin | activity and can cause orthostatic |
| drugs) | Clomipramine, Lofepramine, | hypotension. |
| | Nortriptyline, Trimipramine | All are antihistamines and cause |
| | | drowsiness, impaired balance and |
| | Mirtazapine, Mianserin | slow reaction times. |
| | Trazodone | Double the rate of falling |
| Monoamine Oxidase | Phenelzine, Isocarboxazid, | MAOIs are little now used; all |
| Inhibitors | Tranylcypromine | (except moclobemide) cause |
| | | severe OH |
| Drugs for psychosis and | Chlorpromazine, Haloperidol, | All have some alpha receptor |

| Agitation | Eluphonazina Picnoridana | blocking activity and can cause | | | |
|--|---|--|--|--|--|
| Agitation | Fluphenazine, Risperidone Quetiapine, Olanzapine | blocking activity and can cause orthostatic hypotension. | | | |
| | Quetiapine, Olanzapine | Sedation, slow reflexes, loss of | | | |
| | | balance. | | | |
| CCDI entidorenegante | Controlino, Citologuera | | | | |
| SSRI antidepressants | Sertraline, Citalopram, | Cause falls as much as other | | | |
| | Paroxetine, Fluoxetine | antidepressants in population | | | |
| | | studies. | | | |
| | nave shown that SSRIs are consisten | • | | | |
| rate of falls and fractures, but there are no prospective trials. The mechanism of such an effect is | | | | | |
| - | nd bradycardia only rarely as an idio | osyncratic side effect. They do not | | | |
| normally sedate. They impa | | | | | |
| SNRI antidepressants | Venlafaxine, Duloxetine | As for SSRIs but also commonly | | | |
| | | cause orthostatic | | | |
| A combination of an SSRI | | hypotension (through | | | |
| and a noradrenaline re- | | noradrenaline re-uptake | | | |
| uptake inhibitor | | blockade) | | | |
| Opiate analgesics | All opiate and related analgesics | Sedate, slow reactions, impair | | | |
| | – Codeine, Morphine, Tramadol | balance, cause delirium, | | | |
| Anti-epileptics | Phenytoin | Phenytoin may cause permanent | | | |
| | | cerebellar damage and | | | |
| | | unsteadiness in long term use at | | | |
| | | therapeutic dose. | | | |
| | | Excess blood levels cause | | | |
| | | unsteadiness and ataxia. | | | |
| | Carbamazepine | Sedation, slow reactions. Excess | | | |
| | Phenobarbitone, | blood levels cause unsteadiness | | | |
| | | and ataxia. | | | |
| | Sodium valproate, Gabapentin | Some data on falls association. | | | |
| | Lamotrigine, Pregabalin | Insufficient data to know if these | | | |
| | Levatiracetam, Topiramate, | newer agents cause falls | | | |
| Parkinson's disease: | Ropinirole, Pramipexole | May cause delirium and OH | | | |
| Dopamine agonists | | | | | |
| MAOI-B inhibitors | Selegiline | Causes OH | | | |
| The subject of drugs and fal | ls in PD is difficult, as falls are so co | mmon, and OH is part of the | | | |
| disease. In general only defi | nite drug related OH would lead to | a change in medication | | | |
| Muscle relaxants | Baclofen, Dantrolene | Sedative. Reduced muscle tone. | | | |
| | . Tend to be used in conditions ass | | | | |
| Vestibular sedatives | Prochlorperazine | Dopamine antagonist – may cause | | | |
| Phenothiazines | | movement disorder in long term | | | |
| | | use. Alpha receptor blocker and | | | |
| | | antihistamine. | | | |
| Vestibular sedatives | Cinnarazine, Betahistine | Sedating. No evidence of benefit | | | |
| Antihistamines | | in long term use. | | | |
| Sedating Antihistamines | Chlorphenamine, Hydroxizine, | No data, but sedation likely to | | | |
| for allergy | Promethazine, Trimeprazine | contribute to falls. Long half lives. | | | |
| Anticholinergics acting on | Oxybutinin, Tolterodine, | No data, but have a known CNS | | | |
| the bladder | Solifenacin | effects | | | |
| | Somenacin | CHECIS | | | |

Drugs acting on the heart and circulation

Maintaining consciousness and an upright posture requires adequate blood flow to the brain. This requires an adequate pulse and blood pressure. In older people a systolic BP of 110mmHg or below is associated with an increased risk of falls.

Any drug that reduces the blood pressure or slows the heart can cause falls (or feeling faint or loss of consciousness or "legs giving way") (3). In some patients the cause is clear – they may be hypotensive, or have a systolic drop on standing. Others may have a normal blood pressure lying and standing, but have syncope or pre-syncope from carotid sinus hypersensitivity or vasovagal syndrome. Stopping cardiovascular medication reduces syncope and falls by 50%, and reduces the prevalence of these 4 syndromes (4, 5).

| Alpha receptor blockers | Doxazosin, Indoramin, Prazosin, Tamsulosin, Terazocin, Alfluzosin | Used for hypertension or for prostatism in men. They commonly cause severe orthostatic hypotension. Stopping them may precipitate |
|---|--|--|
| | Sedating antidepressants Drugs for psychosis and agitation | urinary retention in men. See above. Orthostatic hypotension. |
| Centrally acting alpha 2 receptor agonists | Clonidine, Moxonidine | May cause severe orthostatic hypotension. Sedating |
| Thiazide diuretics | Bendroflumethiazide, Chlorthalidone, Metolazone | Cause OH, weakness due to low potassium. Hyponatraemia |
| Loop diuretics | Furosemide, Bumetanide | Dehydration causes hypotension. Low potassium and sodium |
| Angiotensin converting enzyme inhibitors (ACEIs) | Lisinopril, Ramipril, Enalapril, Captopril, Perindopril | These drugs rely almost entirely on the kidney for their elimination and can accumulate in dehydration or renal failure. |
| | Fosinopril, Trandolapril, Quinapril | Excreted by liver and kidney |

Symptomatic hypotension in systolic cardiac failure

- ACEIs and beta blocker have a survival benefit in systolic cardiac failure and should be maintained whenever possible.
- NICE recommends: stop nitrates, calcium channel blockers and other vasodilators. If no evidence of congestion, reduce diuretics. If problem persists, seek specialist advice.
- The mortality risk from a fall at age 85 is about 1% per fall. The frequency of falls determines the balance between risk and benefit.
- Most cardiac failure in older people is diastolic (preserved left ventricular function). ACEIs and beta blockers have little survival benefit in diastolic failure.

| Angiotensin receptor blockers (ARBs) | Losartan, Candesartan, Valsartan, Irbesartan, Olmesartan, Telmesartan, Eprosartan | May cause less OH then ACEIs. Excreted by liver and kidney. |
|---|--|---|
| Beta blockers | Atenolol, Sotalol - Renally excreted. May accumulate Bisoprolol, Metoprolol, Propranolol, Carvedilol, Timolol eye drops | Can cause bradycardia, hypotension, CSH, OH and VVS |
| Antianginals | GTN | A common cause of syncope due to sudden BP drop |
| | Isosorbide mononitrate, Nicorandil | Cause hypotension and paroxysmal hypotension |
| Calcium channel blockers that only reduce blood pressure | Amlodipine, Felodipine, Nifedipine, Lercanidipine | |
| Calcium channel blockers which slow the pulse and reduce BP | Diltiazem, Verapamil | May cause hypotension or bradycardia |
| Other antidysrhythmics | Digoxin, Amiodarone, Flecainide | May cause bradycardia and other arrhythmias. Data on digoxin and falls probably spurious due to confounding by indication |
| Acetylcholinesterase inhibitors (for dementia) | Donepezil, Rivastigmine, Galantamine | Cause symptomatic bradycardia and syncope |

 Campbell AJ, Robertson MC, Gardner MM, Norton RN, Buchner DM. Psychotropic medication withdrawal and a home-based exercise program to prevent falls: a randomized, controlled trial. J Am Geriatr Soc 1999; 47: 850–3.
Darowski A, Chambers SCF and Chambers DJ. Antidepressants and falls. Drugs and Aging 2009 26 (5) 381-394
Darowski A and Whiting R. Cardiovascular drugs and falls. Reviews in Clinical Gerontology 2011, 21 (2), 170-179
Van der Velde N, van den Meiracker AH, Pols HA, Stricker BH, van der Cammen TJ. Withdrawal of fall-risk-increasing drugs in older persons: effect on tilt-table test outcomes. J Am Geriatr Soc 2007;55:734–739.
Alsop K, MacMahon M. Withdrawing cardiovascular medications at a syncope clinic. Postgrad MJ 2001; 77:403-5.

Dr Adam Darowski, Consultant Physician, Clinical Lead, The FallSafe Project Dr Jeremy Dwight, Consultant Cardiologist Dr John Reynolds, Consultant in Clinical Pharmacology John Radcliffe Hospital, Oxford.

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