Managing Complex Pharmacological Scenarios – Deprescribing and Polypharmacy

2023 Toronto Geriatrics Update Course

Friday November 10th; 1:10 – 2:10 PM EST

Facilitators: Katie Mok & Andrea Hudson





Facilitators

- Katie Mok: BScPhm, ACPR, BCGP (Internal Medicine/Geriatrics Pharmacist, Mount Sinai Hospital)
- Andrea Hudson: B.A Psych, BScPhm, Pharm D, (Pharmacist, Home and Community Care Support Services)
- We have no disclosures





Poll question (#1): What is your health care role?

- a) Family physician
- b) Nurse practitioner
- c) Specialist
- d) Social worker
- e) Occupational therapist
- f) Physiotherapist
- g) Pharmacist
- h) Other allied health
- i) Non-direct patient care





Poll question (#2): How often do you discuss polypharmacy and deprescribing with your patients/clients?

- a) Every visit and chance I get!
- b) If patient/client raises it as a concern
- c) After a hospitalization or new diagnosis
- d) Whenever I prescribe a new medication
- e) Never
- f) Doesn't really apply to me





Learning objectives

- 1. Review key concepts of inappropriate polypharmacy, prescribing cascades and deprescribing.
- 2. Utilize a complex patient case to apply a framework for deprescribing medications.
- 3. Appreciate differences in the opportunities and timeframes of deprescribing that occur as outpatients compared to inpatients.





Polypharmacy

- WHO definition: concurrent use of multiple medications by a patient
 - Typically defined as routine use of <u>five</u> or more medications

Appropriate Polypharmacy

- Medications have clear indications
- Medications are safe in older adults
- Aligned with goals of care

Inappropriate Polypharmacy

- No evidence based indication
- Medication does not achieve therapeutic objectives
- Cause of adverse drug reactions





Polypharmacy





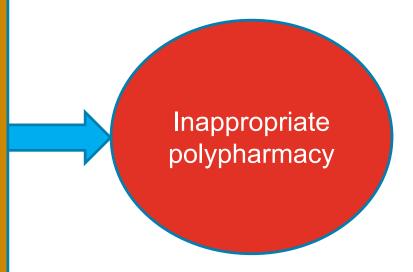
Canadian Institute for Health Information. Drug use among seniors in Canada. Accessed October 17, 2023





Risk factors for inappropriate prescribing

- Age >62 (homeostasis changes, organ dysfunction)
- Cognitive impairment
- Frailty
- Multiple chronic conditions
- Mental Health conditions
- Lack of primary care
- Multiple prescribers (primary care, specialists)
- Acute illness
- Poor care transitions







Age-related changes impacting medications

Changes in physiology:

- ↑ body fat
- ↓ body water
- ↓ albumin
- ↓ liver metabolism

Pharmacodynamic changes:

- Receptor binding
- Number or receptors and activity
- Increased or decreased efficacy
- Increased toxicity

Pharmacokinetic changes:

- Absorption
- Distribution
- Metabolism
- Excretion

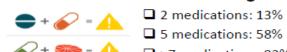
Drenth-van Maanen AC et, Wilting al I,. Br J Clin Pharmacol. 2020 Oct;86(10):1921-1930.





Consequences of inappropriate polypharmacy

↑ Risk Adverse Drug Events



☐ 5 medications: 58%



Prescribing cascade



Unnecessary drug expenses



ED visits







Functional decline



↑ Mortality risk

Zed et al. CMAJ 2008;178:1563-1569; Bourgeois et al. Pharmacoepidemiol Drug Saf 2010;19:901-910; Gnjidic D et al. J Clin Epidemiol 2012;65(9):989-95; Maher RL, et al. Expert Opin Drug Saf 2014;13(1):57-65; Patterson SM et al. Cochrane Database Syst Rev 2014;(10): CD008165; Johansson T et al. Br J Clin Pharmacol 2016;82:532-548





WELL, THE WHITE PILL LOWERS MY BLOOD PRESSURE BUT MAKES MY LEGS SWELL, THE YELLOW PILL LOWERS THE SWELLING BUT CAUSES ME TO PEE, THE BLUE PILL STOPS ME FROM PEEING BUT MAKES ME CONFUSED, THE TAN PILL IMPROVES MY MEMORY BUT MAKES MY NOSE FROM RUNNING BUT MAKES ME SLEEPY, THE ORANGE PILL WAKES ME UP BUT INCREASES MY BLOOD PRESSURE, SO THE WHITE PILL LOWERS MY BLOOD PRESSURE BUT...



By Edwin Tan (c) 2015 www.facebook.com/edsrant

Prescribing cascades

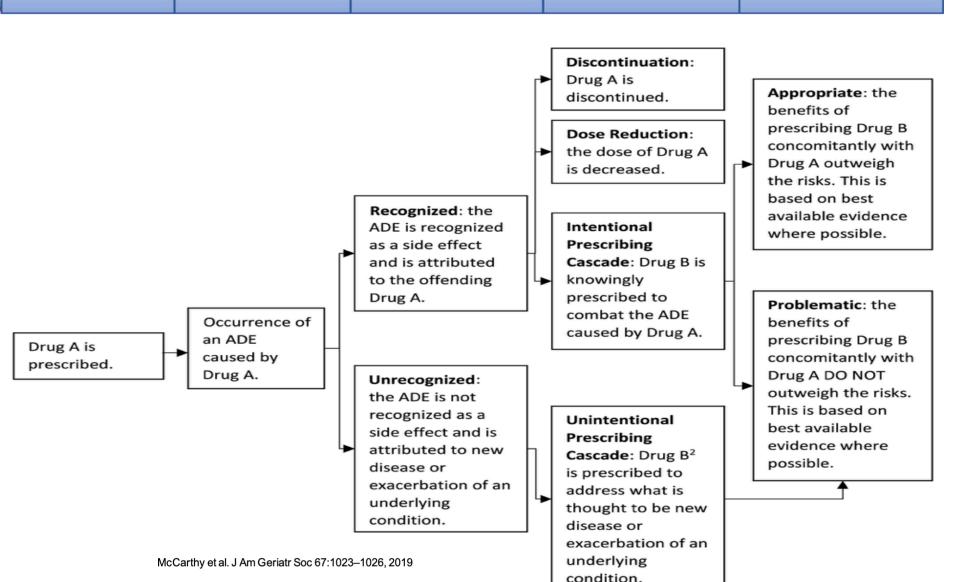
PRESCRIPTION of DRUG

ADE¹
OCCURRENCE

CLINICIAN'S INTERPRETATION of the ADE

CLINICIAN'S ACTION

APPROPRIATENESS of the PRESCRIBING CASCADE



Poll question (#3): Which of the following is an example of a possible inappropriate prescribing cascade?

- a) pregabalin --> edema --> diuretic
- b) levodopa-carbidopa --> hallucinations --> quetiapine
- c) over the counter NSAID --> GERD --> PPI
- d) all of the above



Prescribing cascade examples

Drug A	Side effect	Drug B
Appropriate cascade		
methotrexate	Mouth sores, folate deficiency	folic acid
opioid	constipation	laxative
Inappropriate cascade		
calcium channel blocker	peripheral edema	diuretic
diuretic	urinary incontinence	overactive bladder medication
NSAID	hypertension fluid retention	antihypertensive diuretic





Strategies to reduce potential harms of polypharmacy

- 1. Assess risk of polypharmacy
- 2. Regular medication review
- 3. Inform caregivers of medication changes
- 4. Choose medications with fewest side effects
- 5. DEPRESCRIBE- Stop or reduce unnecessary medication
- Consider impact of medications on QOL
- 7. Consider ability to take and remember to take medications
- 8. Multidisciplinary case conferencing

Medication Safety in Polypharmacy. Geneva: World Health Organization; 2019 (WHO/UHC/SDS/2019.11).





What can patients, families and caregivers do?

- Bring someone to appointments
- List questions before appointments
- Inform about over-the-counter medications, supplements and herbal products
- Report any new or unexpected symptoms
- Ask questions about your medication (prescribers and pharmacists)
- Use one pharmacy

Medication Safety in Polypharmacy. Geneva: World Health Organization; 2019 (WHO/UHC/SDS/2019.11).





QUESTIONS TO ASK ABOUT YOUR MEDICATIONS

when you see your doctor, nurse, or pharmacist.

1. CHANGES?

Have any medications been added, stopped or changed, and why?

2. CONTINUE?

What medications do I need to keep taking, and why?

3. PROPER USE?

How do I take my medications, and for how long?

4. MONITOR?

How will I know if my medication is working, and what side effects do I watch for?

5. FOLLOW-UP?

Do I need any tests and when do I book my next visit?



Remember to include:

- √ drug allergies
- ✓ vitamins and minerals
- herbal/natural products
- all medications including non-prescription products

Ask your doctor, nurse or pharmacist to review all your medications to see if any can be stopped or reduced.

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Deprescribing

"Deprescribing is the <u>planned and supervised</u> process of <u>dose reduction or stopping of</u> <u>medication</u> that might be <u>causing harm</u>, or no <u>longer be of benefit</u>.

Deprescribing is part of good prescribing – backing off when doses are too high, or stopping medications that are no longer needed."

What is deprescribing? deprescribing.org. Accessed Oct 30, 2023





10 step deprescribing algorithm (British Medical Journal Group)

1. Accurately ascertain all current drug use · 'brown paper bag' medication reconciliation 2. Identify patients at risk of, or suffering, ADR All three at-risk criteria – aim for ≤ 5 drugs at risk: ≥8 medications advance high-risl or) assess for curre 3. Estimate life e clinical prognost 4. Define overall consider current S **I**reference 5. Verify current perform diagnos 1 or confirm diagnos · ascertain, for ea Determine nee efit estimate clinical compare this es 7. Determine abs reconcile estima tools (se ses 8. Review the rel rank drugs according to the relative utility from high to low based on Discontinue drugs of low utility predicted benefit, harm, administration and monitoring burden 9. Identify drugs to be discontinued and seek patient consent reconcile drugs for discontinuation with patient preferences Discontinue drugs patients are not in favour of taking

10. Devise and implement drug discontinuation plan with close

monitoring

Poll question (#4): What is the biggest challenge that affects your ability to deprescribe?

- a) Not enough time
- b) Knowledge and skills, access to resources
- c) Patient hesitancy
- d) Lack of information from other providers
- e) Not part of my role





Barriers and Facilitators to deprescribing

	Barrier	Facilitator
Cultural	Prescribing culture Lack of non-pharmaceutical alternatives	More prudent prescribing culture
Organizational/ system	Disease focused evidence-based guidance Lack of guidance on tapering and interactions Incomplete information sharing	Better evidence-based multimorbidities guidance Accessibility of resources and tools
Interpersonal	Fragmented care Prescriber uncertainties (e.g., reluctance to cease medication, concerns of poor outcomes) Lack of time	Improved communication Continuity of care Interdisciplinary collaboration
Individual	Patient may not want to stop medication Cognitive impairment	Patient-centred care Involvement of patient in decision making Decreased pill burden





Identifying potentially inappropriate medications

Resources:

- Beers Criteria updated May 2023
- STOPP/START version 3 updated July 2023
 - 133 STOPP criteria and 57 START criteria
- Medication Appropriateness Index (MAI)



Deprescribing resources/tools

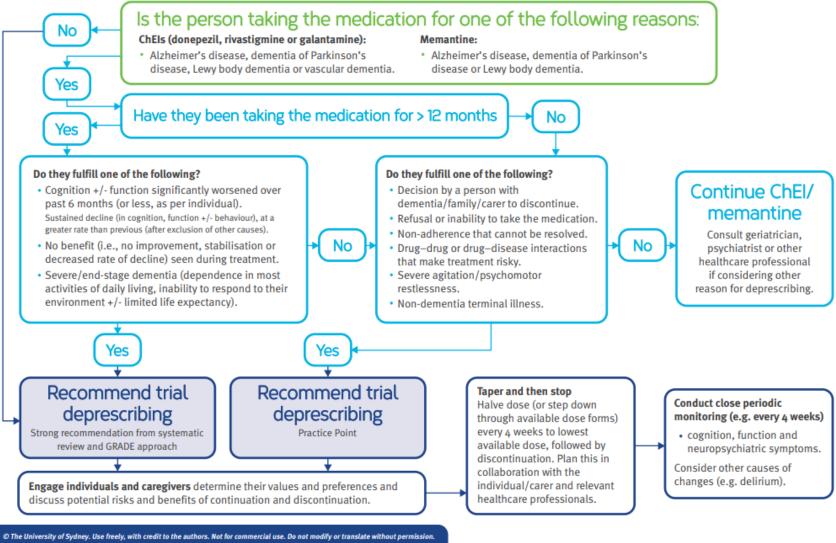
- <u>Deprescribing.org</u>: resources and algorithms (e.g., PPIs, sedative hypnotics, antihyperglycemics, antipsychotics, AChEIs)
- The Canadian Medication Appropriateness and Deprescribing Network: resources, tools and algorithms
- <u>Medstopper.com</u>: prioritization and tapering recommendations
- Centre for Effective Practice: evidence-based tools, resources and programs
- Choosing Wisely: recommendations and toolkits (e.g., Bye-bye PPI)
 for treatments not supported by evidence







deprescribing.org | Cholinesterase Inhibitor (ChEI) and Memantine Deprescribing Algorithm



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Contact emily.reeve@sydney.edu.au for more information.

Reeve E, Farrell B, Thompson W, at al Evidence-based Clinical Practice Guideline for Deprescribing Cholinesterase Inhibitors and Memantine. 2018. ISBN-13: 978-0-6482658-0-1 Available from: http://sydney.edu.au/medicine/cdpc/resources/deprescribing-guidelines.php

















MedStopper is a deprescribing resource for healthcare professionals and their patients.

- **1** Frail elderly?
- Generic or Brand Name:



3 Select Condition Treated:



Generic Name	Brand Name	Condition Treated	Add to MedStopper
gabapentin	Neurontin	Select Condition 🔻	<u>ADD</u>
pregabalin	Lyrica	Select Condition 🔻	<u>ADD</u>
vigabatrin	Sabril	Select Condition 🔻	<u>ADD</u>



◆ Previous Next ▶

MedStopper Plan

Arrange medications by:

Stopping Priority

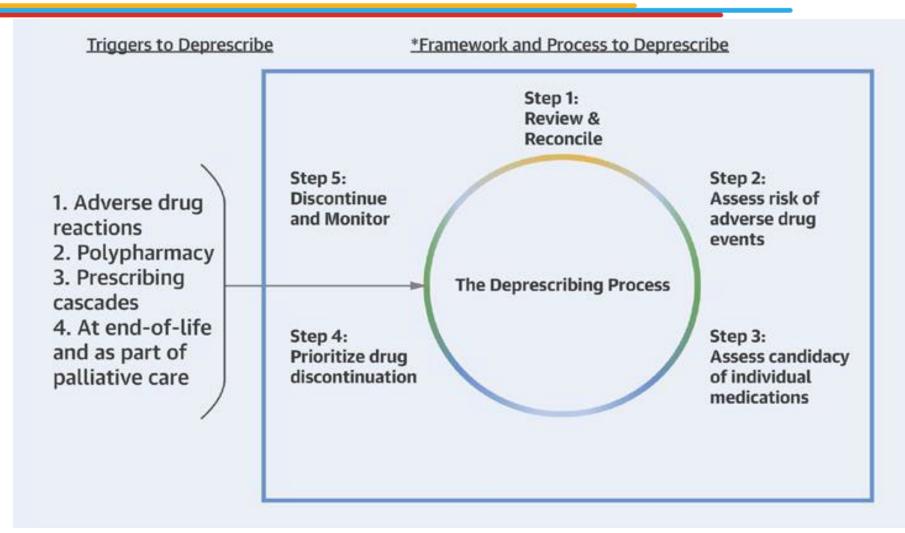
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CLEAR ALL MEDICATIONS

PRINT PLAN

Stopping Priority RED=Highest GREEN=Lowest	Medication/ Category/ Condition	May Improve Symptoms?	May Reduce Risk for Future Illness?	May Cause Harm?	Suggested Taper Approach	Possible Symptoms when Stopping or Tapering	Beers/STOPP Criteria
	duloxetine (Cymbalta) / SNRI / chronic pain	([:)	(<u>;</u>)	(3)	If used daily for more than 3-4 weeks. Reduce dose by 25% every week (i.e. week 1-75%, week 2-50%, week 3-25%) and this can be extended or decreased (10% dose reductions) if needed. If intolerable withdrawal symptoms occur (usually 1-3 days after a dose change), go back to the previously tolerated dose until symptoms resolve and plan for a more gradual taper with the patient. Dose reduction may need to slow down as one gets to smaller doses (i.e. 25% of the original dose). Overall, the rate of discontinuation needs to be controlled by the person taking the medication.	nausea, diarrhea, abdominal pain, sweating, headache, dizziness, cold and flulike symptoms, anxiety, irritability, trouble sleeping, unusual sensory experiences (e.g. electric shock-like feelings, visual after images), sound and light sensitivity, muscle aches and pains, chills, confusion, pounding heart (palpitations), unusual movements, mood changes, agitation, distress, restlessness, rarely suicidal ideation	None
	gabapentin (Neurontin) / Antiepileptic / pain	(I:)	(;)	(3)	If used daily for more than 3-4 weeks. Reduce dose by 25% every week (i.e. week 1-75%, week 2-50%, week 3-25%) and this can be extended or decreased (10% dose reductions) if needed. If intolerable withdrawal symptoms occur (usually 1-3 days after a dose change), go back to the previously tolerated dose until symptoms resolve and plan for a more gradual taper with the patient. Dose reduction may need to slow down as one gets to smaller doses (i.e. 25% of the original dose). Overall, the rate of	return of symptoms, pain	None

Triggers for deprescribing







Problem-based deprescribing

- 1. Consider medications as a possible cause
- 2. Prioritize according to risk (e.g., falls, delirium)
- 3. Consider the risk and benefits of deprescribing
- 4. Develop a plan for deprescribing and monitoring (collaborate)

Molnar FJ, et al. Can Geriatr Soc J CME 2018;8(2):1-14







CASE STUDY

Case

- 85-year-old female presents to primary care clinic for dizziness
- Accompanied by daughter who is concerned about her mom being able to keep track of recent medication changes

Past medical history: atrial fibrillation, hypertension, type 2 diabetes with polyneuropathy, hypothyroidism, osteoarthritis, mild cognitive impairment, falls

Social history: no alcohol, smoking or recreational drug use

Allergies: none known

Weight: 61 kg Height: 165 cm





Case

Medications on file	Labs and vitals:
 Bisoprolol 5 mg daily Apixaban 5 mg BID Amlodipine 10 mg daily Rosuvastatin 5 mg daily Levothyroxine 0.075 mg daily Metformin 500 mg twice daily Insulin glargine 8 units daily Duloxetine DR 30 mg daily Gabapentin 600 mg BID Donepezil 10 mg daily Pantoprazole 20 mg daily Vitamin D 1000 units daily Acetaminophen PRN 	 BP 110/78 HR 62 (lying), 91/80 HR 70 (standing) Na= 138 (135-145) K= 3.6 (3.5-5.) SCr= 78 (42-102) eGFR- 64 (>60) A1C= 6.7 (9.2 6 months ago) TSH= 2.83 (0.4-5.00) B12 > 1400 (>220) MOCA 21/30 CHADS2- 5







Consider medications as a cause

- Ensure list of medications is accurate and complete
- List all prescription and non-prescription medications
- Ensure list is what is actually being taken versus what is prescribed
- You review your list and the blister pack and ask about over-thecounter medications, supplements and other natural health products and discover additional medications:
 - Furosemide 20 mg daily- started 4 weeks ago and increased to 40 mg last week. Patient had lower extremity swelling- was seen at a walk-in clinic
 - 2. Vitamin B12 1000 mcg daily- family started due to cognitive impairment
 - 3. Over-the-counter sleep aid- diphenhydramine 50 mg at bedtime when needed





Poll question (#5): Which of the following medications could be contributing to orthostatic hypotension?

- a) bisoprolol, amlodipine and apixaban
- b) bisoprolol, furosemide and duloxetine
- c) donepezil, furosemide and vitamin D
- d) amlodipine, levothyroxine and diphenhydramine





Drug-induced orthostatic hypotension

Drug	How
Diuretics (e.g., furosemide, thiazides)	Volume depletion
Alpha-1-blockers (e.g., terazosin, tamsulosin, prazosin)	Vasodilation
Beta-blockers (e.g., metoprolol, bisoprolol) and calcium channel blockers (e.g., verapamil, diltiazem)	Reduced cardiac output and vascular resistance
RAS inhibitors (e.g., ramipril, valsartan)	Vasodilation
Antidepressants (e.g., TCA, trazodone, paroxetine)	Vasodilation
SGLT2 inhibitors (e.g., empagliflozin, canagliflozin)	Volume depletion
Antipsychotics (e.g., olanzapine, risperidone, quetiapine)	Vasodilation

Possible medication causes

- 1. Bisoprolol 5 mg daily
- 2. Apixaban 5 mg BID
- 3. Amlodipine 10 mg daily
- 4. Furosemide 40 mg daily
- Rosuvastatin 5 mg daily
- 6. Levothyroxine 0.075 mg daily
- 7. Metformin 500 mg twice daily
- 8. Insulin glargine 8 units daily
- 9. Duloxetine DR 30 mg daily
- 10. Gabapentin 600 mg BID
- 11. Donepezil 10 mg daily
- 12. Pantoprazole 20 mg daily
- 13. Vitamin D 1000 units daily
- 14. Vitamin B12 1000 mcg daily
- 15. Diphenhydramine 50 mg PRN





Prioritize and consider risks/benefits of deprescribing

- Temporal relationship- furosemide most likely
- Several other possible causes
- Risks
 - Falls and falls complications
 - Return of swelling- possible prescribing cascade:

Amlodipine --> edema --> furosemide --> dizziness

Gabapentin --> edema --> furosemide --> dizziness

 You decide to focus on stopping furosemide, reassessing and consider additional deprescribing at the next visit





Develop a plan and monitor

- Discontinue furosemide
- Discourage diphenhydramine use for sleep
- Encourage fluid intake
- Monitor dizziness, blood pressure and leg swelling
- Return to office in 1 week for reassessment
 - Reassess other causes of dizziness at next appointment (donepezil, duloxetine, gabapentin, bisoprolol)
 - Consider alternatives for amlodipine





Case follow-up

- Prior to her follow-up appointment, patient presents to ER for a mechanical fall
- RFR: falls and confusion no acute fractures
- Patient admitted to hospital under GIM and flagged for a geriatrics consult
- Pharmacist completes medication reconciliation and review, and discovers the following:
 - Patient was taking medications from old blister pack, containing furosemide, which was stopped about 5 days ago by GP

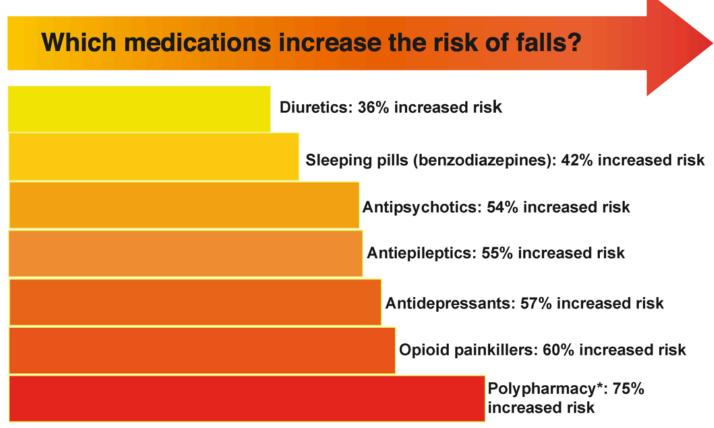




Case: on admission

Medications	Labs
 Bisoprolol 5 mg daily Apixaban 5 mg BID Amlodipine 10 mg daily Furosemide 40 mg daily Rosuvastatin 5 mg daily Levothyroxine 0.075 mg daily Metformin 500 mg twice daily Insulin glargine 8 units daily Duloxetine DR 30 mg daily Gabapentin 600 mg twice daily Donepezil 10 mg daily Pantoprazole 20 mg daily Vitamin D 1000 units daily Vitamin B12 1000 mcg daily Acetaminophen PRN Diphenhydramine 50 mg PRN – last used 2 nights ago 	 BP = 105/73, HR=62 Na=134 K=3.8 SCr= 105 (was 78 last week) BG=3.6 Recent HbA1C=6.7% TSH=2.83
	Tarento Reha

Polypharmacy and falls



*In this analysis, the most commonly used definition for polypharmacy was 4 or more medications. Sources: de Vries M et al. 2018, Seppala LJ et al. 2018, Seppala LJ et al. 2018





FRIDs: Fall-Risk-Increasing-Drugs

- Antihypertensives
- Anticonvulsants
- Antipsychotics
- Antidepressants
- Benzodiazepines/Sedative/hypnotics
- Opioids
- Antispasmodics, muscle relaxants
- Antidiabetic agents
- Anticholinergic agents





Problem-based deprescribing

- 1. Consider medications as a possible cause
- 2. Prioritize according to risk (e.g., falls, delirium)
- 3. Consider the risk and benefits of deprescribing
- 4. Develop a plan for deprescribing and monitoring (collaborate)

Molnar FJ, et al. Can Geriatr Soc J CME 2018;8(2):1-14



Case: Medication causes of falls

- 1. Bisoprolol 5 mg daily
- 2. Apixaban 5 mg BID
- 3. Amlodipine 10 mg daily
- 4. Furosemide 40 mg daily
- Rosuvastatin 5 mg daily
- 6. Levothyroxine 0.075 mg daily
- 7. Metformin 500 mg twice daily
- 8. Insulin glargine 8 units daily
- 9. Duloxetine DR 30 mg daily
- 10. Donepezil 10 mg daily
- 11. Pantoprazole 20 mg daily
- 12. Vitamin D 1000 units daily
- 13. Vitamin B12 1000 mcg daily
- 14. Acetaminophen PRN
- 15. Diphenhydramine 50 mg PRN sleep





Poll Question (#6): Which medication would you prioritize for deprescribing for our patient?

- a) donepezil
- b) pantoprazole
- c) insulin glargine
- d) furosemide
- e) amlodipine





Framework for deprescribing medications

CEASE protocol: 5 steps

- Confirm current medications
- Estimate risk
- Assess each medication
- Sort: Prioritize medications to deprescribe
- Eliminate: discontinue and monitor





Prioritize according to risk

- 1) Risk for recurrent falls:
 - Low blood pressure and orthostasis: furosemide, amlodipine, bisoprolol, duloxetine, gabapentin, donepezil
 - Hypoglycemia: insulin glargine
 - Anticholinergic: diphenhydramine
- 2) Medications with no indications:
 - Pantoprazole
 - Vitamin B12





Discharge plan/medication changes

Changes to medications	For follow-up and monitoring
STOPPED: 1) furosemide 2) amlodipine 3) diphenhydramine 4) insulin glargine 5) vitamin B12 6) pantoprazole NEW: 1) melatonin 5 mg SL QHS	 BP, orthostatic vitals> if BP control required, consider ACEi as patient has DM2 Sleep BG and recheck HbA1C in 3 months Recurrence of GERD symptoms Reassess with GP need for duloxetine and gabapentin Consider tapering donepezil with monitoring of cognitive function





Considerations for seamless care:

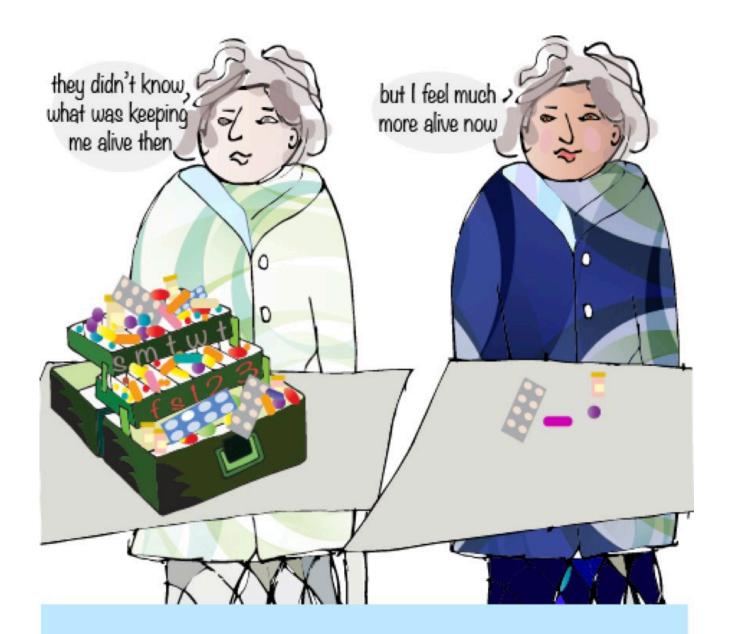
- Ensure medication changes are communicated to patient (or family/caregivers), community pharmacy and primary care provider
- When discontinuing medications, include a prescription order to "stop all refills on _____" to ensure patient does not continue to have stopped medications filled
- Include rationale for changes where possible

Post-discharge resources:

- Homecare referral for OT home assessment
- Geriatric Hub referral
- House Calls
- MedsCheck Follow-up with community pharmacy
- Pharmacist home visit







Thank you!

Questions/Comments???



