### **Approach to Polypharmacy**

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### Impact of Aging on polypharmacy:

 Understanding age-related changes associated with pharmacokinetics and pharmacodynamics helps one optimize safe prescribing and appreciate the impact of polypharmacy. Table 1 gives summary of these changes.

| Parameter        | Age Effect                                                                                                                                                                                                                                                 | Example                                                                                          |
|------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| Absorption       | Aging change: decrease rate of gastric emptying and slowing Impact: slows absorption capacity however bioavailability unchanged                                                                                                                            | Decreased absorption Vitamin B12, iron, calcium                                                  |
| Distribution     | Hydrophilic medications: decrease Vd due to decreased total body water                                                                                                                                                                                     | Increased serum levels of digoxin and alcohol                                                    |
|                  | Hydrophobic meds: increase Vd due to increased fat                                                                                                                                                                                                         | Increased half-life of<br>Benzodiazepines                                                        |
|                  | Decreased muscle mass                                                                                                                                                                                                                                      | Digoxin as it binds to skeletal muscle                                                           |
|                  | Decrease plasma protein especially albumin: higher proportion of unbound drug- more active                                                                                                                                                                 | Diazepam                                                                                         |
| Metabolism       | Aging Change: Decrease in liver mass and liver blood flow affects drug metabolism Impact: Decrease first pass metabolism which increases bioavailability of certain drugs Change in renal function also impacts drug metabolism in liver.                  | Increased bioavailability of<br>Propranolol, labetalol due to<br>decreased first pass metabolism |
| Elimination      | Changes in the kidney: decrease kidney size, decrease blood flow and decrease functioning nephrons lead to decrease GFR Impact: takes longer for drug to eliminate. Hence advisable to dose medications in elderly as if they have chronic kidney disease. | Decreased clearance of diuretics, digoxin                                                        |
| Pharmacodynamics | With aging there is increased sensitivity to drugs especially in brain                                                                                                                                                                                     | Morphine- can have the same effect at half the dose given to younger patients Benzodiazepines    |

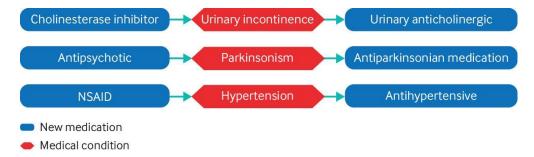
# **Definition of polypharmacy:**

- Most accepted definition based on systematic review by Masnoon et al. is > 5 medications. However, depending on the patient it could be anywhere between > 2 to >11
- Impact of polypharmacy: adverse outcomes such as falls, frailty, increased length of stay in hospital, readmission, mortality and ultimately death
- Adverse outcomes are more pronounced with psychotropic medications
  - Definition of psychotropic medications: these are medications that effect mental function, behavior and experience. Ex: antipsychotics, anticholinergics, antidepressants, anxiolytics.

### **Prescribing Cascade:**

Definition: an adverse event is addressed by giving another medication as depicted in Figure 1

**Impact:** increased pill burden and adverse events which further increases need for hospitalizations and readmissions.



**Figure 1: Prescribing Cascade**. Pigott K L, Mehta N. et al. Using a clinical process map to identify prescribing cascades in your patient. BMJ 2020; 368 doi: https://doi.org/10.1136/bmj.m261 (Published 19 February 2020)

### How to prevent polypharmacy and prescribing cascade

- Thorough medication review at least every 6-12 months
- Optimize prescribing by eliminating potentially inappropriate medications using Beers' criteria table (hospital pharmacy should have this available upon request)
- If an adverse event is identified such as urinary continence, taper off of discontinue as appropriate
- Avoid new psychotropic medications if possible
- De-prescribe psychotropic medications if appropriate
  - Guide to de-prescribing
    - Identify medications and the indication
    - Assess drug-related harm
    - Identify benefit vs harm risk for patient
    - Prioritize which medications to discontinue
    - Initiate de-prescribing and monitor for improvement or any adverse effects as a result
  - Various efforts are happening in the area of deprescribing.
    - De-prescribing algorithms developed by Canadian Deprescribing networks.
    - De-prescribing Research Network in US which promotes research and has a compilation of resources to assist with de-prescribing.
  - o <u>De-prescribing algorithms</u> are available for the following classes:
    - Proton Pump inhibitors
    - Antihyperglycemics
    - Antipsychotics
    - Benzodiazepine receptor agonist
    - Cholinesterase inhibitors and memantine

## Choosing Wisely Campaign with respect to medications in elderly

- 1. Antipsychotic medications are not the first choice to use in dementia with behaviors such as aggression, agitation and disruptive behaviors
- 2. Avoid tight glycemic control in the elderly; they are more prone to hypoglycemia
- 3. Don't prescribe AchEIs in dementia patients unless they are being assessed periodically and being monitored for GI adverse events
- 4. Don't use BZD as first line for insomnia, anxiety, or delirium in the elderly
- 5. Don't prescribe a new medication without a thorough review of med list

#### References

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