# The Concept of Deprescribing

Myra Belgeri, Pharm.D., BCGP, BCPS, FASCP
Clinical Pharmacist
Optum Hospice Pharmacy Services

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# Disclosure

- I have no relevant financial relationships with manufacturers of any commercial products and/or providers of commercial services discussed in this presentation.
- This discussion will include the use of medications for off-label indications.

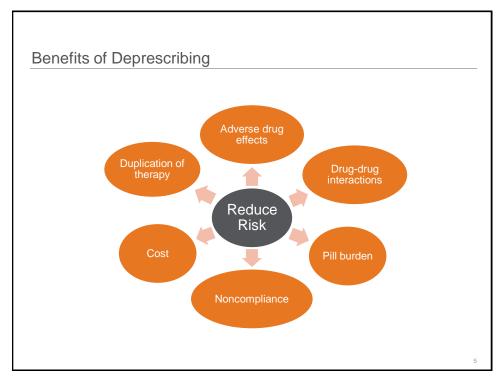
# Objectives

- Describe common barriers to deprescribing medications
- Evaluate the appropriateness and strategies for deprescribing medications
- Describe communication techniques for deprescribing medications

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# Too much stuff?





# Barriers to Deprescribing

# Patient/Family/Caregivers

- · Changing goals of care
- · Attachment to medications
- · Risk of abandonment
- · Influential family members
- · Confrontation with mortality
- Belief that medication discontinuing is suboptimal care

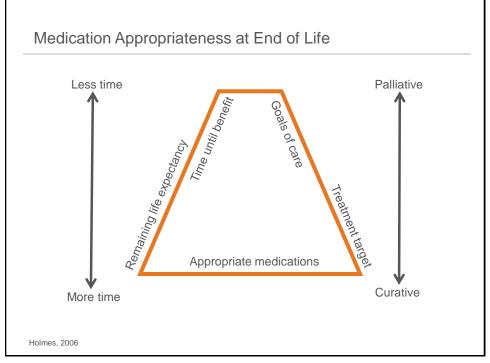
### **Prescribers**

- · Clinical complexity
- Multiple prescribers
- Risk of withdrawal effects
- · Risk of return of symptoms
- Limited information on harm of continuation or discontinuation
- · Limited guidelines on deprescribing

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### Factors to Consider

- Functional status
  - Swallowing ability
  - Ambulation status
  - Cognitive ability
  - Environment
- Time to benefit
- · Medication safety profile
- · Changes in medication effectiveness
- · Changes in goals of care

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# Mrs. Davis

- 78 years old with chief complaint of dyspnea
- She is a fall risk, peripheral edema, dysphagia
- Goal: comfort and reduce medications
- · Primary diagnosis: CHF
- PMH:
  - Atrial fibrillation
  - Type 2 diabetes
  - Hypothyroidism
  - Hyperlipidemia
  - Renal insufficiency
  - Peptic ulcer disease
- PPS: 30
- BP 100/58; HR 70s

- · Medication List:
  - Amlodipine (Norvasc®) 10 mg PO daily
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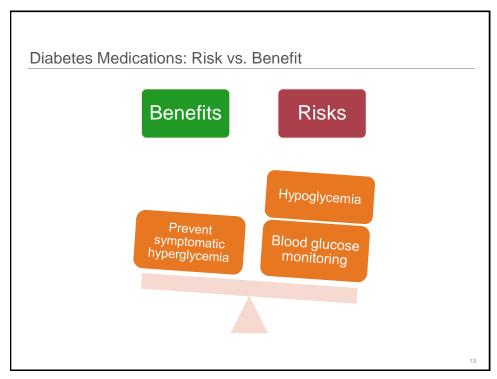
  - Hydralazine 25 g PO TID
  - Clopidogrel (Plavix®) 75 mg daily
  - Metformin (Glucophage<sup>®</sup>) 500 mg PO BID
  - Glipizide (Glucotrol®) 5 mg PO qam
  - Insulin glargine (Lantus®) 20 units QHS
  - Levothyroxine (Synthroid®) 50 mcg PO daily
  - Atorvastatin (Lipitor®) 40 mg PO QHS
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  - Calcium/vitamin D PO BID
  - Omeprazole 40 mg BID

# Medications to Consider Deprescribing

- · Diabetes Medications
- Antihypertensives
- Dementia Medications
- Proton-pump inhibitors

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**Diabetes Medications** 



# Diabetes: Hyperglycemia vs. Hypoglycemia Hyperglycemia Symptoms · Blurred vision · Clumsy or jerky movements · Difficulty concentrating Confusion · Dry mouth · Difficult concentrating • Fatigue Dizziness Headaches Falls Polydipsia Headache Polyphagia Hunger Lethargy • Polyuria • Weight loss · Mental or behavior changes · Pale skin Palpitations Seizures Shakiness Sweating

# Diabetes Medications: Literature

### ADA Guideline Summary

- "For patients with advanced diabetes complications, life-limiting comorbid illness, or substantial cognitive or functional impairment, it is reasonable to set less intensive glycemic target goals"
- "Providers should be vigilant in preventing severe hypoglycemia in patients with advanced disease and should not aggressively attempt to achieve nearnormal A1C levels in patients in whom such targets cannot be safely and reasonably achieved"

# Canadian guidelines:

- · Deprescribe antihyperglycemic agents that cause hypoglycemia
- Deprescribe antihyperglycemic agents in patients experiencing or at risk of adverse effects
- Individual glycemic targets to goal of care for terminally ill patients

American Diabetes Association, 2020; Canadian Diabetes Association, 2013

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# Diabetes Medications: Literature

Patient characteristics/ health status	Rationale	Reasonable A1c goal	Fasting or pre-prandial glucose	Bedtime glucose
Healthy	Longer life expectancy	< 7.5%	90-130 mg/dL	90-150 mg/dL
Complex/ intermediate	Intermediate life expectancy	< 8%	90-150 mg/dL	100-180 mg/dL
Very complex/ poor health	Limited life expectancy	< 8.5%	100-180 mg/dL	110-200 mg/dL
Patients at end of life		Avoid hypoglycemia		

American Diabetes Association, 2020

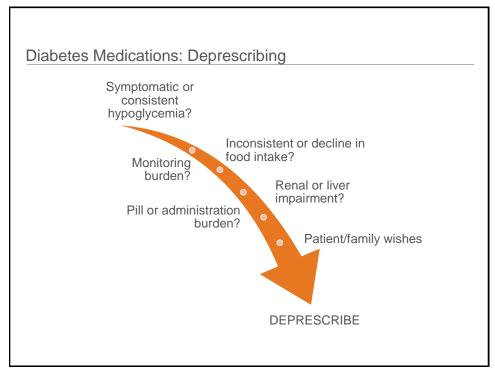
Patient Categories	Life Expectancy	Enteral Intake	Treatment Approach
Active Disease but Relatively Stable	Months to > 1 year	Fair with sporadic improvements or worsening	Goal: prevent hypoglycemia and symptomatic hyperglycemi.      Management:     Adjust medication doses for nausea/vomiting, hepatic/renainpairment, decreased oral intake, weight loss, hypoglycemia
Impending Death or Organ or System Failure	Days to weeks	Declining calorie intake with anorexia	Goal: Prevent hypoglycemia  Management:  • Adjust medication doses for nausea/vomiting, hepatic/renimpairment, decreased oral intake, weight loss, hypoglycemia  • T2DM: Discontinue finger-stick glucose checks  • T1DM: May continue for insulin needs.
Actively Dying	Hours to days	None	Goal: Patient comfort  Medications: Consensus is lacking for patients in this stage. Most prescribers recommend to DC medications T1DM: May be reasonable to continue insulin with liberal BG targets

### Diabetes Medications: Which cause hypoglycemia? **Examples** Cause Hypoglycemia? Drug Alpha-glucosidase Acarbose, miglitol No inhibitors Dipeptidyl peptidase 4 (ex. sitagliptin, saxagliptin, No (DPP-4) Inhibitors linagliptin) Glucagon-like peptide-1 (ex. dulaglutide, liraglutide) No (GLP-1) agonists Yes; Highest risk with regular and Insulins NPH insulin (ex. repaglinide, nateglinide) Yes (low risk) Meglitinides Biguanides (ex. metformin) No Sodium-glucose linked (ex. canagliflozin, dapagliflozin, transporter 2 (SGLT<sub>2</sub>) No empagliflozin) inhibitors (ex. glipizide, glyburide, Sulfonylureas Yes; Highest risk with glyburide glimepiride) Thiazolidinediones (TZDs) (ex: pioglitazone, rosiglitazone)

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Adapted from Farrell, 2017

\*Not a complete list



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# Mrs. Davis - Poll Question 1

- Patient continues on three antihyperglycemic agents:
  - Metformin (Glucophage®) 500 mg PO BID
  - Glipizide (Glucotrol®) 5 mg PO gam
  - Insulin glargine (Lantus®) 20 units QHS

### Which statement is correct?

- A. Discontinue metformin due to decline in renal function
- B. Discontinue glipizide due to risk of lactic acidosis
- C. Insulin glargine should be changed to insulin NPH to reduce risk of hypoglycemia.
- D. Glipizide should be changed to glyburide to reduce risk of hypoglycemia.

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# Mrs. Davis – What should we do?

- Patient continues on three antihyperglycemic agents:
  - Metformin (Glucophage®) 500 mg PO BID
  - Glipizide (Glucotrol®) 5 mg PO qam
  - Insulin glargine (Lantus®) 20 units QHS
- · Agents:
  - Metformin: renally cleared and has risk of lactic acidosis
  - Glipizide: risk of hypoglycemia
  - Insulin glargine: risk of hypoglycemia and burden of injections
- · What are our options?
  - 1) Discontinue one agent
  - 2) Reduce doses







# Antihypertensives: Literature

2017 ACC/AHA Guidelines

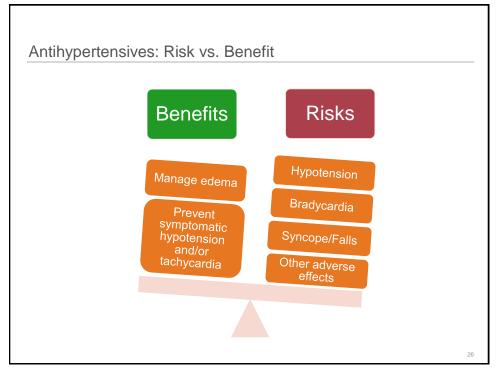
Recommendations for Treatment of Hypertension in Older Persons (≥65 years of age)

- For noninstitutionalized ambulatory community-dwelling adults
  - Treatment of hypertension with a SBP treatment goal of <130 mmHg is recommended
- For older adults with hypertension & high burden of comorbidity & limited life expectancy
  - Clinical judgement, patient preference, and team-based approach to assess risk/benefit is appropriate for decisions regarding BP lowering and choice of antihypertensive medications

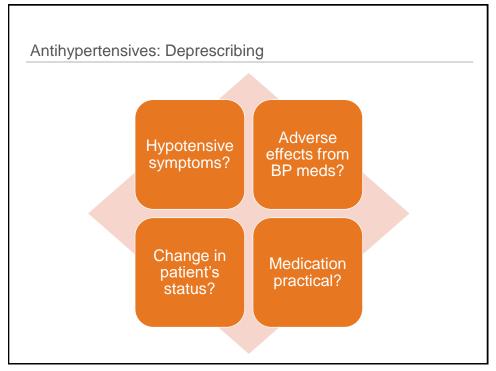
J Am Coll Cardiol. 2018

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Select Antihyper	7 (470)	
Antihypertensive Class	Examples	Adverse Effects Hypotension, dizziness, fatigue
ACE-Inhibitors	lisinopril, ramipril, enalapril	Hyperkalemia, dry cough, angioedema
Angiotensin Receptor Blockers	irbesartan, losartan, valsartan	Hyperkalemia, angioedema, chest pain, diarrhea
Alpha-1 Blockers	doxazosin, terazosin	Orthostatic hypotension, edema
Alpha-2 Agonist	clonidine	Orthostatic hypotension, anticholinergic effects, edema
Beta-Blockers	atenolol, carvedilol, metoprolol	Bradycardia, bronchospasms (non-selective), cold extremities, mask symptoms of hypoglycemia
Calcium Channel Blockers  – Dihydropyridine	amlodipine, felodipine, nifedipine	Edema, flushing, headache
Calcium Channel Blockers  - Non-Dihydropyridine	diltiazem, verapamil	Edema, bradycardia, constipation, flushing
Diuretics	furosemide, hydrochlorothiazide	Electrolyte imbalance, dehydration, nocturnal diuresis
Direct Vasodilators	hydralazine	Headache, palpitations, angina, sodium & water retention



# Antihypertensives: Deprescribing

- · Risk of discontinuation
  - · Rebound hypertension
- · How to discontinue?
  - Abrupt discontinuation
  - · Taper (ex. beta blockers, clonidine)
- · What to discontinue first?
  - Medication that is causing adverse drug effects
  - · Medication with least benefit or without dual benefit
  - · Medication that is inconvenient for patient/family
  - · Medication that was last started

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# Deprescribing Antihypertensives – Which To Taper?

### Beta-blockers

- · Abrupt discontinuation may result in rebound hypertension, tachycardia, or angina
- Gradually taper over 1-2 weeks

### Alpha-2 agonist (clonidine)

- Abrupt discontinuation may result in rebound hypertension, nervousness, agitation, headache, or tremor
  - Less likely to occur with clonidine transdermal patch (compared to oral therapy)
- Gradually taper over 6-10 days by reducing dose by 30-50% every 2-3 days
- If patient is taking a beta-blocker <u>and</u> clonidine, taper and discontinue the beta-blocker several days before tapering clonidine

Calcium channel blockers - non-dihydropyridine (diltiazem, verapamil)

- · Abrupt discontinue may result in angina
- · Consider a gradual dose reduction

Lexi-Comp

Diabetes	Chronic Kidney Disease	Atrial fibrillation	Heart Failure
<ul> <li>ACE-Inhibitors</li> <li>ARBs</li> </ul>	<ul> <li>ACE-Inhibitors</li> <li>ARBs</li> </ul>	Beta Blockers     Non-DHP CCB (ex. diltiazem)	Benefit:  ACE-Inhibitors  ARBs  Beta blockers  Loop diuretics  Possible Benefit  Vasodilators  Aldosterone antagonist  Thiazide diuretics

# Mrs. Davis - Poll Question 2

- Medication List:
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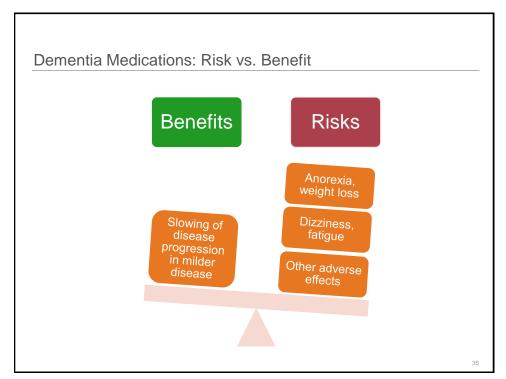
Which medication would you consider reducing or discontinuing FIRST?

- A. Amlodipine (Norvasc®) 10 mg PO daily
- B. Carvedilol (Coreg®) 6.25 mg PO BID
- C. Furosemide (Lasix®) 40 mg PO daily
- D. Hydralazine 25 mg PO TID



# **Dementia Medications**

Acetylcholinesterase inhibitors NMDA-receptor antagonist



Medication	Adverse Effects	Notes
Acetylcholines	sterase Inhibitors (AChEI)	
Donepezil (Aricept®)	Anorexia, diarrhea, nausea, bradycardia, heart block, urinary incontinence, insomnia, headache, hallucinations, agitation, confusion	Mild to severe Alzheimer's Dementia
Rivastigmine (Exelon®)	Anorexia, diarrhea, nausea, bradycardia, heart block, urinary incontinence, insomnia, headache, hallucinations, agitation, confusion, dizziness, fatigue	Mild to severe Alzheimer's dementia; mild to moderate Parkinson's dementia     Capsule must be swallowed whole     Apply patch to the upper or lower back (alternatively may be applied to upper arm or chest)     Safety risk with patches – patient picking; medication error with other patches
Galantamine (Razadyne®)	Anorexia, diarrhea, nausea, bradycardia, heart block, urinary incontinence, insomnia, headache, hallucinations, agitation, confusion, dizziness, fatigue, depression	Mild to moderate Alzheimer's dementia     Caution in renal, hepatic impairment
NMDA Recept	or Antagonist	
Memantine (Namenda <sup>®</sup> )	Dizziness, headache, constipation, hallucinations, confusion	Moderate to severe Alzheimer's dementia     Caution in renal, hepatic impairment     Entire content of capsules may be sprinkled on applesauce and swallowed immediately.     Do not chew, crush or divide     Do not mix oral solution with any other liquid

### **Dementia Medications**

- In nursing home patients with advanced dementia, 53.9% received at least one medication with questionable benefit
  - · Cholinesterase inhibitors were most commonly used
- · 2009 Survey of hospice medical directors
  - · 20% of patients were taking a cholinesterase inhibitor and 20% were taking memantine
  - Majority did not believe either medication class was effective in patients with end-stage dementia
  - 80% recommended discontinuing these therapies
  - · Some benefits noted in the study (for continuing therapy):
    - Improve or stabilize cognition and function
    - Decrease challenging behaviors
    - Improve quality of life and energy
    - Reduce time spent caregiving or need for nursing home placement
    - Reduce caregiver burden or improve caregiver quality of life

Shega, 2009

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# Dementia Medications: Deprescribing

### **Considerations to Continue**

### Patient has demonstrated beneficial effects of the drug:

- Improved cognition
- Managed behaviors
- Improved verbal communication
- Improved functional ability
- Family or patient resistant to discontinuation

### **Considerations to Discontinue**

- Decline (rapid or over the past 6 months) in cognition and functional status
- Severe/end-stage dementia (dependent on all ADLs, inability to respond to stimuli)
- Adverse effects
  - Diarrhea, nausea/vomiting, dizziness, headache, insomnia, weight loss, falls
- Drug interactions
- > Reduce pill burden
- Reduce cost of care
- No longer part of goals of care
- > Family or patient decision to discontinue

Liao, 2018; Reeve, 2018; Renn, 2017

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# Dementia Medications – Taper or No Taper?

- Reduce dose by 50% (or the next available dosage form) every 4 weeks
- Consider a faster taper if time is limited: reduce the dose by 25-50% every 1-2 weeks
- If there is no time to taper, abrupt discontinuation is appropriate
- After taper and/or discontinuation
  - Monitor cognition, function, and neuropsychiatric symptoms
  - If patient has changes, consider other causes of changes (i.e., infection, delirium, dehydration)

Liao, 2018; Reeve, 2018

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# Dementia Medications – Monitoring After Discontinuation

Less than 1 week Severe symptoms including agitation, aggression, hallucinations, or reduced consciousness

- Cause: Likely withdrawal of medication(s)
- Plan: Restart medication at previous dose

2 – 6 weeks

Worsening of cognition, behavioral or psychological symptoms, or function

- Cause: Re-emergence of symptoms that were controlled by medication(s)
- Plan: Consider restarting medication at previous dose and evaluate

6 weeks to 3 months Worsening of cognition, behavioral or psychological symptoms, or function

- Cause: Re-emergence of symptoms that were controlled by medication(s)
- · Plan: Evaluate

> 3 months

Re-emergence of any symptoms

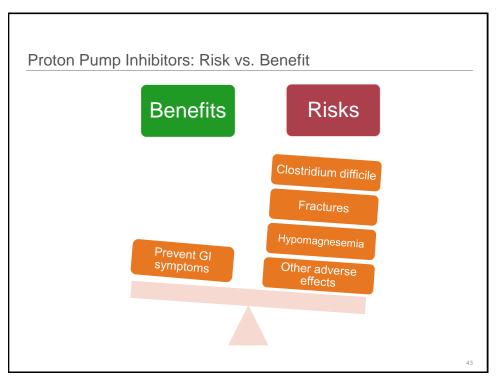
- · Cause: Progression of condition
- Plan: Evaluate

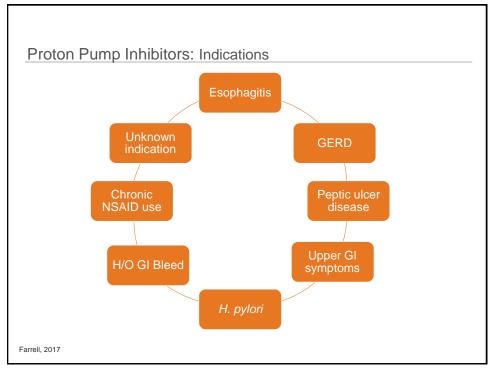
Liao, 2018; Reeve, 2018

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Proton Pump Inhibitors (PPI)





# Proton Pump Inhibitors: Literature

American College of Gastroenterologists guideline

- · Management of GERD and peptic ulcer disease
  - Suggest short-term treatment (2-12 weeks) for most patients
  - GERD 4-8 weeks
  - Peptic ulcer disease 2-12 weeks
- · Recommendations are to discontinue PPI after recommended duration of therapy
- · Maintenance/continued therapy is warranted for a compelling indication
  - Erosive esophagitis or Barrett esophagus
  - If continued, use at lowest dose or changing to on-demand or intermittent PPI use

Farrell, 2017

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# Proton Pump Inhibitors: Deprescribing

- Discontinuation options
  - Taper dose of PPI
    - · Not many indications warrant BID dosing
  - Stop abruptly
  - Change PPI to "as needed" dose
  - Change PPI to an H2 antagonist (ex. famotidine) as an alternative
- Monitor for heartburn, dyspepsia, regurgitation, epigastric pain, decreased appetite, weight loss

Farrell, 2017

# Mrs. Davis - Poll Question 3

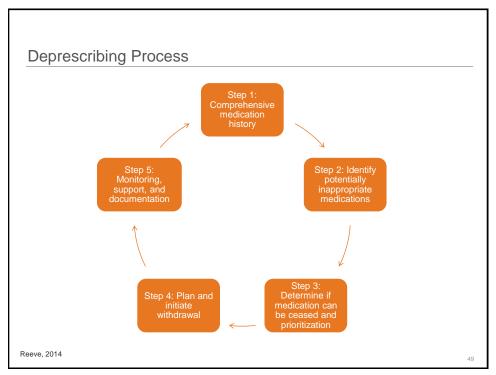
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  - Hyperlipidemia
  - Renal insufficiency
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This patient is on omeprazole for peptic ulcer disease. What is the typical recommended duration of therapy?

- A) 2-12 weeks
- B) 8-16 weeks
- C) Indefinitely
- D) There are no recommendations on duration of therapy.

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# Too much stuff?





# End of Life Communication

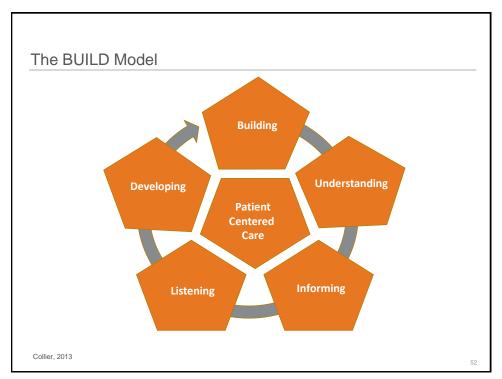
### Studies have found...

- Avoiding discussions can lead to poor patient satisfaction and psychological morbidity
- Key areas of patient satisfaction include:
  - Talking honestly and straightforward
  - Talking about dying
  - Providing information in a sensitive manner
  - Listening and encouraging questions
  - Being considerate with timing of discussions have been identified as key areas of patient satisfaction

Linsky A, 2015; Reeve, 2014

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# The BUILD Model

**Build** a foundation of trust and respect.

 $\underline{\underline{\mathbf{U}}}$ nderstand what the patient and caregiver know about the medication and disease progression.

**Inform** the patient and caregiver of evidence-based information.

**<u>Listen</u>** to the patient's and caregiver's goals and expectations.

**Develop** a plan of care in collaboration with the patient and caregiver.

Collier, 2013

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# The BUILD Model: Key Phrases

**Building** a foundation of trust and respect:

- "I appreciate you sitting down to talk with me today."
- "Your dad is really lucky to have you looking out for him."

**Understanding** what the patient and caregiver know about the medication and disease:

- "What do you know about what this medication is supposed to do?"
- "How do you think your husband's symptoms will look once this medication is no longer working?"

**Informing** the patient and caregiver about appropriateness of medications:

- "We know that this drug works well in milder disease, but it usually stops helping after awhile."
- "We'll need to keep making adjustments to your medications as things change."

Collier, 2013

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# The BUILD Model: Key Phrases

**Listening** to the patient and caregiver as they share goals and expectations:

- "What other information can I provide to help you decide what is best?"
- "Did you and your wife ever talk about what she would want if she couldn't make her own medical decisions?"

**Developing** a plan of care in collaboration with the patient and caregiver:

- "I can't tell you what to do or make the choice for you, but I can provide you with tools so you can make an informed decision."
- "We have some options: we can stop the medication now, we can reduce the dose and follow-up next week, or we can leave everything the same for now."

Collier, 2013

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# Planned Discussions

- · At time of admission
- · Prior to recertification
- · During a family or facility care conference
- · When it is time to re-order a potentially disease-delaying medication
- · When filling the patient's pillbox or ordering refills
- · Change in location or level of care
- · Change in patient condition

# **Key Points**

- · Continuously reevaluate medication appropriateness at end of life.
- Deprescribing guidelines are lacking and though some clinical guidelines mention medication use at end of life, recommendations are not specific. Clinical judgement is required.
- Medication use of end of life should be prioritized based on the patient's preferences, functional status, goals of care, and prognosis.
- Practicing effective communication leads to increased patient satisfaction, better understanding, and improved clinical outcomes.



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# Thank you.

Myra Belgeri, Pharm.D., BCGP, BCPS, FASCP

Clinical Pharmacist

myra.belgeri@optum.com



Hospice Pharmacy Services

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