# Updates in Overactive Bladder

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### Disclosure

- I have no relevant relationships with ineligible companies to disclose within the past 24 months
- The case studies in this lecture are fictional

## Objectives

- At the conclusion of this session participants should be able to:
  - Definition of OAB/ Stress Urinary Incontinence / Urge urinary incontinence
  - Discuss signs and symptoms of OAB and how to recognize it in clinical practice
  - Discuss Treatment options
    - Behavioral
    - Medications
    - Procedures / surgical
  - Review American Geriatrics Society Beers Criteria for OAB medications
  - Case Studies for review

### Question 1

- All of the following are TRUE about OAB (Overactive Bladder) EXCEPT:
- 1. Self limiting
- 2. Urodynamic Studies are required to make the diagnosis
- 3. Behavioral modifications are first line therapy
- 4. Symptoms include urgency and frequency of urination

### Question 2

- Which statement BEST defines Beer's Criteria
- 1. A list of medications used to treat OAB
- 2. Inclusion Criteria to assist in making the diagnosis of OAB
- 3. A list of beers that have a lower likelihood of causing OAB symptoms
- 4. A list of potentially inappropriate medications for the elderly

### Question 3

• All of the following are 3<sup>rd</sup> line treatment options for Overactive Bladder EXCEPT:

- 1. Botox
- 2. Percutaneous Tibial Nerve Stimulation
- 3. Urinary diversion
- 4. Sacral Neuromodulation

### Overactive Bladder

- "urinary urgency, usually accompanied by frequency and nocturia, with or without urgency urinary incontinence, in the absence of UTI or other obvious pathology"
- Self limiting
- Clinical diagnosis

IUGA (International Urogynecological Association) and ICA (International Continence Society)

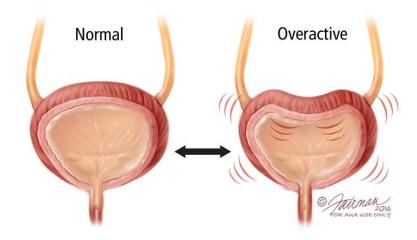
- Prevalence up to 27% men / 47% women (AUA guidelines 2019)
- Pathophysiology: abnormal urothelium and suburothelial signaling that leads to pathologic sensation of urgency

### OAB Terms

- Detrusor Overactivity urodynamic observation characterized by involuntary / spontaneous or provoked detrusor contractions during the filling phase
- Nocturnal polyuria excessive (>20-30%) urine output at night
- Polyuria > 40 ml urine / kg body weight during a 24 hr period
- **Postvoid residual (PVR)** volume of fluid remaining in the bladder after completion of micturition
- **Urgency** sudden desire to void that is difficult to defer
- Urinary Frequency > 8 micturitions in 24 hr period
- **Urgency urinary incontinence** involuntary loss of urine associated with urgency (different than SUI)

## Overactive Bladder (OAB)

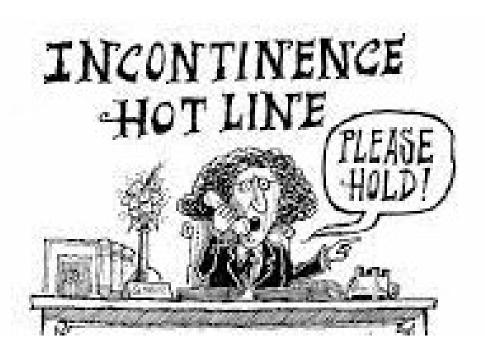
#### **OVERACTIVE BLADDER**



#### Definition

- Syndrome NOT a disease
- Different from stress urinary incontinence (SUI)
- Can be associated with Urge Urinary Incontinence (UUI)
- Urgency
- Urinary Frequency
- Nocturia
- Urgency urinary incontinence

## Urge Urinary Incontinence



- Definition
  - Urge to void immediately and often associated with involuntary urine leakage
  - Temporary or persistent
  - · Quality of life

## Questionnaire for patients

Name:	Date:	MRN #

#### OAB-q short form symptom bother

This questionnaire asks about how much you have been bothered by selected bladder symptoms during the past 4 weeks. Please place a  $\checkmark$  or  $\times$  in the box that best describes the extent to which you were bothered by each symptom during the past 4 weeks. There are no right or wrong answers. Please be sure to answer every question.

	aring the past 4 weeks, how bothered are you by	Not at all	A little bit	Some- what	Quite a bit	A great deal	A very great deal
1.	An uncomfortable urge to urinate?		2	3	4	5	6
2.	A sudden urge to urinate with little or no warning?		2	3	4	5	6
3.	Accidental loss of small amounts of urine?		2	3	4	5	6
4.	Nighttime urination?		2	3	4	5	6
5.	Waking up at night because you had to urinate?		2	3	4	5	6
6.	Urine loss associated with a strong desire to urinate?		2	3	4	5	6

## Symptoms

- Urgency with urination
- Urge urinary incontinence (Wet OAB)
- Difficult to control urination
- Frequency of urination usually > 8 x in 24 hr period
- Nocturia 2+ per night
- Worsens with age
- Prevention: healthy weight / exercise / limit caffeine and etoh / no smoking / pelvic floor exercises

## Physical Exam

- Abdominal / Rectal / GU / pelvic
  - Rule out pelvic prolapse
- Cognitive function
  - Rule out dementia
- Post void Residual to rule out urinary retention

## Treatment Options

- Shared Decision Making
  - Based on QoL
- First Line Behavioral Modifications
- Second Line pharmacotherapy
- Third Line

- Mrs LUTs is a 83 yo female with PMhx for HTN
  - 2 vaginal deliveries Full Term
  - Medications: Lisinopril
- CC: Urge urinary incontinence worsening x 5 years with rare SUI (stress urinary incontinence) only when she laughs a lot (which she laughs often)
- OAB questionnaire
- Work up
  - UA micro negative
  - PVR 2 ml
  - PE: pelvic organ prolapse
  - Obtain Voiding Diary



## Voiding Diary



- Look at frequency volume chart
- What is NORMAL
  - Voiding every 3-4 hours / median 6x a day
- Polydipsia related frequency -> managed with behavior / limit fluids
  - Calculate the urine output overnight
  - Excessive fluid intake can present as OAB

## **Voiding Diary Specifics**

- Voiding Diary Prefer 2-3 days of information
  - On each day: wake up time // Bedtime
  - Measure each void. Write it down in ml or oz, recorded on the chart to the nearest hour
  - Document degree of urgency for each void (0-3): while, 0=1 no urgency, 1=10 urgent)
  - Record of all "wet" events and degree of wetness. For example: dry, damp/dribble, wet/stream, soaked/flood or by pad weighing.
    - What you were doing when you leak examples standing up / getting your leggings off etc
  - Record of pad/underwear alterations
  - Record fluid intake in ml / cc and what it is
    - Coffee / water / wine etc.

- Reviewed Voiding Diary
- Discussed behavioral modifications / Bladder training
- Healthy bladder diet education provided
- Discussed bladder training
- She has been doing pelvic floor exercises with her hot yoga instructor but states "its not helping"
- What to do?



- Treatment:
  - Healthy bladder diet / pelvic floor exercises
  - Rx Virbegron 75 mg daily
- Follow up in ~6 weeks to re-evaluate



- Follow up appointment
  - PVR 0 ml
  - Some improvement in urgency / frequency / leakage
  - Continue to encourage behavioral modifications
    - Referral to pelvic floor PT in addition to yoga
    - Bladder training discussed



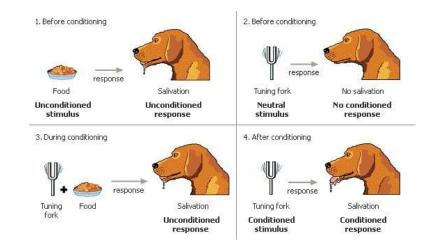
## OAB Workup Review

- Voiding Diary
- Urinalysis / Urine cx
- No role for urodynamic study / cystoscopy / imaging in the initial workup
- Discuss treatment options

- Shared decision making when discussion treatment options
  - Every patient is different!

## First Line Therapy

- Behavioral Modifications
  - Diet healthy bladder diet
  - Bladder Training (change bladder habits / delayed voiding)
  - Pelvic Floor Exercises / referral to PT
  - Weight loss
    - 8% weight loss in obese women caused 42% decrease in s/sx compared to 26% in control
  - Vaginal atrophy topical estrogen cream
  - Timeframe 6-12 weeks



## Healthy Bladder Diet

	Avoid the	ese bladder i	irritants	
All alcoholic beverages	Carbonated Drinks	Cranberries	NutraSweet	Saccharin
Apples	Champagne	Fava beans	Onions (raw)	Sour cream
Apple juice	Cheese	Grapes	Peaches	Soy sauce
Bananas	Chicken livers	Guava	Pickled herring	Strawberries
Beer	Chilies/Spicy foods	Lemon juice	Pineapple	Tea
Brewer's Yeast	Chocolate	Lentils	Plums	Tomatoes
Canned Figs	Citrus fruits	Lima Beans	Prunes	Vinegar
Cantaloupes	Coffee	Nuts	Raisins	Vitamins- buffered with aspartame
	Corned beef	Mayonnaise	Rye bread	Yogurt

## Bladder Training



#### STEP 1

For one or two days, take note of how many times you urinate or leak.

#### STEP 3

Based on your calculations, choose an interval long enough to hold between urinating.

#### STEP 2

Each day, calculate the number of hours you wait between each urination.

#### STEP 4

Empty your bladder first thing in the morning. Add 15 minutes to each interval each day. Continue with this training.

#### STEP 5

Once comfortable with this schedule, increase the intervals by another 15 minutes.

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## Pelvic Floor Exercises – not just kegels





## Second Line Therapy

- Can combine with 1<sup>st</sup> line therapy
- Medications
  - Anti-muscarinic agents
    - ER (extended-release medications preferred)
    - Higher risk of dementia
  - Beta 3 adrenergic agonists
    - \$\$\$
  - COMBO
  - Takes up to 12 weeks to notice full effects
  - Risk of urinary retention, monitor with PVR
  - Use in caution with PVRs > 250-300 ml

## Pharmacological Treatments

#### Anti-muscarinic agents

- MOA stimulates
   acetylcholine to reduce
   smooth muscle contraction
   in the bladder
- Increase bladder capacity / decrease urgency
- Generic options / cheaper
- Can cause cognitive dysfunction
- Side Effects: dry mouth and eyes, constipation
- Contraindications
- Examples: trospium / darifenacin

### • Beta 3 adrenergic agonists

- Mirabegron \* (risk of HTN)
- Vibegron
- MOA smooth muscle relaxation in the bladder
- Less side effects compared to anti-muscarinic agents
- \$\$\$
- Contraindications
  - Uncontrolled HTN
  - Child Pugh class B / ESRD GFR < 30</li>
  - Flecainide / propafenone cannot take 50 mg dose

## Anti-Muscarinic agents

### • Oxybutynin / Tolterodine

- Comes in immediate or extended release
  - Prefer extended release to min. SE
- Cheap / generic
- Not well tolerated

### Oxybutynin

- Highly lipophilic / crosses the blood brain barrier resulting in CNS adverse effects
- Can be given transdermal or ER dosing which decreases SE
- Avoid in elderly

## AUA Update Series 2021 OAB

### • Trospium

- less likely to pass the blood brain barrier / take at least ONE hour prior to food
- Food significantly decreases bioavailability
- M2 / M3
- No need to adjust for hepatic dz / only AM NOT metabolized by CYp3A4
- Study UK patients who use AM (antimuscarinic agents) have 20% increased risk of Dementia in the future

## AUA Update Series 2021 OAB

- Combo Beta 3 adrenergic agonists and AM drugs
  - Improvements in volume voided / frequency / urgency and QOL
- Role of PDE5 inhibitors
  - Tadalafil FDA approved for LUTS in men with BPH

Liu, Bonnie et al. Drug Therapy for Overactive Bladder. AUA Update Series 2021, Vol 40 page 193.

## Pharmacological Treatment

- Recommend Follow up 4-6 weeks after starting medication
- If no improvement titrate medication / combo
  - Solifenacin / Trospium PLUS mirabegron
- If some improvement titrate medication
- Obtain PVR if >1/3 total voided amount watch closely
- Cannot tolerate side effects -> 3<sup>rd</sup> line therapy
- Side Effects: dry mouth / constipation / dry eyes / blurred vision/ dyspepsia/ Urinary Retention / impaired cognitive function

### **Beers Criteria**

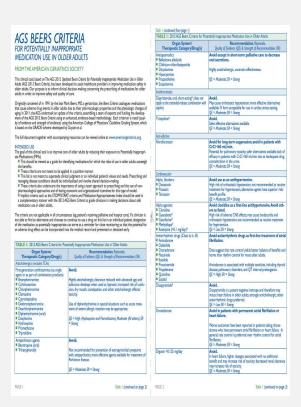
- American Geriatric Society updates Beers Criteria
  - Criteria:
    - Potentially inappropriate medications in older adults
    - Potentially inappropriate medications to avoid in older adults with certain conditions
    - Medications to be used with caution in older adults
    - Medication combinations that may lead to harmful interactions
    - List of medications that should be avoided / dosed differently in those with poor renal function





### OAB and Beer

- Updated in 2019
- Anticholinergics / Anti-muscarinic agents made the LIST
- Prescribe with caution in the elderly
  - anti-muscarinic agents are contraindicated in elderly on oral potassium supplements d/t slowing gastric motility
  - Trospium is considered the safest (lowest DDI – drug drug interactions)



### **AUA Beers List**

https://www.auanet.org/guidelines/guidelines/beers-criteria

Potentially Inappropriate Medication for Use in Older Adults (developed by the AUA Beers Criteria White Paper Workgroup based on  $^{\prime}$ ).

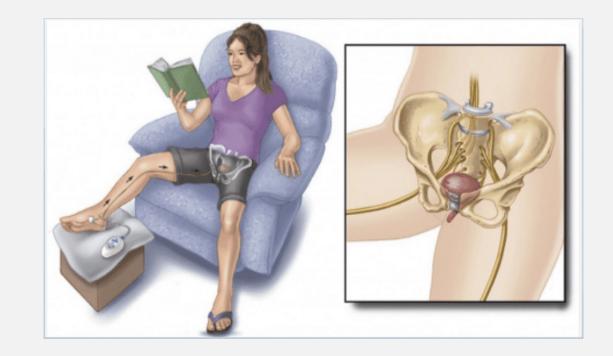
Medication	Urologic Indication			
Nitrofurantoin	Acute treatment of uncomplicated urinary tract infection without systemic symptoms in individuals living in communities with an identified high risk of quinolone-resistant organisms OR due to multiply-resistant bacteria with identified sensitivity to nitrofurantoin			
Alpha-blockers	Medical management of bothersome benign prostate enlargement symptoms while monitoring for efficacy and adverse events			
Estrogenics	Topical (vaginal) use for symptomatic vaginal atrophy due to low estrogenic states Risk reduction for chronic recurrent urinary tract infections in postmenopausal women			
Anti-muscarinics	Trial of antimuscarinics is appropriate as second line therapy in patients with high bother from overactive bladder symptoms, with monitoring of benefits, risks, and adverse effects for that individual patients. Trial of antimuscarinics is appropriate for male patients with benign prostatic enlargement in whom the symptom complex includes high bother from urgency and frequency symptoms in the absence of significant urinary retention (post-void residual urine volume <200 mL) and in patients for whom first-line therapy for OAB fails.			

## Third Line Therapy

- Percutaneous Tibial Nerve Stimulation
- Botox
- Implantable Tibial Nerve Stimulation
- SNS

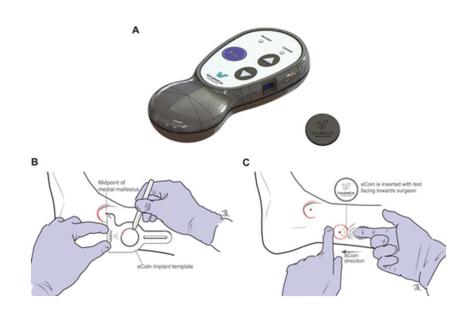
### **PTNS**

- PTNS Peripheral tibial nerve stimulation
- Started in 1983 / FDA approval 2005
  - Less invasive
  - Acupuncture-like electrical nerve stimulation
  - Weekly for 12 weeks / 30 minutes each session
  - Needle placed medially behind the ankle with mild electrical stimulation
  - Shown to reduce OAB s/sx and improve quality of life



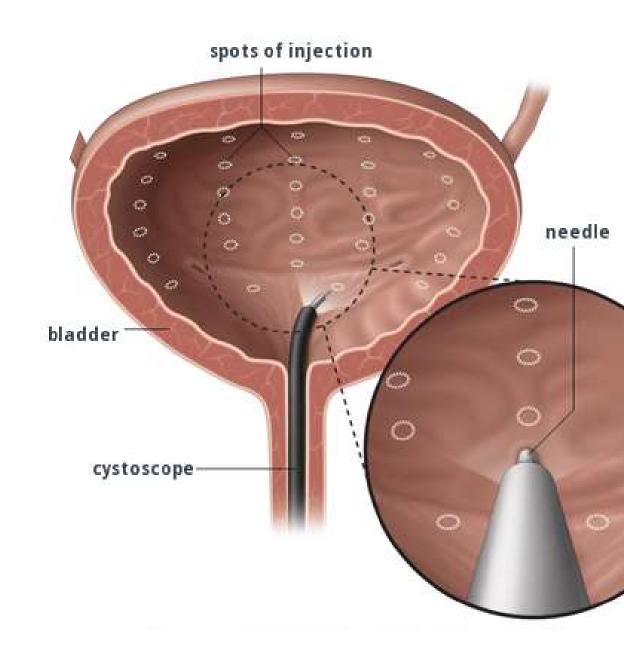
# Implantable / Peripheral Neurostimulator

- eCoin Peripheral Neurostimulator System
  - FDA Approved March 2022
  - Indication Urge Urinary Incontinence
  - Implanted into the ankle
  - Not MRI compatible
  - Provides Tibial nerve stimulation 2x a week
  - 3 year battery life



## BOTOX

- Botulinum toxin
  - Consider if failed pharmacologic therapy
  - Botox administered under local anesthesia
  - Results are seen within 2 weeks and last for 3-12 months
  - Can cause increased risk of UTIs
  - AUA guidelines 2019
    - Pt must be able to do SIC if needed



## Sacral Neuromodulator

- Sacral Neuromodulation (SNS)
  - FDA approval 1997
  - Min. invasive surgical electrical stimulation
  - InterStim / Axonics
  - Patients must be able to learn to adjust the setting with a small device
  - Wire is placed into S3 foreman and connected to stimulation device
    - Two phase procedure
      - Test phase need to see > 50% improvement in S/sx
      - Second stage implantation phase





## Case Study

- 74 yo male with hx of BPH with LUTS (lower urinary tract symptoms)
  - PMHx: HTN BMI 32
  - Workup / GU history
    - Urodynamic study showed bladder outlet obstruction
    - PVR 35 ml
    - Prostate US 55 cc
    - I-PSS score 25/35 Q 4
  - S/p TURP 12/2020 / path benign



### International Prostate Symptom Score

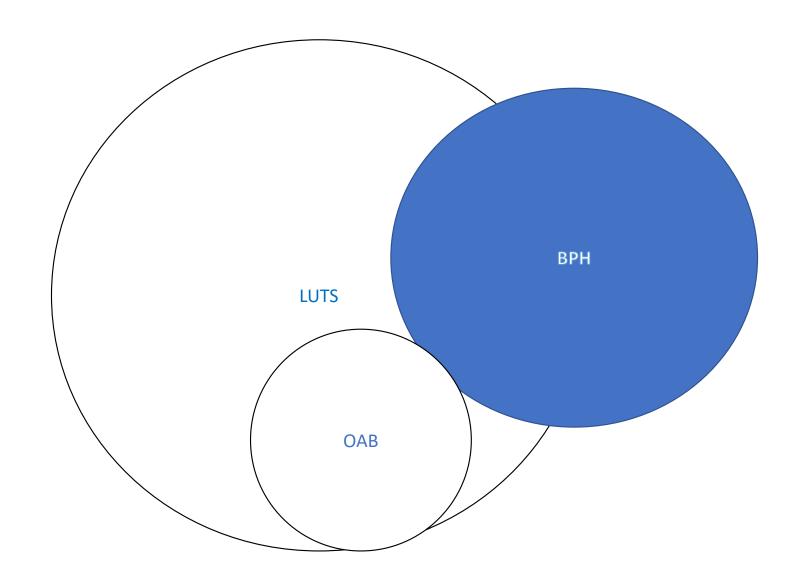
#### International Prostate Symptom Score (I-PSS)

Patient Name: \_\_\_\_\_ Date of birth: \_\_\_\_\_ Date completed \_\_\_\_

In the past month:	Not at All	Less than 1 in 5 Times	Less than Half the Time	About Half the Time	More than Half the Time	Almost Always	Your
<ol> <li>Incomplete Emptying         How often have you had the sensation of not emptying your bladder?     </li> </ol>	0	1	2	3	4	5	
2. Frequency How often have you had to urinate less than every two hours?	0	1	2	3	4	5	
3. Intermittency How often have you found you stopped and started again several times when you urinated?	0	1	2	3	4	5	
4. Urgency How often have you found it difficult to postpone urination?	0	1	2	3	4	5	
5. Weak Stream How often have you had a weak urinary stream?	0	1	2	3	4	5	
6. Straining How often have you had to strain to start urination?	0	1	2	3	4	5	
	None	1 Time	2 Times	3 Times	4 Times	5 Times	
7. Nocturia How many times did you typically get up at night to urinate?	0	1	2	3	4	5	
Total I-PSS Score							

Score: 1-7: Mild 8-19: Moderate 20-35: Severe

Quality of Life Due to Urinary Symptoms	Delighted	Pleased	Mostly Satisfied	Mixed	Mostly Dissatisfied	Ushappy	Terrible
If you were to spend the rest of your life with your urinary condition just the way it is now, how would you feel about that?	0	1	2	3	4	5	6



## Case Study

- Initially did very well with TURP but 6+ months later developed increasing frequency / urgency and severe urge incontinence with some lack of sensory awareness
- 3 depends a day / accidents
- Effected his quality of life
- I-PSS 15/35 (urgency / frequency / nocturia)
- Treatment options



## Case Study

- Behavioral Modifications / weight loss
- Started on mirabegron 25 mg x 2 months
- No significant improvement
- Discussed 3<sup>rd</sup> line therapies Botox / PTNS
- Elected to proceed with PTNS

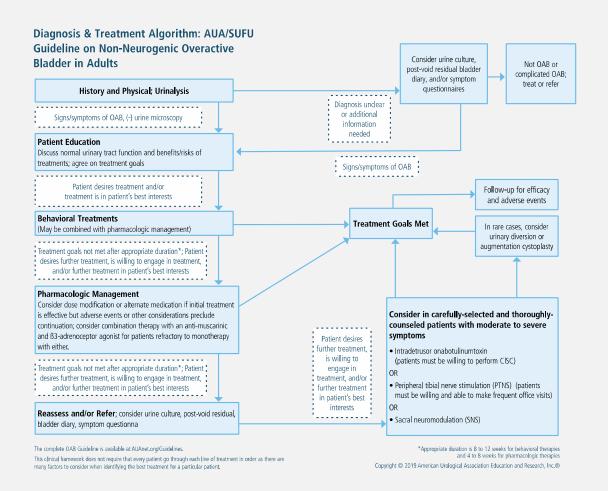




#### **PTNS**

- Prior to starting PTNS
  - Voids 5-6x a day
  - 3-4x nocturia
  - High Urgency
  - Severe urge urinary incontinence / 3 depends daily / daily accidents
- After completing 12 weekly sessions
  - Voids 4-5x a day
  - 1x nocturia
  - 1 depends a day, sometimes stays dry
  - Mild urgency

agreed to have his picture taken for this presentation



## Tips and Tricks

- Nocturia
  - Causes: OAB / Obstructive Sleep apnea / excessive nighttime urine production
- Nocturnal Polyuria
  - Rule out Low nocturnal bladder capacity
- In patients taking OAB meds, they may need bowel regimen to prevent constipation

### Question 1

- All of the following are TRUE about OAB (Overactive Bladder) EXCEPT:
- 1. Self limiting
- 2. Urodynamic Studies are required to make the diagnosis
- 3. Behavioral modifications are first line therapy
- 4. Symptoms include urgency and frequency of urination

### Question 2

- Which statement BEST defines Beer's Criteria
- 1. A list of medications used to treat OAB
- 2. Inclusion Criteria to assist in making the diagnosis of OAB
- 3. A list of beers that have a lower likelihood of causing OAB symptoms
- 4. A list of potentially inappropriate medications for the elderly

## Question 3

• All of the following are 3<sup>rd</sup> line treatment options for Overactive Bladder EXCEPT:

- 1. Botox
- 2. Percutaneous Tibial Nerve Stimulation
- 3. Urinary diversion
- 4. Sacral Neuromodulation

# Clinical Pearls

- When to refer to urology?
  - No response with 1<sup>st</sup> line therapy
  - Neurological disease
  - Hematuria / pelvic mass / underlying disease that could be contributing
- OAB is a clinical diagnosis
- Treatment plans:
  - require shared decision making
  - step by step approach
  - individualized
- Screen for Dementia in OAB patients
- Indwelling catheters are NOT recommended as management for OAB
- Uncontrolled diabetes is a FACTOR with OAB
- Encourage all women to do pelvic floor exercises

## Thank YOU