Urinary Incontinence in Older Adults: Taking a Patient Centered Approach Martha Spencer, MD, MHSEd, FRCPC

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We would like to acknowledge that we are gathered today on the traditional territories of the Musqueam, Squamish and Tsleil-Waututh peoples.





1. Review the **normal physiological changes** of aging that occur in the lower urinary tract.

2. Consider how **medical comorbidities** and **medications** can contribute to urinary incontinence.

3. Discuss evidence-based **non-pharmacological treatments** for overactive bladder and their utility for different patient populations of older adults.

4. Review **pharmacological treatments** for overactive bladder in older adults, considering appropriateness based on the individual patient characteristics.

Spread the Word!

Urinary incontinence is NOT a normal part of aging!!!



Aging and UI

- Very little longitudinal data to support increase in lower urinary tract symptoms (LUTS) with aging + physiological changes with aging
- Cross-sectional studies (EPIC, EPICONT)
 - Increasing prevalence of LUTS
 - Transition from urgency urgency incontinence detrusor failure
- Difficult to separating effects of declining estrogen and aging
 - Pelvic organs and their surrounding structures are estrogen responsive
 - Menopause if a major risk factor for developing pelvic floor disorders (epidemiological studies)
 - LUTS and LUTS severity increase after menopause

Age (years)	Prevalence (%)		
18-29	5.9		
30-39	4.2		
40-49	8.5		
50-59	11.8		
60-69	12.3		
70-79	15.6		
Age-standardized	8.6		

Tab. III. Prevalence (%) of overactive bladder by age of women [15]

- Most significant increase in OAB wet
- Most significant increase in OAB severity between 60-69yo
- Nocturia most bothersome symptoms for woman (>urgency or urgency incontinence)

Urination: Physiological changes with aging

• Bladder

- Decreased capacity
- Decreased sensation to filling
- Increased detrusor overactivity
- Decreased contractile function
- Increased residual
- Decreased urethral closure pressure (women)
- Increased night-time urine production (blunting of ADH peak)
- Altered central and peripheral neurotransmitter concentrations/actions
- Increased white matter hypodensities in the brain

Abrams et al. Incontinence. 5th International Consultation on Incontinence. 5th Edition, 2013.

The Brain and UI



Increased periventricular WM= Increased incidence of LUTS + LUTS severity

Gibson, W., & Wagg, A. (2017). Incontinence in the elderly, 'normal'ageing, or unaddressed pathology?. *Nature Reviews Urology*, 14(7), 440.

Associated conditions and UI

- Peripheral vascular disease
- Diabetes mellitus
- Congestive heart failure
- Venous insufficiency
- Chronic lung disease
- Falls and contractures
- Sleep disordered breathing
- Stroke

- Dementia
- Diffuse Lewy body disease
- Parkinson's disease
- Normal Pressure Hydrocephalus
- Recurrent infection
- Constipation
- Obesity

Culprit medications

- ACEI- cough (SUI)
- Anticholinergics- impaired bladder emptying, constipation, sedation
- Calcium channel blockers- impaired emptying, constipation, leg edema
- Cholinesterase inhibitors- increase bladder contractility
- Diuretics- increase urinary frequency
- Lithium- polyuria
- Opioids- impaired bladder contractility, constipation, sedation

Culprit medications

- Psychotropics (benzodiazepines, antipsychotics, z-drugs, trazadone)sedation, enuresis
 - Many also have anticholinergic effects (ex. quetiapine, trazadone)
- SSRIs- increase bladder contractility, decreased tone of the urethral sphincter (increases cholinergic transmission)
- Gapabentin/pregabalin-leg edema
- NSAIDs- leg edema

Case Example

Case 2: Ms. G

- 88yo female, living in Assisted Living for 5 years after a fall and hip fracture
- Needs help for bathing, dependent IADLs
- Mobilizes with 4-wheeled walker
- Loves to play bridge, spending time with her Yorkie Maisey
- Urinary symptoms:
 - Frequency- 9x/day, 1-2x/night (with enuresis)
 - Urgency with urgency incontinence if waits >5min past urge to void
 - Leaking with coughing and sneezing
 - No hesitancy, no sensation of incomplete emptying
 - 3-4 pads/day, large pad on bed at night
 - Daily BM but hard (Bristol Stool Chart 1-2)
 - Bother/QOL- stopped going on outings with friends



Ms. G.

• Past medical history:

- CAD (NSTEMI 2011, 2016)
- HTN
- Dyslipidemia
- TIA (2009)
- CHF (EF 40%)
- GERD
- Depression
- Anxiety



Ms. G. ■Medications

Ramipril 10 mg po daily ■Furosemide 80 mg po daily ■Amlodipine 5mg po daily ■Bisoprolol 5 mg po daily Atorvastatin 20 mg po daily ■ASA 81 mg po daily ■Alendronate 70mg po weekly Amitriptyline- 20mg po qhs Clonazepam- 0.5mg po qhs ■Calcium 1250mg po bid ■Ferrous fumarate 300mg po daily

Physical exam



- Independent for transfers, unable to rise without use of arms
- ■Timed up-and-go 18 seconds
- ■BP 120/70 lying, 106/66 standing
- ■Bilateral pitting leg edema
- Sacral innervation intact
- Anal wink intact, anal tone present but weak, hard stool on DRE
- Perineum excoriated, no sacral wounds, no prolapse
- Weak pelvic floor muscles
- ■Positive stress test in the supine position
- ■Post-void residual urine volume 50 mL

Investigations

GFR 50

HbA1C= 6.2%

Urinalysis negative

What Type of Incontinence?

- Urgency incontinence
- Stress incontinence
- Functional incontinence

Multifactorial Incontinence

Treatment

Clinical Frailty Scale*

I Very Fit – People who are robust, active, energetic and motivated. These people commonly exercise regularly. They are among the fittest for their age.

 Well – People who have no active disease symptoms but are less fit than category 1. Often, they exercise or are very active occasionally, e.g. seasonally.

3 Managing Well – People whose medical problems are well controlled, but are not regularly active beyond routine walking.

4 Vulnerable – While not dependent on others for daily help, often symptoms limit activities. A common complaint is being "slowed up", and/or being tired during the day.

5 Mildly Frail – These people often have more evident slowing, and need help in high order IADLs (finances, transportation, heavy housework, medications). Typically, mild frailty progressively impairs shopping and walking outside alone, meal preparation and housework.

6 Moderately Frail – People need help with all outside activities and with keeping house. Inside, they often have problems with stairs and need help with bathing and might need minimal assistance (ouing, standby) with dressing.



7 Severely Frail – Completely dependent for personal care, from whatever cause (physical or cognitive). Even so, they seem stable and not at high risk of dying (within ~ 6 months).

8 Very Severely Frail – Completely dependent, approaching the end of life. Typically, they could not recover even from a minor illness.



Ferminally III - Approaching the end of life. This category applies to people with a life expectancy
 6 months, who are not otherwise evidently frail.

Scoring fraility in people with dementia

The degree of frailty corresponds to the degree of dementia. Common **symptoms in mild dementia** include forgetting the details of a recent event, though still remembering the event itself, repeating the same question/story and social withdrawal.

In moderate dementia, recent memory is very impaired, even though they seemingly can remember their past life events well. They can do personal care with prompting.

In severe dementia, they cannot do personal care without help.

* I. Canadian Study on Health & Aging Revised 2008. 2.K. Rockwood et al.A. global clinical measure of fitness and frailty in elderly people. CMAJ 2005;173:489-495.

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Lifestyle interventions

 No recommendations are possible regarding lifestyle interventions for UI in the frail elderly (Level 4)

Best evidence- Increased hydration for incontinent frail elderly may decrease UI

Abrams et al. Incontinence. 5th International Consultation on Incontinence. 5th Edition, 2013.

Behavioral modifications

- Prompted voiding
 - May be effective in nursing home and home care
 - Ineffective if need more than 1PA for transfer
 - Do not continue if <20% reduction in wet checks or toilet successfully <2/3 of times after 3 day trial
- Timed voiding- insufficient evidence
- Habit retraining- insufficient evidence
- No proven interventions for night-time UI or UI in hospitalized patients

Behavioural interventions

- Pelvic floor muscle training (PFMT)
 - Effective as a **stand-alone treatment**, as part of a **multicomponent strategy**
 - PFMT may be just as effective as drug therapy and that combination of drug therapy + PFMT more effective than either treatment alone
 - Supervised PFMT should be offered as a first-line conservative therapy for women of all ages with UI (Level 1 evidence, Grade A recommendation)

Abrams et al. (2017). Incontinence, 6th International Consultation on Incontinence.

Behavioural interventions Scheduled voiding regimens:

- Bladder training (BT)- should be recommended as a first-line conservation therapy for UI in women
 - Start with 1hr intervals and increase by 15-30min intervals until q2-3hr voiding intervals achieved
 - Consider self-monitoring with diary/log
 - Importance of supervising HCP
 - Consider different strategy if no improvement after 3 weeks
 - BT likely as effective as drug therapy

Abrams et al. (2017). Incontinence, 6th International Consultation on Incontinence.

Behavioral interventions

- Behavioural interventions + physical activity (Talley et al, 2017)
 - 42 cognitively intact elderly females (mean age 84.9)
 - Frail- >3 on Vulnerable Elders Survey, slow gait speed/assistive device
 - Single-blind, two-arm pilot RCT
 - Intervention- 12 week program:
 - Customized behavioral treatments
 - 150 minutes/week of walking
 - 2x/week strength training



Behavioral interventions

- Population- mean age 84.9, 98% white, 83% living independently, 62% mixed UI and 22% urgency
- Results
 - 50% decrease in daily leaks (bladder diary)
 - "Overall do you feel you ae better?- 81% treatment group vs. 36% control
 - % improvement- 65% treatment group vs. 34% control
 - Greater improvement in self-reported and objectively measured toilet skills + in balance and gait speed (not statistically significant)
 - No different in improvement in QOL

Treatment

Skin protection

- Barrier cream
- Avoiding long periods of wetness

Treat constipation

- Daily PEG/lactulose (increase frequency prn to achieve 1 soft BM/day)
- Stop Calcium and Iron
- Increase fluid intake
- Physical exercise + behavioural interventions, including individual work with a pelvic floor physiotherapist

Treatment

Medication review

- Lasix at 4pm, consider adjusting dose if able
- Stop Amlodipine (BP too low for level of frailty)
- Taper off Amitriptyline- monitor mood
- Taper off Clonazepam (VERY slowly)

• ?Pharmacological therapy for incontinence?

- Anticholinergic
- Beta-3 agonist

Pharmacological Rx for OAB

Antimuscarinics:

- -Oxybutynin (Ditropan)
- -Tolterodine (Detrol)
- -Darifenacin (Enablex)
- -Solifenacin (Vesicare)
- -Fesoterodine (Toviaz)
- -Trospium (Santura)

Beta-3 agonist: -Mirabegron (Myrbetriq)

Anticholinergic Agents for OAB: Potential Crossing of Blood-Brain Barrier



Chancellor MB et al. Drugs Aging. 2012 April; 29(4):259-273





Geriatr Gerontol Int 2015; 15: 521-534

REVIEW ARTICLE

Effect of pharmacological treatment for urinary incontinence in the elderly and frail elderly: A systematic review

Eva Samuelsson,¹ Jenny Odeberg,² Karin Stenzelius,³ Ulla Molander,⁴ Margareta Hammarström,⁵ Karin Franzen,⁶ Gunnel Andersson⁶ and Patrik Midlöv⁷

- 13 trials of high/moderate quality
 - 11- Antimuscarinic
 - 2- Duloxetine

Systematic review- pharmacological Rx elderly/frail elderly

Results

- Oxybutynin (only drug studied in frail elderly at time of publication)no effect on UI or QOL (4 trials)
- Other anticholinergics (Darifenacin, Fesoterodine, Solifenacin, Tolterodine, Trospium)- decrease in UI (mean= 1/2 leak/24hrs) (7 trials)
 - Adverse effects- dry mouth, constipation
- Data insufficient for quality of life, cognitive effects
- Data insufficient for Duloxetine (SUI)
- No studies on Mirebegron or estrogen (at time of publication)

FORTA Classifications

Class A (absolutely)	Indispensable drug, clear-cut benefit in terms of efficacy/safety ratio proven in elderly patients for a given indication
Class B (beneficial)	Drugs with proven or obvious efficacy in the elderly, but limited extent of effect or safety concerns
Class C (careful)	Drugs with questionable efficacy/safety profiles in the elderly, to be avoided or omitted in the presence of too many drugs, lack of benefits or emerging side effects; review/find alternatives
Class D (don't)	Avoid in the elderly, omit first, review/find alternatives

Wehling M. J Am Geriatr Soc. 2009; 57(3): 560 -

LUTS FORTA Classification: OAB drugs

Class A (absolutely) Indispensable drug, clear-cut benefit in terms of efficacy/ safety ratio proven in elderly patients for a given indication	
Class B (beneficial) Drugs with proven or obvious efficacy in the elderly, but limited extent of effect or safety concerns	• Fesoterodine
Class C (careful) Drugs with questionable efficacy/safety profiles in the elderly, to be avoided or omitted in the presence of too many drugs, lack of benefits or emerging side effects; review/find alternatives	 Darifenacin Mirabegron Extended-release oxybutynin Solifenacin Tolterodine Trospium
Class D (don't) Avoid in the elderly, omit first, review/find alternatives	 Immediate release oxybutynin Propiverine

Fesoterodine in the Frail Elderly

562 frail elderly with urgency urinary incontinence, average age 75 (range 65-91), cognitively intact, with a mean of 8-9 health conditions, 1-in-4 taking \geq 11 meds



DuBeau CE, Kraus SR, Griebling TL et. al. Effect of Fesoterodine in Vulnerable Elderly Subjects with Urgency Incontinence: A Double-Blind, Placebo Controlled Trial. J Urol (2014). 191:2; 395-404)

	No. Placebo (%)	No. Fesoterodine (%)
Subjects with AEs	120 (42.7)	158 (56.2)
Discontinued due to AEs	14 (5.0)	26 (9.3)
Dry mouth*	17 (6.0)	66 (23.5)
Constipation	12 (4.3)	31 (11.1)†
Urinary retention	0	9 (3.2)‡
Diarrhea	7 (2.5)	8 (2.8)
Fatigue	3 (1.1)	8 (2.8)
Dyspepsia	1 (0.4)	7 (2.5)
Headache	5 (1.8)	7 (2.5)
Cough	2 (0.7)	7 (2.5)

Table 3. Treatment emergent AEs with an incidence exceedingthe placebo rate and occurring in 2% or more of subjects

* Reported as severe by 2 subjects in the fesoterodine group and 1 in the placebo group.

† Reported as severe by 1 subject in the fesoterodine group.

‡ Catheterization required by 3 (1%) subjects with urinary retention.

DuBeau CE, Kraus SR, Griebling TL et. al. Effect of Fesoterodine in Vulnerable Elderly Subjects with Urgency Incontinence: A Double-Blind, Placebo Controlled Trial. J Urol (2014). 191:2; 395-404)

Efficacy, safety, and tolerability of mirabegron in patients aged \geq 65 yr with overactive bladder wet: a phase IV, double-blind, randomised, placebo-controlled study (PILLAR)

Adrian Wagg^{a,*}, David Staskin^b, Eli Engel^c, Sender Herschorn^d, Rita M. Kristy^e, Carol R. Schermer^e

Wagg, A., Staskin, D., Engel, E., Herschorn, S., Kristy, R. M., & Schermer, C. R. (2020). Efficacy, safety, and tolerability of mirabegron in patients aged≥ 65 yr with overactive bladder wet: a phase IV, double-blind, randomised, placebo-controlled study (PILLAR). *European urology*, 77(2), 211-220.



Fig. 1 – Study flow diagram. EOT = end of treatment (week 12).

Wagg, A., Staskin, D., Engel, E., Herschorn, S., Kristy, R. M., & Schermer, C. R. (2020). Efficacy, safety, and tolerability of mirabegron in patients aged≥ 65 yr with overactive bladder wet: a phase IV, double-blind, randomised, placebo-controlled study (PILLAR). *European urology*, 77(2), 211-220.

Table 1 – Baseline demographic and OAB characteristics.

Demographic characteristics (safety analysis set)	Placebo $(n = 442)$	Mirabegron 25 mg^{a} (<i>n</i> = 226)	Mirabegron 50 mg ^a ($n = 219$)	Mirabegron total $(n = 445)$
	(1 112)	25 mg (n 220)	50 mg (n 215)	
Female sex, n (%)	324 (73)	168 (74)	149 (68)	317 (71)
Age, mean \pm SD	71.9 ± 6.0	71.6 ± 5.8	71.7 ± 5.2	71.7 ± 5.5
Age \geq 75 yr, <i>n</i> (%)	124 (28)	66 (29)	59 (27)	125 (28)
BMI category, n (%)				
<25	91 (21)	60 (27)	48 (22)	108 (24)
≥25-<30	150 (34)	84 (37)	73 (33)	157 (35)
≥30	201 (46)	82 (36)	98 (45)	180 (40)
Race, <i>n</i> (%)				
White	357 (81)	151 (67)	197 (90)	348 (78)
Asian	54 (12)	58 (26)	1 (0.46)	59 (13.3)
Black or African American	25 (5.7)	16 (7.1)	17 (7.8)	33 (7.4)
Other	6 (1.4)	1 (0.4)	4 (1.8)	5 (1.1)
MoCA total score category, n (%) ^b				
Normal (≥ 26)	305 (69)	168 (74)	142 (65)	310 (70)
Mild (18–25)	103 (23)	44 (20)	68 (31)	112 (25)
Moderate (10–17)	3 (0.68)	1 (0.44)	2 (0.91)	3 (0.67)
Severe (<10)	0	0	0	0
Missing	29 (6.6)	13 (5.8)	5 (2.3)	18 (4.1)
Medical history, most frequent conditions, $n(\%)^{c}$				
Hypertension	243 (55.0)	134 (59.3)	125 (57.1)	259 (58.2)
Osteoarthritis	173 (39.1)	60 (26.5)	87 (39.7)	147 (33.0)
Hypertonic bladder ^d	145 (32.8)	86 (38.1)	72 (32.9)	158 (35.5)
Gastro-oesophageal reflux disease	135 (30.5)	54 (23.9)	77 (35.2)	131 (29.4)
OAB characteristics (full analysis set—incontinence)	Placebo	Mirabegron	Mirabegron	Mirabegron
	(n = 431)	25 mg^{a} (<i>n</i> = 220)	50 mg^{a} (<i>n</i> = 217)	total ($n = 437$)
Duration of symptoms (mo), mean \pm SD	120 ± 112	119 + 119	123 + 113	121 + 116
Number of micturitions/24 h. mean $+$ SD ^e	10.5 ± 3.1	10.7 ± 2.3	10.5 ± 2.5	10.6 + 2.4
Number of incontinence episodes/24 h. mean \pm SD ^e	3.4 ± 3.2	3.2 ± 3.1	3.7 ± 3.1	3.5 ± 3.1
Number of urgency episodes/24 h, mean \pm SD ^{e,1}	5.8 ± 3.9	6.7 ± 3.9	5.0 ± 3.0	5.9 ± 3.6
Number of urgency incontinence episodes/24 h. mean $+$ SD ^e	3.4 ± 3.1	3.2 ± 3.1	3.7 ± 3.1	3.4 ± 3.1
Mean volume voided/micturition (ml), mean \pm SD	185 ± 75.2	175 ± 59.2	199 ± 90.1	186 ± 76.3

B. Number of incontinence episodes/24 h*



p < 0.001

All ages



Wagg, A., Staskin, D., Engel, E., Herschorn, S., Kristy, R. M., & Schermer, C. R. (2020). Efficacy, safety, and tolerability of mirabegron in patients aged≥ 65 yr with overactive bladder wet: a phase IV, double-blind, randomised, placebo-controlled study (PILLAR). *European urology*, 77(2), 211-220.

Side effects

- UTI
- Headache
- Diarrhea
- Nausea
- No change in MOCA scores

?Pharmacological treatment



- Trial non-pharmacological symptoms + medication reduction/discontinuation x 6-8 weeks
- Treat constipation
- Brain problem vs. lower urinary tract problem??
- Reassess degree of bother and effects on QOL
- Consider trial of anticholinergic or Mirabegron
- Follow-up in 2-3 months

Conclusions

- Urinary incontinence is **NOT a normal part of aging** but physiological changes in the lower urinary tract do occur
- The approach to UI must be **individualized**, and will depend on the patient's degree of **frailty, medical comorbidities and medications**
- Non-pharmacological treatments should be offered to all patients and tailored to their individual profile, considering frailty and cognitive status
- Pharmacological therapies can be considered based on response to non-pharmacological treatments, bother and medical, functional and cognitive profile

