

From wet to dry in primary care



A case based workshop

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Doreen is 83 years old. She has had "many years" of urinary frequency and 2 years at least of urinary urgency and urgency incontinence.

She is particularly bothered by nocturnal voiding; she gets up 4 times at night, which disturbs her sleep. She drinks 1L fluid daily and her bowels are kept soft with polyethylene glycol.

Her family has been concerned that she has fallen on the way to the lavatory at night and her daughter has noticed that she occasionally "smells of urine". They report her being more vague and forgetful.

The family is concerned that she is no longer able to look after herself.

- Her medical history includes:
 - myocardial infarction
 - chronic heart failure
 - COPD
 - hypertension
 - diabetes mellitus
 - mild renal impairment
 - chronic lower back pain

- Her surgical history includes:
 - Appendicectomy
 - Tonsillectomy
 - Cholecystectomy
 - Abdominal hysterectomy and "bladder repair" 1998
- She is G6, P5
- No caesarian, ventouse, forceps



Bowels

BO: 4/ week

Formed / hard, followed by "soft stuff"

No faecal urgency

No flatus or stool incontinence

Occasional use of prunes to "help out"

Medication

- Atorvastatin 80 mg OD
- Ramipril 10 mg OD
- Metformin 1000 mg BID
- Dapagliflozin 10 mg OD
- Furosemide 40 mg OM
- Bisoprolol 5 mg OD
- Tiotropium bromide 18 mcg OD
- Amitriptyline 25 mg nocte

- Vitamin D 1000 IU OD
- Calcium carbonate 500 mg OD
- Alendronic acid 70 mg weekly
- Salbutamol 200 mcg prn inh
- Aspirin 75 mg OD
- PEG 3350, 17g OD
- Clopidogrel 75 mg OD



Medication class	Prevalence n (%)
Any medication class	236 (60.5%)
Calcium channel blockers	85 (21.8%)
Benzodiazepines	68 (17.4%)
Other centrally active agents*	65 (16.7%)
ACE inhibitors	56 (14.4%)
Oral estrogens	50 (12.8%)
Oral estrogen + progesterone	12 (3.1%)
NSAIDS	38 (9.7%)
GABAergic analgesics	11 (2.8%)
Loop diuretics	7 (1.8%)
Thiazolinedione hypoglycemic agents	3 (0.8%)
Alpha blocking antihypertensives	1 (0.3%)

Distribution of medications potentially contributing to urinary symptoms in 390 older patients seeking healthcare for their UI

^{*}Antidepressants, antipsychotics and narcotics were combined.

Predictor	Crude odds ratio	Adjusted odds ratio*
	(95% CI)	(95% CI)
Age	1.0 (1.0 - 1.1)	1.0 (1.0 - 1.1)
Sex	1.5 (0.7 - 3.0)	1.1 (0.5 - 2.3)
Polypharmacy (≥ 5 drugs)	5.3 (3.4 – 8.2)	4.9 (3.1 – 7.9)
Multimorbidity (≥ 3)	2.0 (1.2 - 3.3)	1.2 (0.6 – 2.03)
Depression	1.1 (0.7 - 1.67)	0.7 (0.4 - 1.2)
Cardiovascular disease	2.0 (1.3 - 3.0)	1.4 (0.8 - 2.3)
Hypertension	2.2 (1.4 - 3.5)	1.5 (0.9 - 2.7)
Respiratory disease	1.7 (1.0 - 3.1)	1.4 (0.7 - 2.9)
Gastrointestinal disorders	1.5 (1.0 - 2.2)	1.1 (0.6 - 1.9)
Dyslipidemia	1.6 (1.1 - 2.4)	0.9 (0.5 - 1.5)
Renal failure	1.6 (0.6 - 4.1)	0.7 (0.2 - 2.1)
Musculoskeletal disorders	1.3 (1.0 - 1.6)	0.9 (0.6 - 1.2)
Diabetes	1.4 (0.9 - 2.3)	0.9 (0.5 - 1.7)

No associations were found between age or sex and the use of medications potentially contributing to urinary symptoms

No association between between the class of medication and the type or severity of incontinence in multivariate models

Individuals consuming 5 or more drugs were almost 5 times more likely to be taking a medication contributing to urinary symptoms, when adjusting for age, sex and comorbidity (OR = 4.9, 95% CI = 3.1-7.9)

 $^{{\}rm *Adjusted\ for\ age,\ sex,\ polypharmacy\ and\ multimorbidity}.$

Medications	Effects on Continence
Alpha adrenergic agonists	Increase smooth muscle tone in urethra and prostatic capsule and may precipitate obstruction, urinary retention, and related symptoms
Alpha adrenergic antagonists	Decrease smooth muscle tone in the urethra and may precipitate stress urinary incontinence in women
Angiotensin converting enzyme inhibitors	Cause cough that can exacerbate UI
Anticholinergics	May cause impaired emptying, urinary retention, and constipation that can contribute to UI. May cause cognitive impairment and reduce effective toileting ability.
Antidepressants	Associative studies identify link in addition to that with specific groups
Calcium channel blockers	May cause impaired emptying, urinary retention, and constipation that can contribute to UI. May cause dependent ooedema which can contribute to nocturnal polyuria
Cholinesterase inhibitors	Increase bladder contractility and may precipitate urgency UI
Diuretics	Cause diuresis and precipitate UI
Lithium	Polyuria due to diabetes insipidus
Opioid analgesics	May cause urinary retention, constipation, confusion, and immobility, all of which can contribute to UI
Psychotropic drugs Sedatives Hypnotics Antipsychotics Histamine ₁ receptor antagonists	May cause confusion and impaired mobility and precipitate UI Anticholinergic effects Confusion
Selective serotonin re-uptake inhibitors	Increase cholinergic transmission and may lead to urinary UI
Others Gabapentin Glitazones Non-steroidal anti-inflammatory agents	Can cause ooedema, which can lead to noctumal polyuria and cause nocturia and night-time UI

Wagg, A et al "Incontinence in Frail Older Persons" in "Incontinence, 6th Edition, 2017, Abrams, P., Cardozo, L, Wagg, A., Wein, A. (Eds.) International Continence Society, Bristol, UK.

Functional history

She is dependent upon a walking aid in and out of doors.

She can't get into her washroom if she uses it.

She has an electrically operated chair to assist her in standing, but she doesn't use it.

Her walking speed is slower than six months ago.



Containment

- She uses panty liners to help her incontinence by day
- She uses larger pull ups by night
- She uses lavatory paper in her pants in addition to the liner by day
- Her pads leak she reports soreness "down below, Doctor."







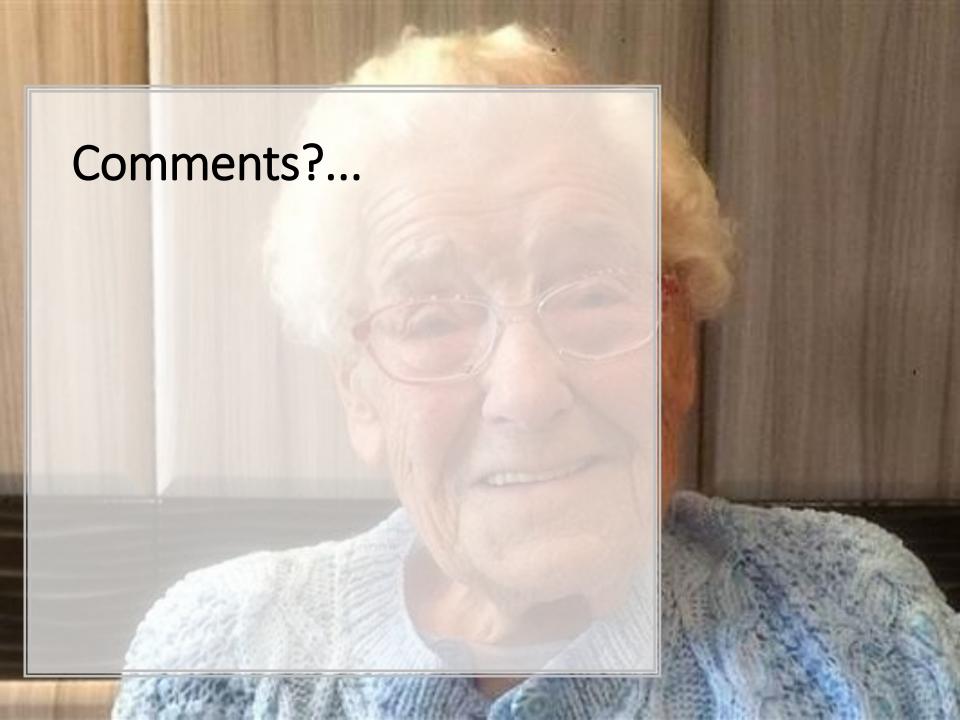
She experiences a small leak on vigorous coughing

A pelvic floor squeeze is pretty nonexistent

Examination

She has:

- trouble transferring from a clinic chair
- a urinalysis positive for glucose ++,
 nitrites +, SG 1.005
- Her urine smells
- Her pad is wet
- Her skin is erythematous with some maceration









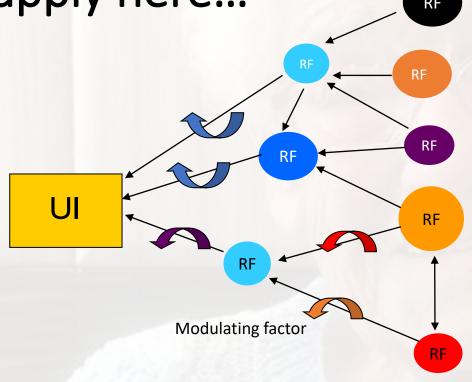








Geriatric syndrome rules apply here...



Tinetti ME, et al. JAMA. 1995;273:1348–53. Inouye SK, et al. J Am Geriatr Soc. 2007;55:780-91. Risk factor







- Bedside commode
- Low level night time lighting
- Home care support
 (example emptying commode in AM)
- Incontinence products
 http://www.continenceproductadvisor.org/



Medication management

Review medications which might compromise successful toileting and potentially add to anticholinergic load

- Anticholinergic load (amitriptyline, tiotropium, furosemide)^{1,2}
- Impaired emptying (amitriptyline, tiotropium)^{2,3}

Polyuria (dapagliflozin,⁴ furosemide⁵)

- Aging Brain Care. Anticholinergic Cognitive Burden Scale 2008, 2012 Update.
- 2. Tiotropium Summary of Product Characteristics. 2016.
- 3. Amitriptyline Summary of Product Characteristics. 2016.
- 4. Dapagliflozin Summary of Product Characteristics. 2017.
- 5. Furosemide Summary of Product Characteristics. 2016.

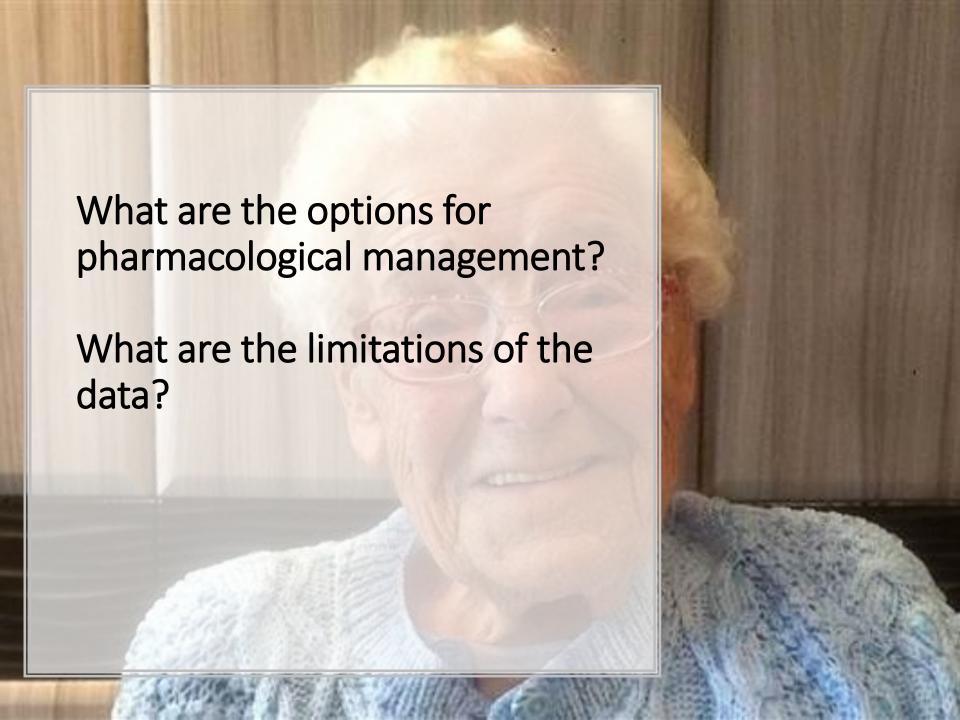




What are the options for management (and the limitations)?

- Limiting fluid consumption¹
- Limiting consumption of caffeine and alcohol¹
- Bladder training urgency suppression techniques¹
- Pelvic floor muscle training¹
- Managing constipation¹
- Smoking cessation (where relevant)²
- Consider vaginal oestrogen¹
- Weight reduction¹

- 1. NICE. Urinary Incontinence in Women Management. Clinical guidelines. Available at https://www.nice.org.uk/guidance/cg171/resources/urinary-incontinence-in-women-management-35109747194821 (accessed February 2017).
- 2. EAU Guidelines on Urinary Incontinence. European Association of Urology. 2016.



Options for pharmacological management?

There are data supporting the efficacy of antimuscarinic drugs in community-dwelling older people (>65y, >75y)¹⁻⁵

From studies planned *a priori* to investigate efficacy in the elderly:

darifenacin¹

fesoterodine^{2,3} (incl. medically complex elderly)

oxybutynin⁴

propiverine (primarily safety)⁵

- 1. Chapple C, et al. Curr Med Res Opin 2007; 23: 2347-58.
- 2. Wagg A, et al. JAGS. 2013;61:185-193.
- 3. Dubeau CE, et al. J Urol. 2014;191:395-404.
- 4. Szonyi G, et al. Age Ageing. 1995; 24:287-91.
- 5. Dorschner W, et al. Aktuel Urol. 2003; 34:102-8.

Efficacy data from pre-planned studies in older nursing home residents:

oxybutynin¹ oxybutynin ER²

From pooled *post-hoc* analyses of community dwelling older people (65+y):

fesoterodine³
oxybutynin ER²
oxybutynin transdermal patch⁴
tolterodine⁵
solifenacin⁶
trospium chloride⁷
1. Ouslander JG
2. Lackner TE, e

- 1. Ouslander JG, et al. J Am Geriatr Soc 1995; 43: 610-17.
- 2. Lackner TE, et al. J Am Med Dir Assoc 2011; 12: 639-47.
- 3. Kraus S, et al. Urology 2010; 76: 1350-57.
- 4. Newman DK. Director 2008; 16: 21-4.
- 5. Malone-Lee J, et al. J Am Geriatr Soc 2001; 49: 700-5.
- 6. Wagg A, et al. Am J Geriatr Pharmacother 2006; 4: 14-24.
- 7. Sand PK, et al. BJU Int 2011; 107: 612-20.

Beta-3- agonist: mirabegron

Pooled analysis of >65 and >75 cohort from registration trials for 12/52-weeks efficacy and 52/52-weeks safety¹

Mirabegron is contraindicated in patients with severe uncontrolled hypertension (defined as systolic blood pressure ≥180 mm Hg and/or diastolic blood pressure ≥110 mm Hg). Blood pressure should be measured at baseline and periodically during treatment with mirabegron, especially in hypertensive patients.²

- 1. Wagg A. et al. Age and ageing. 2014; 43:666-75.
- 2. Betmiga Summary of Product Characteristics. 2016.

Antimuscarinics and cognition

- Data exist for the following drugs in cognitively intact older adults
 - darifenacin¹
 - solifenacin²
 - oxybutynin transdermal gel³
 - tolterodine⁴
 - trospium chloride⁵
 - Fesoterodine⁶

- In older adults with mild cognitive impairment
 - Solifenacin⁷

Trials of fesoterodine^{8,9} and propiverine¹⁰ have monitored MMSE scores and found no change over 12/52 therapy

- 1. Kay G, et al. Eur Urol. 2006; 50:317-26.
- 2. Wesnes KA, et al. Expert Opin Drug Saf. 2009; 8(6):1-12.
- 3. Kay G, et al. Neurourol Urodyn. 2009;28;711-712, ICS2009.
- 4. Kay G, et al. ICS 2006:Abstr 87.
- 5. Staskin D, et al. Int J Clin Pract. 2010; 64:1294-300.
- 6. Kay GG, et al. Postgrad Med. 2012; 124:7-15.
- 7. Wagg A, et al. Eur Urol. 2013; 64:74-81.
- 8. Wagg A, et al. JAGS. 2013;61:185-193.
- 9. . Dubeau CE, et al. J Urol. 2014;191:395-404.
- 10. https://www.ics.org/abstracts/publish/180/000201 poster.pdf

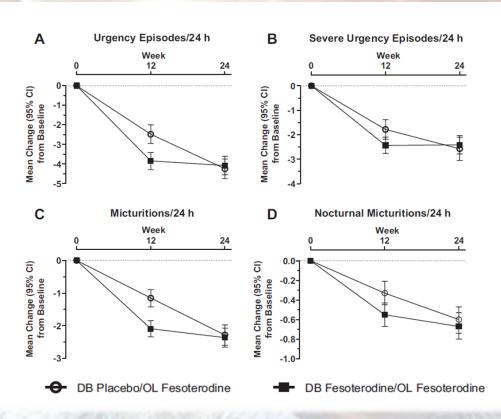
What are the limitations of the evidence?

What else should we take into account?

Can we identify those older persons who might be at more cognitive risk?

Follow up

When should we follow up on Doreen's progress?



DB, double blind; OL, open label.