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# Overactive bladder: Recognition requires vigilance for symptoms

## ■ ABSTRACT

Overactive bladder (OAB) is a prevalent condition in both men and women that imposes significant burdens on the patient and his or her quality of life. Nevertheless, only a small percentage of patients with OAB receive diagnosis and treatment. The identification of OAB is well within the scope of the primary care provider, as it is symptom-based and does not generally require specialized testing. The treatment of OAB relies on behavioral modification and/or pharmacologic options, primarily antimuscarinic therapy. Better identification of OAB symptoms in the primary care setting should reduce the number of patients suffering from untreated OAB.

## ■ DEFINITION OF THE CONDITION

Overactive bladder (OAB) is defined by the International Continence Society as a symptom complex of urinary urgency (intense, sudden desire to void) with or without incontinence, urinary frequency (voiding eight or more times in a 24-hour period), or nocturia (awakening at night to void).<sup>1,2</sup> The symptoms of OAB are present in the absence of any pathologic or metabolic disorders that could cause them.

Although this paper deals primarily with OAB, it is important to recognize other types of lower urinary tract symptoms (LUTS) and dysfunctions that are

common in the primary care setting; for an overview, see the sidebar on the next page.

## ■ PREVALENCE AND SOCIAL IMPLICATIONS

Studies on OAB show that approximately 33 million US adults of all ages are affected.<sup>3</sup> Although OAB historically has been thought of as a disease diagnosed more frequently in women, its prevalence is nearly equal in both sexes, affecting 16.9% of US women and 16.0% of US men.<sup>4</sup> The major difference between the sexes is the frequency of accompanying urinary incontinence (UI) (see sidebar): 55% of women have OAB with UI versus only 16% of men.<sup>4</sup>

The prevalence of OAB increases with age: a multinational population-based survey in Europe showed prevalence rates of 31% and 42% among women and men, respectively, 75 years of age or older compared with rates of 17% and 16% among women and men 40 years of age or older.<sup>5</sup>

### Most cases go untreated

Despite the large number of both men and women with OAB, only 15% of all patients with symptoms of OAB receive treatment.<sup>6</sup> Kinchen and colleagues noted that only one of four women with symptoms of OAB with UI seeks clinical help.<sup>7</sup>

Patients want their primary care provider (PCP) to discuss the issue, yet there appears to be a communication gap.<sup>8</sup> A recent online survey of 1,228 women aged 40 to 65 years (898 of whom had symptoms of OAB) found that more than half of the women who discussed OAB with a health care provider (56%) waited longer than 1 year to seek treatment; many attempted self-management of their symptoms.<sup>9</sup> A contributing factor is the stigma surrounding bladder control problems and the many misconceptions that patients have about their condition that may prevent them from seeking care.<sup>10</sup>

### Social cost of OAB

OAB significantly affects many aspects of a patient's life, including self-esteem, sexual relations, family

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## Lower urinary tract symptoms: Defining terms

Following are some common types of lower urinary tract symptoms (LUTS) and dysfunctions seen in the primary care setting that should be kept in mind when evaluating complaints suggestive of overactive bladder:

**Urge urinary incontinence** is involuntary leakage accompanied or immediately preceded by urgency, and is most commonly associated with overactive bladder.

**Stress urinary incontinence** is involuntary leakage associated with physical activities that increase intra-abdominal pressure, such as coughing, sneezing, or laughing, or upon effort or exertion, such as lifting objects or exercising. Stress urinary incontinence reflects a dysfunction of the bladder outlet, the urethral sphincter.

**Mixed urinary incontinence** involves the coexistence of urge and stress incontinence.

**Overflow incontinence** occurs when the bladder becomes overdistended as a result of outlet obstruction and may manifest as constant or postvoid dribbling. It is seen most often in men with benign prostatic hyperplasia but can be seen in women with pelvic organ prolapse.

relations, lifestyle, professional life, health perception, and sleep.<sup>9</sup> “Bathroom mapping” is a common behavioral technique of patients with OAB. Since they need to void frequently, they will consciously or unconsciously conduct a search for all the bathrooms in the vicinity in order to prevent an emergency. It is not uncommon for OAB patients to avoid prolonged social activities so as to not embarrass themselves with the frequency of toilet use.

### Financial cost of OAB: Direct, indirect, and intangible

The social implications of OAB can be explained by its direct, indirect, and intangible costs. Direct costs include those associated with treatment, diagnosis, routine care, and the consequences of the disease. Indirect costs encompass lost wages and productivity. Intangible costs are associated with suffering, embarrassment, and overall decreased quality of life.<sup>11</sup>

A study by Hu et al estimated that the direct cost of OAB in the United States was \$12.6 billion during 2000.<sup>12</sup> Although providers are familiar with the costs of treatment and routine care of OAB, they may not be as aware of the costs associated with the consequences of the disease, which make up greater than 50% of the overall costs. These consequences include skin irrita-

tion, urinary tract infections, falls, additional admissions to institutions, and prolonged hospital stays.<sup>13</sup> The odds ratio of a hip fracture is two times greater in an elderly woman with urge UI than in the general population.<sup>14</sup> One can only speculate on the reason for this increased risk; however, the authors have treated OAB patients who have tripped in a dark room as they raced to the bathroom and others who have slipped in a urine puddle on their way back.

The indirect costs and intangible costs of OAB are more difficult to quantify. It may be helpful to think of the indirect costs as the wages lost to missed work as a result of OAB or the decreased productivity from continually needing to find the bathroom. Although intangible costs defy being assigned an actual price by their very definition, they nevertheless can be devastating. In a national community survey, Coyne et al reported that both continent and incontinent OAB patients suffered in all health-related quality-of-life measures compared with controls who did not have OAB.<sup>15</sup> Additionally, the prevalence of depression is markedly higher in patients suffering OAB, with or without UI, than in the general population.<sup>8</sup>

Although the social implications of OAB clearly reflect a large societal burden, there is good news in the form of opportunities for intervention in treatment-naïve patients. A 2006 study by Balkrishnan et al shows that compliance with medications for OAB can result in a significant decrease in older adults' health care costs.<sup>16</sup>

## ■ PATHOGENESIS

OAB is a syndrome with a varied pathophysiology that may be multifactorial. The detrusor is composed of smooth muscle under voluntary neurologic control. Idiopathic OAB has been proposed to be secondary to myogenic or subclinical neurogenic abnormalities. A “neurogenic bladder” is a result of neurologic dysfunction. Although the etiology of OAB is not clear, the cause of its symptoms is better understood and will be reviewed below.

### How the bladder normally functions

To understand the abnormal function suffered by the patient with OAB, it is instructive to first review normal bladder function. Micturition involves two important and discrete processes: (1) bladder filling and storage, and (2) bladder emptying.<sup>17</sup> The filling and storage phase requires accommodation of increasing volumes of urine at low intravesical pressures with appropriate sensation, a closed bladder outlet (adequate outlet resistance), and absence of involuntary contractions (which result in urgency or leakage). The process of

bladder emptying requires a coordinated contraction of the bladder muscle, a lowering of the resistance of the outlet (sphincter), and an absence of anatomic obstruction. All types of voiding dysfunctions may be classified by an abnormality of one or more of the factors listed, alone or in combination.

Although its specific etiology is not known, OAB can be explained as the inability to accommodate increasing volumes of urine as a result of high intravesical pressures, along with increased sensation causing symptoms of urgency and frequency with or without a contraction.

### Abnormal urge sensation in OAB

In the OAB patient, signals to the bladder allow contraction and subsequent micturition before the bladder reaches full capacity. When this signal is sudden, intense, and difficult to deter, patients report the urgency associated with OAB. This is opposed to a normal urge sensation that allows the patient adequate warning to prepare for bladder emptying. Whether this abnormal signaling is an amplification (ie, increased sensitivity) of the afferent “sensory” fibers or increased output of the efferent motor fibers is not known. Antimuscarinic therapy aims to block these pathways from overresponding,<sup>18</sup> as discussed below.

### OAB, benign prostatic hyperplasia, and bladder outlet obstruction are interrelated

In discussing the pathophysiology of OAB, the connection between OAB, benign prostatic hyperplasia (BPH), and bladder outlet obstruction is important to note. The incidence of OAB increases with age, and many men develop OAB symptoms concomitantly with BPH. The most common cause of voiding symptoms in men is related to urethral obstruction from the prostate gland, secondary to BPH. Approximately 50% of men with some type of prostatic obstruction also have detrusor overactivity, but conversely, men younger than 60 years who present with LUTS tend not to have an enlarged prostate or a history of BPH.<sup>19</sup> The relationship between symptoms and OAB, bladder outlet obstruction, and BPH remains unclear.

### PRESENTING SYMPTOMS

Patients may avoid seeking medical care for OAB for several reasons, including embarrassment, a belief that the symptoms are part of normal aging, or the perception that it is not a valid medical condition.<sup>20,21</sup> Physicians tend not to raise the issue for lack of time, concern that the evaluation is difficult, or concern that the treatment options are minimal.

Several studies show that women are more likely to

use inconvenient and lifestyle-altering coping strategies than to seek treatment from a health care provider.<sup>20</sup> The most common strategies are wearing absorbent incontinence pads or feminine hygiene pads and always locating the nearest bathroom when away from home. In addition, women may wear special clothes or use deodorant powders or sprays to help conceal wetting accidents, and some may carry a change of clothes wherever they go.

Patients generally present because their symptoms have simply become intolerable. Women tend to present when urge UI develops and their quality of life is affected. Men will start to complain when nocturia becomes excessive. The likely reason for the difference is that UI tends to be more common in OAB among women than among men (approximately 50% vs 15% of OAB cases, respectively).<sup>4</sup>

### EVALUATION

The evaluation of the patient with OAB should focus on the history, the physical examination, and a limited laboratory evaluation. During the physical examination, it is useful to pay attention to items that may be transient or reversible.

#### Screen for symptoms

The history may be the most important component in the evaluation of the patient with OAB, and the symptoms of urgency, frequency, nocturia, and UI are paramount. Screening for OAB requires minimal time from a provider, as a self-administered screener or questionnaire can be used in most clinical settings.<sup>22</sup> In this context, a screening tool or questionnaire is not meant to diagnose OAB or UI, but rather to identify symptoms that may require treatment. Onset, duration, severity, and bother can be noted with a few key questions. **Table 1** lists examples of questions that may be useful.<sup>23</sup>

#### Components of the history

**A full neurologic history** should be taken to explore the possibility of dementia, Parkinson disease, spinal cord injury or stenosis, multiple sclerosis, or stroke. Functional and cognitive assessment should be performed in older patients.

**The gastrointestinal history** is important, as constipation can cause OAB and the medications used to treat OAB can cause or exacerbate constipation.<sup>24,25</sup>

**Dietary habits**, especially regarding fluid intake, have long been thought to be associated with urinary symptoms and should be addressed in the history.<sup>2</sup> A relationship between LUTS and consumption of caffeinated beverages or alcohol is often seen in clinical

**TABLE 1**  
Simple screening questions for evaluation of overactive bladder (OAB) and incontinence

- Do you get sudden urges to go to the bathroom that are so strong you can't ignore them? (suggests OAB)
- How often do you go to the bathroom? More than eight times in a 24-hour period? (suggests OAB)
- Do you have uncontrollable urges to urinate that sometimes result in wetting accidents? (suggests UUI)
- Do you leak urine on the way to the bathroom? (suggests UUI)
- Do you frequently get up two or more times during the night to go to the bathroom? (suggests OAB)
- Do you avoid places that you think won't have a nearby restroom? (suggests OAB or UUI)
- When you're in an unfamiliar place, do you make sure you know where the restroom is? (suggests OAB or UUI)
- Do you leak urine when you laugh, cough, or sneeze? (suggests SUI)
- Do you use absorbent pads to keep from wetting your clothes? (suggests SUI or UUI)

UUI = urge urinary incontinence; SUI = stress urinary incontinence  
Adapted, with permission, from Newman DK, Giovannini D. The overactive bladder: a nursing perspective. *Am J Nurs* 2002; 102:36–45.<sup>23</sup>

practice, although there is a lack of clear-cut data supporting such a relationship, except with tea consumption.<sup>26</sup>

**Prior surgeries** need to be considered, especially any genitourinary interventions (eg, hysterectomy or bladder suspensions). Orthopedic procedures can be the cause of transient OAB as a result of temporary mobility issues.

**Obstetrical history** should be addressed in women, as a history of several or difficult vaginal deliveries can predispose a woman to OAB or stress UI.

**Medications** should be reviewed to explore a potential association with symptoms. For example, the timing of when a diuretic is taken can have profound effects on urinary habits.<sup>2</sup> Medications that can affect urinary function are listed in **Table 2**.<sup>27</sup>

**Pearls for conducting the physical exam**

The physical examination should focus on detecting anatomic and neurologic abnormalities that could contribute to the patient's symptoms. The neurologic examination should start by observing the patient's gait as he or she walks into the room or down the hall. Limping, poor coordination, dysarthria, facial asymmetry, or other

**TABLE 2**  
Medications that affect bladder function

Medication	Effect
ACE inhibitors	Cough leading to stress UI
Alpha-adrenergic agonists	Increased urethral resistance, causing postvoid dribbling, straining, hesitancy in urine flow
Alpha-receptor agonists (pseudoephedrine, ephedrine)	Urethral constriction, urinary retention (males)
Alpha-receptor antagonists	Urethral relaxation and decreased urethral resistance, causing stress UI (females) and UI with cough, sneeze, or other activity
Anticholinergics (H1 antihistamines, antiparkinsonian agents)	Urinary retention with symptoms of postvoid dribbling, straining, hesitancy in urine flow, overflow incontinence, fecal impaction
Antipsychotics/sedatives	Sedative effect, causing confusion; may relax detrusor muscle, leading to urinary retention
Beta-receptor antagonists	Urinary retention
Calcium channel blockers	Urinary retention, fecal impaction
Diuretics	Increased urine production (polyuria) and volume, leading to urgency and frequency
Methylxanthines (caffeine, theophylline)	Polyuria, bladder irritation
Neuroleptics (thioridazine, chlorpromazine)	Anticholinergic effect, sedation
Other (caffeine and alcohol)	Diuretic effect, leading to urgency and frequency; possible sedation
Opioids	Urinary retention, fecal impaction, sedation, delirium
Sedatives and hypnotics	Sedative effect, which may relax detrusor muscle
Tricyclic antidepressants	Anticholinergic and alpha-receptor antagonist effects, causing postvoid dribbling, straining, hesitancy in urine flow

UI = urinary incontinence  
Adapted from Newman.<sup>27</sup>

findings may indicate neurologic conditions such as a stroke or multiple sclerosis. A brief mental status examination can be performed by observing the patient's general appearance and his or her response to questions. Alertness, orientation, memory, and thought content can be useful parameters in patient assessment.



If the patient appears overweight, consider calculating the body mass index (BMI). The relationship between increased BMI ( $> 30 \text{ kg/m}^2$ ) and the likelihood of UI in females is strong.<sup>26</sup> Identifying this correlation provides an opportunity for the PCP to discuss lifestyle changes with the patient, as research has shown that moderately obese women who lose 5% to 10% of their weight have a decrease in LUTS.<sup>28</sup>

Check the abdomen for masses, hernias, or a distended bladder. In women, the genitalia should be assessed for abnormalities such as prolapse of the bladder or uterus, atrophic vaginitis, or urogenital atrophy, and rectal sphincter tone should be checked. It may be useful to examine the female patient when her bladder is full in order to identify stress UI. In men, assess prostate size (an enlarged prostate can lead to OAB symptoms), the penis and scrotum for abnormalities (such as urethral discharge, epididymitis, or even urethral stricture), and rectal sphincter tone. A basic neurologic examination focusing on motor and sensory components (eg, anal wink and bulbocavernosus reflex) should be performed.

A voiding diary is a simple and practical method of obtaining detailed information about a patient's voiding habits,<sup>2</sup> and can be helpful in evaluating the extent of the problem and offering clues on how best to proceed with evaluation and treatment. The diary or log should be structured to keep track of voiding, urgency, and UI patterns over a 3-day period<sup>29</sup> (Figure 1).

### Judicious use of ancillary studies

The number of ancillary studies required is controversial. It may be prudent to check chemistries, especially renal function and blood glucose, in certain patients. For example, the onset of polyuria/polydipsia in the diabetic patient could certainly mimic the symptoms of OAB. Furthermore, if the clinician suspects obstruction in a man, renal function studies may identify upper urinary tract involvement. A urinalysis should be performed to rule out urinary tract infection. Many elderly women will have asymptomatic bacteriuria that does not require treatment. The role of further studies is questionable.

Checking the patient's postvoid residual urine volume using portable ultrasonographic equipment is useful for detecting retention, but it has a limited role in the neurologically normal female. In the male, it becomes more important because the symptoms of a large, obstructing prostate are similar to those of OAB. A postvoid residual volume check is necessary in any patient in whom there is concern about incomplete bladder emptying as a result of neurologic dysfunction, anatomic abnormality, or a pharmacologic cause. It is

VOIDING RECORD					
Time interval	Urinated in toilet	Amount of incontinent episode (large/small)	Reason for urine leakage	Changed wet pad (damp/wet/saturated)	Type/amount of liquid intake
6 AM					
7 AM					
8 AM					
9 AM					
10 AM					
11 AM					
NOON					
1 PM					
2 PM					
3 PM					
4 PM					
5 PM					
6 PM					
7 PM					
8 PM					
9 PM					
10 PM–MIDNIGHT					
MIDNIGHT–2 AM					
2–4 AM					
4–6 AM					
Type of pad _____		No. pads used _____			
Used with permission of Diane K. Newman.					

**FIGURE 1.** Example of a voiding diary to assess symptoms of overactive bladder.

also necessary in the postoperative patient who develops OAB. The incidence of postoperative urinary retention is recognized but poorly understood, with rates ranging from 4% to 25%.<sup>18</sup> Some patients may present in frank retention, whereas others may present with OAB symptoms from incomplete emptying.

Urodynamic studies are not necessary in most patients, especially those without neurologic compromise. If the patient's symptoms are refractory to therapy, if they worsen, or if there is significant postvoid residual volume, then urodynamic studies may be considered as one looks for other causes, such as detrusor sphincter dyssynergia. In our view, a significant postvoid residual volume is any amount greater than 75 to 100 mL in persons younger than 65 years; in the elderly, we consider a volume less than 150 mL to be acceptable. There are no studies supporting specific values for acceptable postvoid residual volume, but there are guidelines.<sup>30</sup>

Cystoscopy has a role only in the patient with hematuria or the patient who is refractory to therapy.

**TABLE 3**  
**Differential diagnosis of symptoms suggestive of overactive bladder**

In women	In men
Urinary tract infection	Benign prostatic hyperplasia
Prolapse	Bladder outlet obstruction
Urethral obstruction	Urethral stricture
Atrophic vaginitis	Bladder stones
Bladder cancer	Bladder cancer
Interstitial cystitis	Prostate cancer
Postsurgical incontinence	Diabetes
Diabetes	Congestive heart failure
Congestive heart failure	Medications/diuretics
Multiple sclerosis	Neurogenic bladder
Medications/diuretics	Postsurgical incontinence
Neurogenic bladder	
Recent pelvic surgery	
Stress urinary incontinence	

Radiologic evaluation beyond portable bladder ultrasonography is reserved for those with hematuria or a palpable mass noted on examination.

**DIFFERENTIAL DIAGNOSIS**

As mentioned, the diagnosis of OAB can be based on symptoms. As with any symptom complex, however, a differential diagnosis should be considered. A history combined with a directed physical examination, urinalysis, and chemistries will exclude most of the alternate diagnoses listed in **Table 3**. **Table 4** presents findings that should prompt further evaluation or referral to a specialist.

**TREATMENT**

The goal of treatment is to teach the patient to inhibit urgency and to improve voluntary control over bladder function.

**Behavioral treatment**

Behavioral modification involves educating patients about the normal process of micturition and how their specific symptoms define an abnormal situation. If patients are actively involved in the diagnosis and subsequent treatment, their expectations are more readily attainable.

Behavioral therapy may involve pelvic floor muscle exercises, bladder retraining and urge-suppression

techniques, changing the timing of various medications (eg, diuretics), or encouraging exercise and weight loss. Although most patients will be treated using drug therapy, the combination of behavioral and pharmacologic therapies greatly enhances the likelihood of a positive outcome compared with either intervention alone. Burgio et al conducted a crossover study among older women with UI to assess the effects of behavioral therapy, drug therapy, and their combination on patients' baseline frequency of UI episodes.<sup>31</sup> Patients receiving behavioral therapy alone in the initial study phase had a 57% reduction in the frequency of UI, which increased to an 89% reduction after the addition of drug therapy. Similarly, patients receiving drug therapy alone in the initial phase had a 73% reduction in the frequency of UI, which increased to an 84% reduction after the addition of behavioral therapy. The authors concluded that combination therapy yields better outcomes.<sup>31</sup>

**Pharmacologic management: Antimuscarinics are first-line**

The principle behind pharmacologic management of OAB is inhibition of the disturbed bladder contraction, and the antimuscarinics are the primary medications used for this effect.

Antimuscarinics exert their clinical effect through two potential pathways: one on the motor pathway via central and peripheral actions that block a facilitatory mechanism and stimulate an inhibitory mechanism; and the other on the sensory pathway via central and peripheral actions that modulate afferent innervations.<sup>19</sup>

As a class, antimuscarinics are safe and effective. Comparisons among these agents have been limited, but, as with any drug class, there are subtle differences that PCPs should be aware of. Dose adjustment, side effects, or metabolism may be important to consider on an agent-by-agent basis for the individual patient. The various antimuscarinics and their properties are detailed in **Table 5**.<sup>32</sup> All of these agents are administered orally; in addition, oxybutynin is also available for delivery by transdermal patch.

Most antimuscarinics have not been directly compared in clinical trials, and outcome measures and patient characteristics differ between trials, making comparisons difficult. Two comparative efficacy studies deserve attention, the OPERA (Overactive Bladder: Performance of Extended Release Agents) trial and the STAR (Solifenacin and Tolterodine as an Active Comparator in a Randomised) trial.<sup>33,34</sup>

The **OPERA trial** compared the long-acting oral versions of oxybutynin (10 mg) and tolterodine (4 mg). There was a reduction of UI with oxybutynin

over tolterodine, but it came at a slight and almost proportional increase in dry mouth.<sup>33</sup>

The STAR trial compared a single dosage strength of long-acting tolterodine (4 mg) with a flexible-dose regimen of solifenacin (5 or 10 mg). Using pooled data for the two solifenacin doses, the study found that titratable solifenacin was associated with slightly better efficacy than tolterodine with an almost proportional increase in side effects.<sup>34</sup>

Another recent head-to-head study compared the effects of darifenacin (7.5 mg) and long-acting oral oxybutynin (10 mg) on cognitive end points in subjects 60 years of age or older.<sup>35</sup> The premise of the study was that an agent with selectivity for the M3 receptor (the prominent muscarinic receptor on the bladder) would not affect the M1 receptors in the brain. The outcome studied was performance on the Name-Face Association Test, which measures delayed recall, at week 3 of treatment. Subjects randomized to oxybutynin performed statistically worse on this test than did placebo recipients, indicating significant memory impairment, whereas no reduction in performance was seen in darifenacin recipients.<sup>35</sup>

### Secondary medications

Although the antimuscarinic class is the first line of pharmacologic therapy for OAB, other medications have a role. There is some evidence that the stress component of mixed UI may respond to the tricyclic antidepressant imipramine or to an alpha-adrenergic agonist such as pseudoephedrine, which increases outlet resistance.<sup>36</sup> Furthermore, these drugs may work synergistically with antimuscarinic therapy in patients with mixed UI. However, imipramine is not indicated in the elderly and pseudoephedrine should be used with caution in the elderly. Treatment should be geared to the symptom that appears to be most bothersome.<sup>37</sup> Neither imipramine nor pseudoephedrine is approved by the US Food and Drug Administration for treating OAB or stress UI.

Transvaginal estrogen therapy also may have a role in treating the irritative symptoms of urgency and frequency associated with vaginal and urogenital atrophy; however, data are lacking to support any particular dosing regimen, route of administration, or treatment duration.<sup>38</sup> A recent analysis from the Women's Health Initiative found that oral estrogen replacement not only failed to improve UI but may actually worsen symptoms.<sup>39</sup>

### Risk of urinary retention with therapy is low

A common concern among PCPs is that antimuscarinic therapy may place a patient at risk of urinary

**TABLE 4**  
Red flags that should prompt further studies or referral to a specialist

Uncertain diagnosis and inability to develop a reasonable management plan
Lack of response to an adequate trial of conservative therapies (eg, bladder training, pelvic muscle exercises, and drug therapy)
Hematuria without infection
Severe (beyond the introitus) pelvic organ prolapse
Abnormal postvoid residual urine volume
Prostate nodule/enlargement
Neurologic condition (eg, multiple sclerosis, spinal cord lesions) in which a component of neurogenic bladder is suspected
History of pelvic surgery

Adapted from Fantl et al.<sup>30</sup>

retention. However, the incidence of retention in both men and women in clinical trials is low.<sup>19,40</sup> Kaplan et al evaluated the safety and efficacy of antimuscarinic therapy in men with BPH and LUTS in whom alpha-antagonist therapy failed to relieve LUTS.<sup>40</sup> In the 39 men who completed this 6-month trial, there was a significant decrease in urinary frequency, nocturia, and postvoid residual volume, as well as an improvement in symptom scores. There were no reports of urinary retention. These findings suggest that the inhibitory effect of antimuscarinic agents on detrusor muscle contraction is unlikely to aggravate voiding difficulties in men with OAB symptoms and possible obstruction.

We believe it is prudent to do a postvoid residual volume check in a man being treated for LUTS, both initially and at follow-up. However, this practice is controversial, and the most appropriate candidates for these checks remain to be better defined.

### Symptom-based treatment can be successful

This paper has addressed the empiric diagnosis and treatment of OAB. The concept of empiric diagnosis and treatment was assessed in the recently published IMPACT (Improvement in Patients: Assessing Symptomatic Control with Tolterodine) trial.<sup>41</sup> In this study, the diagnosis of OAB was made in several hundred patients in primary care and obstetric/gynecologic offices on the basis of symptoms. Patients with OAB symptoms for at least 3 months were treated with extended-release tolterodine for 12 weeks in this

**TABLE 5**  
**Properties of available antimuscarinic agents**

	Darifenacin	Oxybutynin	Oxybutynin TDS	Solifenacin	Tolterodine	Trospium
Chemical structure	Tertiary amine	Tertiary amine	Tertiary amine	Tertiary amine	Tertiary amine	Quaternary amine
Efficacy	Not all products have been directly compared in head-to-head trials; outcome measures and patient baseline characteristics differ among trials, making comparisons difficult. Efficacy similar among products: 60%–75% median reduction of urge urinary incontinence episodes.					
Side effects	Dry mouth, constipation	Dry mouth, constipation	Skin irritation	Dry mouth, constipation	Dry mouth, constipation	Dry mouth, constipation
Dose	7.5 or 15 mg	5, 10, or 15 mg	3.9 mg	5 or 10 mg	2 or 4 mg	20 mg
Regimen	Once daily	IR: twice daily to four times daily ER: once daily	Twice weekly	Once daily	Once daily	Twice daily
Half-life	12.43 hr (7.5 mg) 12.05 hr (15 mg)	13.2 hr (10 mg)	36 hr	45–68 hr	8.8 hr (active metabolite)	12.2 ± 3.2 hr
Metabolism	Extensively metabolized in liver by CYP450 isoforms	Extensively metabolized in liver by CYP450 isoforms	No first-pass clearance, but then extensively metabolized in liver by CYP450 isoforms	Extensively metabolized in liver by CYP450 isoforms	Extensively metabolized in liver by CYP450 isoforms	Active renal tubular secretion is a major route of elimination; global hepatic esterification only, not CYP450
Potential metabolic drug-drug interactions	Flecainide, thioridazine, TCAs; do not exceed 7.5 mg when given with ketoconazole, itraconazole, ritonavir, nelfinavir, clarithromycin, or nefazodone	Itraconazole, miconazole, macrolide antibiotics (eg, erythromycin, clarithromycin)	Itraconazole, miconazole, macrolide antibiotics (eg, erythromycin, clarithromycin)	Do not exceed 5 mg when given with ketoconazole or other potent inhibitors of CYP3A4 (eg, nefazodone, fluvoxamine, TCAs)	Ketoconazole, itraconazole, miconazole, macrolide antibiotics (eg, erythromycin, clarithromycin), cyclosporine, vinblastine	None; active renal tubular excretion; caution if giving with digoxin, metformin, or other drugs that may compete for active renal excretion
Contra-indications	Urinary retention; gastric retention and other conditions of severe decreased gastrointestinal motility; uncontrolled narrow-angle glaucoma; hypersensitivity to the drug or its ingredients					

TDS = transdermal delivery system; IR = immediate-release; ER = extended-release; CYP = cytochrome P; TCAs = tricyclic antidepressants  
 Data are based on the medications' package inserts.<sup>32</sup>

open-label, single-arm trial. At the end of the 12 weeks, there were significant reductions from baseline in urge UI, urgency episodes, nocturnal frequency, and daytime frequency. Common side effects were dry mouth and constipation.

The conclusion drawn from this paper was that patients with OAB in the general population can be readily screened and successfully treated with minimal work-up.<sup>41</sup> The applicability to the PCP is significant, but the study lasted only 12 weeks and longer follow-up would have been useful.

### ■ APPROPRIATE FOLLOW-UP

There is no set rule as to the follow-up interval for the OAB patient. Some clinicians find that a 2-week interval is adequate, whereas others recommend 4

weeks. The interval should be determined by consensus of the provider and the patient. The patient must be educated about what to expect and to not give up hope if these expectations are not met immediately, as a simple drug change or dose titration may provide the desired effect. There are many medication choices, and no one treatment is right for every patient.

### ■ WHEN TO REFER

As noted above, the diagnosis of OAB can be made empirically without the need for specialized evaluation, and treatment likewise can be initiated comfortably by the PCP. If initial treatment with behavioral therapy and medications fails to alleviate the symptoms, it is appropriate to refer the patient for consultation and advanced testing (Table 4), such as urody-



namical evaluation.<sup>30</sup> The definition of “initial treatment” may vary among clinicians. Some may try only one medication, whereas others may be comfortable changing medications and titrating doses. Our view is that any medication should be given for 2 to 4 weeks before it is considered to have failed.

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