# Vaginitis: Diagnosis and Treatment

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Bacterial vaginosis, trichomoniasis, and vulvovaginal candidiasis are the most common infectious causes of vaginitis. Bacterial vaginosis occurs when the normal lactobacilli of the vagina are replaced by mostly anaerobic bacteria. Diagnosis is commonly made using the Amsel criteria, which include vaginal pH greater than 4.5, positive whiff test, milky discharge, and the presence of clue cells on microscopic examination of vaginal fluid. Oral and topical clindamycin and metronidazole are equally effective at eradicating bacterial vaginosis. Symptoms and signs of trichomoniasis are not specific; diagnosis by microscopy is more reliable. Features of trichomoniasis are trichomonads seen microscopically in saline, more leukocytes than epithelial cells, positive whiff test, and vaginal pH greater than 5.4. Any nitroimidazole

drug (e.g., metronidazole) given orally as a single dose or over a longer period resolves 90 percent of trichomoniasis cases. Sex partners should be treated simultaneously. Most patients with vulvovaginal candidiasis are diagnosed by the presence of vulvar inflammation plus vaginal discharge or with microscopic examination of vaginal secretions in 10 percent potassium hydroxide solution. Vaginal pH is usually normal (4.0 to 4.5). Vulvovaginal candidiasis should be treated with one of many topical or oral antifungals, which appear to be equally effective. Rapid point-of-care tests are available to aid in accurate diagnosis of infectious vaginitis. Atrophic vaginitis, a form of vaginitis caused by estrogen deficiency, produces symptoms of vaginal dryness, itching, irritation, discharge, and dyspareunia. Both systemic and topical estrogen treatments are effective. Allergic and irritant contact forms of vaginitis can also occur. (*Am Fam Physician.* 2011;83(7):807-815. Copyright © 2011 American Academy of Family Physicians.)

Patient information: A handout on vaginitis, written by the authors of this article, is provided on page 816. aginitis is defined as a spectrum of conditions that cause vaginal and sometimes vulvar symptoms, such as itching, burning, irritation, odor, and vaginal discharge. Vulvovaginal complaints are one of the most common reasons for women to seek medical advice.

# **Etiology and Diagnosis of Vaginitis**

The most common infectious causes of vaginitis are bacterial vaginosis, vulvovaginal candidiasis, and trichomoniasis. Physicians traditionally diagnose vaginitis using the combination of symptoms, physical examination, pH of vaginal fluid, microscopy, and the whiff test. When combined, these tests have a sensitivity and specificity of 81 and 70 percent, respectively, for bacterial vaginosis; 84 and 85 percent for vulvovaginal candidiasis; and 85 and 100 percent for trichomoniasis when compared with the DNA probe standard.<sup>1</sup> *Table 1* describes common



causes, symptoms, and signs of vaginitis,<sup>2,3</sup> and *Table 2* lists risk factors that contribute to the development of the condition.<sup>3-11</sup>

In a review of studies published between 1966 and 2003, bacterial vaginosis was diagnosed in 22 to 50 percent of symptomatic women, vulvovaginal candidiasis in 17 to 39 percent, and trichomoniasis in 4 to 35 percent. Approximately 30 percent of symptomatic women remained undiagnosed after clinical evaluation.<sup>2</sup> Among multiple individual symptoms and signs, only the following were found to be helpful for the diagnosis of vaginitis in symptomatic women:

• A lack of itching makes diagnosis of vulvovaginal candidiasis unlikely (range of likelihood ratios [LRs], 0.18 [95% confidence interval (CI), 0.05 to 0.70] to 0.79 [95% CI, 0.72 to 0.87]).

• A lack of perceived odor makes bacterial vaginosis unlikely (LR, 0.07 [95% CI, 0.01 to 0.51]).

Clinical recommendation	Evidence rating	Reference
Clindamycin and metronidazole (Flagyl, Metrogel) are equally effective for eradicating symptoms of bacterial vaginosis.	А	33
Nitroimidazole drugs (e.g., metronidazole) given orally in a single dose or over a longer period result in parasitologic cure of trichomoniasis in 90 percent of cases.	A	44
Oral and vaginal antifungals are equally effective for the treatment of uncomplicated vulvovaginal candidiasis.	A	47
All methods of estrogen delivery relieve the symptoms of atrophic vaginitis.	А	49

A = consistent, good-quality patient-oriented evidence; B = inconsistent or limitedquality patient-oriented evidence; C = consensus, disease-oriented evidence, usual practice, expert opinion, or case series. For information about the SORT evidence rating system, go to http://www.aafp.org/afpsort.xml.

• Presence of inflammatory signs is more commonly associated with vulvovaginal candidiasis (range of LRs, 2.1 [95% CI, 1.5 to 2.8] to 8.4 [95% CI, 2.3 to 3.1]).

• Presence of a fishy odor on examination is predictive of bacterial vaginosis (LR, 3.2 [95% CI, 2.1 to 4.7]).

• Lack of odor is associated with vulvovaginal candidiasis (LR, 2.9 [95% CI, 2.4 to 5.0]).<sup>2</sup>

Individual symptoms and signs, pH level, and microscopy results often do not lead to an accurate diagnosis of vaginitis. Laboratory tests perform better than standard office-based evaluation for diagnosing causes of vaginitis,<sup>2</sup> but they do not add substantially to the treatment threshold and are justified only in patients with recurrent or difficult-to-diagnose symptoms. *Table 3* describes laboratory tests used to diagnose infectious causes of vaginitis.<sup>2,12-24</sup>

A cost-effectiveness analysis of diagnostic strategies for vaginitis undiagnosed by pelvic examination, wet-mount preparation, and related office tests showed that the least expensive strategy was to perform yeast culture, gonorrhea and chlamydia probes at the initial visit, and Gram stain and *Trichomonas* 

culture only when the vaginal pH exceeded 4.9. Other strategies cost more and increased duration of symptoms by up to 1.3 days.<sup>25</sup>

# **Bacterial Vaginosis**

Bacterial vaginosis is the most prevalent cause of vaginal discharge or malodor, occurring in up to 30 percent of women.<sup>26</sup> It occurs when the normal *Lactobacillus* species in the vagina are replaced with anaerobic bacteria,

Туре	Etiology	Clinical symptoms			
		Discharge	Pain	Pruritus	
Bacterial vaginosis	Gardnerella vaginalis, Mycoplasma hominis Anaerobic bacteria: Prevotella species, Mobiluncus species	Malodorous; homogenous; clear, white, or gray; fishy odor	Not primary symptom	Not primary symptom	
Trichomoniasis	Trichomonas vaginalis	Green-yellow, frothy	Pain with sexual intercourse, vaginal soreness, dysuria	Not primary symptom	
Vulvovaginal candidiasis	Candida albicans, Candida krusei, Candida glabrata	White, thick, lack of odor	Burning, dysuria, dyspareunia	Frequent	
Atrophic vaginitis	Estrogen deficiency	Yellow, greenish, lack of odor	Vaginal dryness, pain with sexual intercourse	Rare	
Erosive lichen planus	Etiology is unknown	Yellow or gray	Intense pain, dyspareunia, postcoital bleeding	Intense	
rritant or allergic contact dermatitis	Contact irritation or allergic reaction with episodic flares	Minimal	Burning on acute contact, soreness	More likely in allergic reactions	

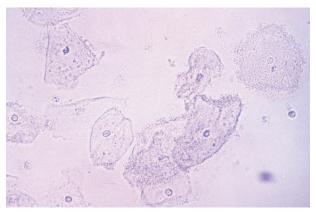
## Table 1. Causes, Symptoms, and Signs of Vaginitis

## Table 2. Risk Factors Contributing to Vaginitis

Type of vaginitis	Risk factors
Bacterial vaginosis	Low socioeconomic status, vaginal douching, smoking, use of an intrautering contraceptive device, new/multiple sex partners, unprotected sexual intercourse, homosexual relationships, frequent use o higher doses of spermicide nonoxynol-9
Trichomoniasis	Low socioeconomic status, multiple sex partners, lifetime frequency of sexual activity, other sexually transmitted infections, lack of barrier contraceptive use, illicit drug use, smoking
Vulvovaginal candidiasis	Vaginal or systemic antibiotic use, diet high in refined sugars, uncontrolled diabetes mellitus
Atrophic vaginitis	Menopause, other conditions associated with estrogen deficiency, oophorectomy, radiation therapy, chemotherapy, immunologic disorders, premature ovaria failure, endocrine disorders, antiestrogen medication
Irritant contact dermatitis	Soaps, tampons, contraceptive devices, sex toys, pessary, topical products, douching, fastidious cleansing, medications, clothing
Allergic contact dermatitis	Sperm, douching, latex condoms or diaphragms, tampons, topical products, medications, clothing, atopic history

Information from references 3 through 11.

Clinical signs	
Vagina	Vulva
No signs of inflammation	Unaffected
Signs of inflammation, "strawberry cervix"	Vestibular erythema may be present
Signs of inflammation, edema	Excoriations
Vagina mildly erythematous, easily traumatized	Vestibule thin and dry; labia majora lose their subcutaneous fat; labia minora irritated and friable
Erythema with friable epithelium	Erosions, white plaques
Vulvar erythema possible	Erythema with or without edema; vesicles or bullae rare



**Figure 1.** Clue cells (400 ×). Vaginal epithelial cells with borders obscured by adherent coccobacilli seen on saline wet-mount preparation.

resulting in reduced levels of hydrogen peroxide and organic acids usually present in the vagina.

The underlying cause of bacterial vaginosis is not fully understood. More than 50 percent of women with bacterial vaginosis are asymptomatic. The fishy odor caused by production of amines from anaerobic bacteria found in many of these patients is predictive of bacterial vaginosis.<sup>2</sup> When vaginal alkalinity increases after sexual intercourse (with the presence of semen) and during menses (with the presence of blood), the odor becomes more prevalent.<sup>27</sup> Vaginal discharge is a more common but less specific symptom. Bacterial vaginosis is not associated with vaginal mucosal inflammation and rarely causes vulvar itch.<sup>27</sup>

Bacterial vaginosis, even when asymptomatic, is associated with a high incidence of endometritis and pelvic inflammatory disease following abortion and gynecologic procedures in the general population. Among women with bacterial vaginosis, no overall increased risk of developing pelvic inflammatory disease has been found.<sup>28</sup> Bacterial vaginosis is associated with late miscarriages, premature rupture of membranes, and preterm birth.<sup>29</sup> Both symptomatic and asymptomatic bacterial vaginosis have been strongly linked with an increased risk of human immunodeficiency virus (HIV)-1 transmission (relative risk, 1.89; 95% CI, 1.46 to 2.43).<sup>30</sup>

## DIAGNOSIS

In clinical practice, bacterial vaginosis is diagnosed by the presence of three out of four Amsel criteria<sup>31</sup>:

- Thin, homogenous vaginal discharge
- Vaginal pH greater than 4.5

• Positive whiff test (fishy amine odor when 10 percent potassium hydroxide solution is added)

• At least 20 percent clue cells (vaginal epithelial cells with borders obscured by adherent coccobacilli on wetmount preparation or Gram stain; *Figure 1*).

In a prospective observational study of 269 women, a vaginal pH of more than 4.5 was found to be the most

# Table 3. Laboratory Testing for Infectious Causes of Vaginitis

Test	Bacterial vaginosis	Trichomoniasis
Point-of-care tests*		
Amsel criteria	Sensitivity, 69%; specificity, 93%	NA
рН	pH > 5: sensitivity, 77%; specificity, 35%	pH > 5.4: sensitivity, 92%; specificity, 51%
Whiff test (the amine odor produced by mixing 10% potassium hydroxide solution with a sample of vaginal discharge)	Positive test: sensitivity, 67%; specificity, 93%	Positive test: sensitivity, 67%; specificity, 65%
Fem Exam card (Cooper Surgical, Shelton, Conn.) Two colorimetric strips: card 1 measures pH and amine levels; card 2 measures proline aminopeptidase activity	Cards 1 and 2 combined: sensitivity, 91%; specificity, 61% Rapid (two minutes), less subjective than whiff test, easily performed	NA
Microscopy (with 10% potassium hydroxide solution, saline)	Clue cells, bacilli with corkscrew motility, scant or absent lactobacilli Sensitivity, 53 to 90%; specificity, 40 to 100%	Motile protozoa with flagella; more leukocytes than epithelial cells Sensitivity, 50 to 70% (may be increased by vaginal lavage to 74%); specificity, 100%
pH, trimethylamine card (QuickVue Advance Quidel, San Diego, Calif.)	Sensitivity, 53%; specificity, 97% Rapid, simple, comparable with pH and whiff test	ΝΑ
Proline aminopeptidase card (Pip Activity TestCard, Litmus Concepts, Inc., Santa Clara, Calif.) Indirect test for a chemical produced by the organisms associated with bacterial vaginosis	Sensitivity, 70%; specificity, 81% Rapid, simple, comparable with Fem Exam card 2	ΝΑ
OSOM Trichomonas Rapid Test (Genzyme Diagnostics, Cambridge, Mass.), uses color immunochromatographic "dipstick" technology with murine monoclonal antibodies	NA	Sensitivity, 90 to 100% 10 minutes to complete test
BD Affirm VPIII Microbial Identification Test (BD Diagnostic Systems, Sparks, Md.)	Sensitivity, 95 to 100% 45 minutes to complete test	Sensitivity, 90 to 100% False-positive results may occur, especially in low-prevalence groups
Polymerase chain reaction: based on DNA amplification (Quest Diagnostics, Madison N.J.), Instagene Matrix (Bio-Rad Laboratories, Hercules, Calif.)	Effective at identifying bacteria responsible for bacterial vaginosis	Sensitivity, 80%; specificity, 97%
Reference laboratory testing* Culture	Predictive value of a positive <i>Gardnerella vaginalis</i> culture is less than 50%; generally not recommended, but may have value in recalcitrant cases	<ul> <li>InPouch Culture System (Biomed, White City, Ore.)</li> <li>Combined wet-mount preparation and culture kit; can be kept at room temperature for up to 18 hours; samples taken during menses were not adversely affected</li> <li>Sensitivity, 81 to 100%</li> </ul>
Patient-performed tests		
Over-the-counter test for vaginal infections (Fem-V; Synova Healthcare, Inc., New York, NY)	Positive test suggests possibility of bacterial vaginosis and need for physician visit (20% false-positive rate)	Positive test suggests possibility of trichomoniasis and need for physician visit (20% false-positive rate)
Over-the-counter rapid yeast detection test (Savyon Diagnostics, Ashdod, Israel): uses the concept of lateral flow immunoassay systems	ΝΑ	ΝΑ

\*—Relative cost of testing does not include physician office charges. Information from references 2, and 12 through 24.

Vulvovaginal candidiasis	Cost*
NA	\$\$
pH < 4.9: sensitivity, 71%; specificity, 90%	\$
Negative test	\$
NA	\$\$

Budding filaments, mycelia with 10% potassium hydroxide solution Sensitivity, 61%; specificity, 77%	\$
NA	\$
NA	\$
NA	\$\$\$
Sensitivity, 90 to 100%	\$\$\$
Polymerase chain reaction more sensitive than culture in detecting <i>Candida</i> ; not yet commercially available as a diagnostic test	\$\$\$
Positive culture alone does not necessarily indicate that the yeast identified are responsible for vaginal symptoms	\$\$\$
Negative test suggests possibility of yeast infection Over-the-counter antifungal treatment recommended (10% false-negative rate)	\$
Positive test: sensitivity, 73%; specificity, 84% Patient-performed tests have results similar to physician-performed tests Negative test: not sensitive enough to rule out yeast infection and avoid a culture	\$

sensitive (89 percent) and a positive whiff test was the most specific (93 percent) method of detecting bacterial vaginosis.<sup>12</sup> The positive presence of these two tests is as sensitive as three or more Amsel criteria.<sup>12</sup> Culture of *Gardnerella vaginalis* is not recommended because of low specificity. Cervical cytology has no clinical value for diagnosing bacterial vaginosis, especially in asymptomatic women, because it has low sensitivity.<sup>13</sup>

## TREATMENT IN NONPREGNANT WOMEN

Current treatment recommendations from the Centers for Disease Control and Prevention (CDC) are listed in *Table 4*.<sup>13,32</sup> Nonpregnant women with symptomatic disease require antibacterial therapy to relieve vaginal symptoms. Other benefits of treatment include decreasing the risk of HIV and other sexually transmitted infections and reducing infectious complications following abortion or hysterectomy.<sup>13</sup>

A Cochrane review of 24 randomized controlled trials (RCTs) showed that clindamycin and metronidazole (Flagyl) are equally effective, achieving clinical cure in 91 and 92 percent of cases, respectively, after two to three weeks of treatment.<sup>33</sup> Six RCTs showed topical and oral antibiotic preparations to be equally effective. One disadvantage of oral regimens is a longer duration of treatment.<sup>33</sup> Intravaginal clindamycin cream is preferred in case of allergy or intolerance to metronidazole. Metronidazole in a single 2-g dose has the lowest effectiveness for treating bacterial vaginosis and is no longer recommended. Metronidazole, 500 mg twice daily for one week, is effective for treating bacterial vaginosis and trichomoniasis.

Although lactobacillus probiotics are safe, there is no conclusive evidence that they are superior to or enhance the effectiveness of antibiotics in the treatment of bacterial vaginosis or prevent its recurrence.<sup>34</sup> Treatment of sex partners and follow-up visits if symptoms are resolved are not recommended.

## BACTERIAL VAGINOSIS IN PREGNANCY

Bacterial vaginosis is present in up to 20 percent of women during pregnancy. The effect of treating bacterial vaginosis in symptomatic or asymptomatic pregnant women on subsequent preterm delivery has produced conflicting results in clinical trials.<sup>13,35</sup> The U.S. Preventive Services Task Force (USPSTF) recommends against routine bacterial vaginosis screening of asymptomatic pregnant women at low risk of preterm delivery (USPSTF grade D recommendation).<sup>36</sup>

#### **RECURRENT BACTERIAL VAGINOSIS**

Most relapses of bacterial vaginosis occur within the first year and strongly correlate with new sex partners.

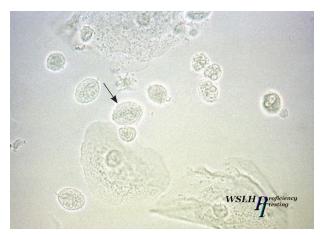
# Table 4. CDC Recommended Treatment of Vaginitis

Medication/dosage	Cost per course of therapy	Medication/dosage	Cost per course of therapy
	шегару		
Bacterial vaginosis		Vulvovaginal candidiasis, uncomplicated (continued)	¢
Recommended regimen	¢.¢	Miconazole 4% cream <sup>†</sup>	\$
Metronidazole (Flagyl)	\$\$	5 g intravaginally once daily for three days	
500 mg orally twice daily for seven days	***	Miconazole vaginal suppository†	¢
Metronidazole gel (Metrogel)	\$\$\$	100-mg vaginal suppository once daily for seven days	\$ ¢¢
One full applicator (5 g) intravaginally once daily for five days		200-mg vaginal suppository once daily for three days 1,200-mg vaginal suppository in a single dose	\$\$ \$\$
Clindamycin 2% cream	\$\$\$	Nystatin vaginal tablet	\$
One full applicator (5 g) intravaginally at bedtime for seven days		100,000-unit vaginal tablet once daily for 14 days Tioconazole 6.5% ointment†	\$\$
Alternative regimen		5 g intravaginally in a single dose	44
Tinidazole (Tindamax)	\$\$\$	Terconazole 0.4% cream	\$\$
2 g orally for two days or 1 g for five days		5 g intravaginally once daily for seven days	44
Clindamycin	\$\$\$	Terconazole 0.8% cream	\$\$
300 mg orally twice daily for seven days		5 g intravaginally once daily for three days	44
Clindamycin ovules	\$\$\$	Terconazole vaginal suppository	\$\$
100 mg intravaginally once daily at bedtime		80-mg vaginal suppository once daily for three days	
for three days		Fluconazole (Diflucan)	\$\$
Pregnancy*		150 mg orally in a single dose	
Metronidazole	\$\$	Vulvovaginal candidiasis, complicated	
500 mg orally twice daily for seven days	**	Recurrent	
Metronidazole	\$\$	Initial regimen	
250 mg orally three times daily for seven days	**	Any topical agent	
Clindamycin	\$\$	Seven to 14 days	
300 mg orally twice daily for seven days		Fluconazole	\$\$\$
Trichomoniasis		100, 150, or 200 mg orally once daily every	444
Recommended regimen		third day for three doses	
Metronidazole	\$	Maintenance regimen	
2 g orally in a single dose		Fluconazole	\$\$\$
Tinidazole	\$\$\$	100, 150, or 200 mg orally once weekly for	
2 g orally in a single dose		six months	
Alternative regimen		Severe	
Metronidazole	\$\$	Any topical azole	\$\$
500 mg orally twice daily for seven days		Intravaginally once daily for seven to 14 days	
Pregnancy* Metronidazole	đ	Fluconazole	\$\$\$
2 g orally in a single dose	\$	150 mg orally once daily in two doses 72 hours apart	
Vulvovaginal candidiasis, uncomplicated		Vulvovaginal candidiasis, nonalbicans	<i>.</i>
Butoconazole 2% cream (Gynazole-1)†	\$\$	Nonfluconazole azole (oral or topical)	\$
5 g intravaginally once daily for three days	<i>.</i>	Seven to 14 days	* *
Butoconazole 2% cream, sustained release	\$	Boric acid gelatin capsule	\$\$
5 g intravaginally in a single dose Clotrimazole 1% cream†	¢	Intravaginally once daily for 14 days	
	\$	Pregnancy*	¢¢
5 g intravaginally once daily for seven to 14 days Clotrimazole 2% cream <sup>†</sup>	\$	Any topical azole Intravaginally once daily for seven days	\$\$
5 g intravaginally once daily for three days	Φ	intravaginally once daily for seven days	
Miconazole 2% cream†	\$		
5 g intravaginally once daily for seven days	Ŷ		
s g intravaginary once daily for seven days			

CDC = Centers for Disease Control and Prevention.

\*—After counseling patients about the potential risks and benefits of treatment, preferably after 37 weeks of gestation. †—Available over the counter.

Information from references 13 and 32.



**Figure 2.** *Trichomonas vaginalis* (400 ×). When vaginal wet-mount preparation is promptly examined, motile trichomonads with flagella slightly larger than a leukocyte may be seen (*arrow*).

Reported recurrence rates are 15 to 30 percent within three months.<sup>37,38</sup> One RCT on persistent bacterial vaginosis indicated that metronidazole gel 0.75% (Metrogel), used twice weekly for six months after initial treatment, effectively maintained a clinical cure for six months.<sup>39</sup>

# Trichomoniasis

Symptoms and signs of trichomoniasis are not specific, and diagnosis by microscopy is more reliable. Features suggestive of trichomoniasis are trichomonads seen with saline, leukocytes more numerous than epithelial cells, positive whiff test, and vaginal pH greater than 5.4.<sup>2</sup> The wet-mount preparation is an inexpensive and quick test with variable sensitivity of 58 to 82 percent,<sup>40</sup> and is influenced by the experience of the examiner and the number of parasites in the vaginal fluid sample (*Figure 2;* a video of motile trichomonads is available at http://www.youtube.com/watch?v=DKJnDz6yBSs&feature=fv sr). Adding examination of the spun urine specimen can increase the detection rate of *Trichomonas vaginalis* from 73 to 85 percent.<sup>41</sup>

Treatment should not be based on a Papanicolaou (Pap) smear finding of trichomonads. Newer point-ofcare tests (*Table 3*<sup>2,12-24</sup>) are more accurate but costly. In one study, the sensitivities of wet-mount preparation, microbiologic culture, rapid antigen testing, and nucleic acid amplification testing were 51, 75, 82, and 98 percent, respectively. Specificity was close to 100 percent for each method.<sup>42</sup> Polymerase chain reaction analysis of samples from tampons and introital specimens is more accurate than vaginal or cervical swabs and Pap smears, and may be preferable for patient comfort.<sup>43</sup>

# TREATMENT

Almost any nitroimidazole drug given orally in a single dose or over a longer period results in parasitologic cure in 90 percent of cases.<sup>44</sup> A single 2-g dose of metronidazole is adequate but can cause dyspepsia and metallic taste; compliant patients may prefer a longer regimen at a lower daily dosage with fewer adverse effects. Metronidazole in a dosage of 500 mg twice daily for seven days will treat bacterial vaginosis and trichomoniasis. Metronidazole in a dosage of 2 to 4 g daily for seven to 14 days is recommended for metronidazole-resistant strains.

The parasitologic cure rate of intravaginal nitroimidazole creams is an unacceptably low 50 percent. In RCTs, combined oral and intravaginal treatments have been more effective than oral treatment alone. Sex partners should be treated simultaneously.<sup>44</sup> To reduce recurrence, partners should avoid resuming sexual intercourse until both have completed treatment and are asymptomatic. Test of cure is not required.<sup>13</sup>

## TRICHOMONIASIS IN PREGNANCY

A Cochrane review found that metronidazole is effective against trichomoniasis when taken by women and their partners during pregnancy. A trial of women treated before 23 weeks of gestation was stopped early because women taking metronidazole were more likely to give birth preterm and have low-birth-weight infants.<sup>45</sup> The CDC (*Table 4*<sup>13,32</sup>) recommends giving 2 g of metronidazole as a single dose, preferably after 37 weeks of gestation, and counseling patients about the potential risks and benefits of treatment.

# **Vulvovaginal Candidiasis**

An estimated 75 percent of women will have at least one episode of vulvovaginal candidiasis, and 40 to 45 percent will have two or more.<sup>46</sup> Changes in the host vaginal environment precipitate and induce pathologic effects of the organism.<sup>14</sup> Several risk factors for vulvovaginal candidiasis are listed in *Table 2*.<sup>3-11</sup>

# DIAGNOSIS

Although symptoms of vulvovaginal candidiasis such as pruritus, vaginal soreness, dyspareunia, and vaginal discharge are common, none of them are specific.<sup>2</sup> Most patients can be diagnosed by microscopic examination (*Figure 3*) of vaginal secretions with a 10% potassium hydroxide solution (sensitivity, 65 to 85 percent). Vaginal pH is usually normal (4.0 to 4.5). Vaginal culture should be considered in recurrently symptomatic women with negative microscopy and a normal pH. The Pap smear, although specific, is insensitive, with positive results in only about 25 percent of patients with culture-positive symptomatic vulvovaginal candidiasis.<sup>14</sup> A rapid yeast detection test (Savyon Diagnostics) can be performed by the patient and costs less than \$10, compared with



**Figure 3.** Candida species (400 ×). Budding yeast visible (arrow).

a mean of \$65 for the yeast culture.<sup>15</sup> Polymerase chain reaction testing is considered the most sensitive method, but is very expensive.

#### TREATMENT

On the basis of clinical presentation, microbiology, host factors, and response to therapy, vulvovaginal candidiasis can be classified as uncomplicated or complicated.<sup>13</sup> Patients with uncomplicated vulvovaginal candidiasis are not pregnant, are otherwise healthy, and have all of the following:

- Mild to moderate disease
- Fewer than four episodes of candidiasis per year
- Pseudohyphae or hyphae visible on microscopy.

Treatment of uncomplicated vulvovaginal candidiasis involves a short course of antifungals<sup>47</sup> (*Table 4*<sup>13,32</sup>); oral and topical preparations are similarly effective.<sup>13,48</sup>

Patients with complicated vulvovaginal candidiasis have one or more of the following:

- Moderate to severe disease
- Four or more episodes of candidiasis per year
- Only budding yeast visible on microscopy

• Adverse host factors (e.g., pregnancy, diabetes mellitus, immunocompromise).

Treatment of complicated vulvovaginal candidiasis involves an intensive, longer course of antifungals (*Table 4*<sup>13,32</sup>).

## **Noninfectious Causes of Vaginitis**

Irritant contact dermatitis and allergic contact dermatitis are two noninfectious causes of vaginitis. They may be associated with use of feminine hygiene products or contraceptive materials, among many other causes. Atrophic vaginitis can manifest clinically with symptoms of vaginal dryness, itching, discharge, irritation, and dyspareunia. It affects 10 to 40 percent of women who have conditions associated with estrogen deficiency.<sup>4</sup> Diagnosis is based on history and physical findings, supplemented by vaginal pH levels, vaginal wetmount preparation (to exclude superimposed infection), and, rarely, culture or cytology. Both systemic and topical estrogen treatments are effective in relieving symptoms. Topical vaginal estrogen is preferred because of the low systemic absorption and reduced risk of adverse effects compared with oral therapy. Estrogen-containing creams, pessaries, intravaginal tablets, and the estradiol vaginal ring appear equally effective for the symptoms of atrophic vaginitis.<sup>49</sup>

**Data Sources:** We reviewed recent relevant publications in the Cochrane Database, InfoPOEMS, and the National Guideline Clearinghouse, and searched Pub Med for vaginitis topics. The search included meta-analyses, randomized controlled trials, clinical reviews, and clinical trials. Terms included vaginitis, trichomoniasis, bacterial vaginosis, candidal vulvovaginitis, and atrophic vaginitis. Search date: March 2010.

Figures 1 through 3 provided by Barbara M. Hill, MT (ASCP), Acting Director, Wisconsin State Laboratory of Hygiene Proficiency Testing.

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#### REFERENCES

- Lowe NK, Neal JL, Ryan-Wenger NA. Accuracy of the clinical diagnosis of vaginitis compared with a DNA probe laboratory standard. *Obstet Gynecol.* 2009;113(1):89-95.
- 2. Anderson MR, Klink K, Cohrssen A. Evaluation of vaginal complaints. *JAMA*. 2004;291(11):1368-1379.
- Farage MA, Miller KW, Ledger WJ. Determining the cause of vulvovaginal symptoms. Obstet Gynecol Surv. 2008;63(7):445-464.
- Greendale GA, Judd HL. The menopause: health implications and clinical management. J Am Geriatr Soc. 1993;41(4):426-436.
- Bradshaw CS, Morton AN, Garland SM, Morris MB, Moss LM, Fairley CK. Higher-risk behavioral practices associated with bacterial vaginosis compared with vaginal candidiasis. *Obstet Gynecol.* 2005;106(1):105-114.
- Ness RB, Hillier SL, Richter HE, et al. Douching in relation to bacterial vaginosis, lactobacilli, and facultative bacteria in the vagina. *Obstet Gynecol.* 2002;100(4):765.
- Sanchez S, Garcia PJ, Thomas KK, Catlin M, Holmes KK. Intravaginal metronidazole gel versus metronidazole plus nystatin ovules for bacterial vaginosis: a randomized controlled trial. *Am J Obstet Gynecol.* 2004;191(6):1898-1906.
- Evans AL, Scally AJ, Wellard SJ, Wilson JD. Prevalence of bacterial vaginosis in lesbians and heterosexual women in a community setting. Sex Transm Infect. 2007;83(6):470-475.
- Schreiber CA, Meyn LA, Creinin MD, Barnhart KT, Hillier SL. Effects of long-term use of nonoxynol-9 on vaginal flora. *Obstet Gynecol.* 2006; 107(1):136-143.

- Cotch MF, Hillier SL, Gibbs RS, Eschenbach DA; Vaginal Infections and Prematurity Study Group. Epidemiology and outcomes associated with moderate to heavy *Candida* colonization during pregnancy. *Am J Obstet Gynecol.* 1998;178(2):374-380.
- Wilton L, Kollarova M, Heeley E, Shakir S. Relative risk of vaginal candidiasis after use of antibiotics compared with antidepressants in women: postmarketing surveillance data in England. *Drug Saf.* 2003;26(8): 589-597.
- Gutman RE, Peipert JF, Weitzen S, Blume J. Evaluation of clinical methods for diagnosing bacterial vaginosis. *Obstet Gynecol.* 2005;105(3): 551-556.
- 13. Centers for Disease Control and Prevention. Sexually transmitted diseases treatment guidelines 2010. Diseases characterized by vaginal discharge. http://www.cdc.gov/std/treatment/2010/toc.htm. Accessed February 28, 2011.
- 14. Sobel JD. Vulvovaginal candidosis. Lancet. 2007;369(9577):1961-1971.
- Chatwani AJ, Mehta R, Hassan S, Rahimi S, Jeronis S, Dandolu V. Rapid testing for vaginal yeast detection: a prospective study. *Am J Obstet Gynecol.* 2007;196(4):309.e1-4.
- West B, Morison L, Schim van der Loeff M, et al. Evaluation of a new rapid diagnostic kit (FemExam) for bacterial vaginosis in patients with vaginal discharge syndrome in The Gambia. *Sex Transm Dis.* 2003;30(6): 483-489.
- 17. Hillier SL. Diagnostic microbiology of bacterial vaginosis. *Am J Obstet Gynecol.* 1993;169(2 pt 2):455-459.
- Brown HL, Fuller DA, Davis TE, Schwebke JR, Hillier SL. Evaluation of the Affirm Ambient Temperature Transport System for the detection and identification of *Trichomonas vaginalis*, *Gardnerella vaginalis*, and *Candida* species from vaginal fluid specimens. J Clin Microbiol. 2001;39(9):3197-3199.
- Huppert JS, Batteiger BE, Braslins P, et al. Use of an immunochromatographic assay for rapid detection of *Trichomonas vaginalis* in vaginal specimens. J Clin Microbiol. 2005;43(2):684-687.
- Radonjic IV, Dzamic AM, Mitrovic SM, Arsic Arsenijevic VS, Popadic DM, Kranjcic Zec IF. Diagnosis of *Trichomonas vaginalis* infection: The sensitivities and specificities of microscopy, culture and PCR assay. *Eur J Obstet Gynecol Reprod Biol.* 2006;126(1):116-120.
- Lara-Torre E, Pinkerton JS. Accuracy of detection of *Trichomonas vaginalis* organisms on a liquid-based Papanicolaou smear. *Am J Obstet Gynecol.* 2003;188(2):354-356.
- 22. Weissenbacher T, Witkin SS, Ledger WJ, et al. Relationship between clinical diagnosis of recurrent vulvovaginal candidiasis and detection of *Candida* species by culture and polymerase chain reaction. *Arch Gynecol Obstet.* 2009;279(2):125-129.
- Kissinger PJ, Dumestre J, Clark RA, et al. Vaginal swabs versus lavage for detection of *Trichomonas vaginalis* and bacterial vaginosis among HIV-positive women. *Sex Transm Dis.* 2005;32(4):227-230.
- 24. Charonis G, Larsson PG. Use of pH/whiff test or QuickVue Advanced pH and Amines test for the diagnosis of bacterial vaginosis and prevention of postabortion pelvic inflammatory disease. Acta Obstet Gynecol Scand. 2006;85(7):837-843.
- Carr PL, Rothberg MB, Friedman RH, Felsenstein D, Pliskin JS. "Shotgun" versus sequential testing. Cost-effectiveness of diagnostic strategies for vaginitis. J Gen Intern Med. 2005;20(9):793-799.
- Allsworth JE, Peipert JF. Prevalance of bacterial vagniosis: 2001-2004 National Health and Nutrition Examination Survey data. *Obstet Gyne*col. 2007;109(1):114-120.
- Livengood CH III, Thomason JL, Hill GB. Bacterial vaginosis: diagnostic and pathogenetic findings during topical clindamycin therapy. Am J Obstet Gynecol. 1990;163(2):515-520.
- Ness RB, Hillier SL, Kip KE, et al. Bacterial vaginosis and risk of pelvic inflammatory disease. Obstet Gynecol. 2004;104(4):761-769.
- 29. Leitich H, Bodner-Adler B, Brunbauer M, Kaider A, Egarter C, Husslein P.

Bacterial vaginosis as a risk factor for preterm delivery: a meta-analysis. *Am J Obstet Gynecol.* 2003;189(1):139-147.

- Martin HL, Richardson BA, Nyange PM, et al. Vaginal lactobacilli, microbial flora, and risk of human immunodeficiency virus type 1 and sexually transmitted disease acquisition. J Infect Dis. 1999;180(6):1863-1868.
- Amsel R, Totten PA, Spiegel CA, Chen KC, Eschenbach D, Holmes KK. Nonspecific vaginitis. Diagnostic criteria and microbial and epidemiologic associations. Am J Med. 1983;74(1):14-22.
- 32. Drugstore.com. http://www.drugstore.com. Accessed September 12, 2010.
- Oduyebo OO, Anorlu RI, Ogunsola FT. The effects of antimicrobial therapy on bacterial vaginosis in non-pregnant women. *Cochrane Database Syst Rev.* 2009;(3):CD006055.
- Senok AC, Verstraelen H, Temmerman M, Botta GA. Probiotics for the treatment of bacterial vaginosis. *Cochrane Database Syst Rev.* 2009; (4):CD006289.
- McDonald HM, Brocklehurst P, Gordon A. Antibiotics for treating bacterial vaginosis in pregnancy. *Cochrane Database Syst Rev.* 2007;(1): CD000262.
- 36. U.S. Preventive Services Task Force. Screening for bacterial vaginosis in pregnancy to prevent preterm delivery: U.S. Preventive Services Task Force recommendation statement. *Ann Intern Med.* 2008;148(3): 214-219.
- Boris J, Påhlson C, Larsson PG. Six years observation after successful treatment of bacterial vaginosis. *Infect Dis Obstet Gynecol*. 1997;5(4):297-302.
- Ralph SG, Rutherford AJ, Wilson JD. Influence of bacterial vaginosis on conception and miscarriage in the first trimester: cohort study. *BMJ*. 1999;319(7204):220-223.
- Sobel JD, Ferris D, Schwebke J, et al. Suppressive antibacterial therapy with 0.75% metronidazole vaginal gel to prevent recurrent bacterial vaginosis. *Am J Obstet Gynecol.* 2006;194(5):1283-1289.
- Wiese W, Patel SR, Patel SC, Ohl CA, Estrada CA. A meta-analysis of the Papanicolaou smear and wet mount for the diagnosis of vaginal trichomoniasis. *Am J Med.* 2000;108(4):301-308.
- Blake DR, Duggan A, Joffe A. Use of spun urine to enhance detection of Trichomonas vaginalis in adolescent women. Arch Pediatr Adolesc Med. 1999;153(12):1222-1225.
- Huppert JS, Mortensen JE, Reed JL, et al. Rapid antigen testing compares favorably with transcription-mediated amplification assay for the detection of *Trichomonas vaginalis* in young women. *Clin Infect Dis.* 2007;45(2):194-198.
- Witkin SS, Inglis SR, Polaneczky M. Detection of Chlamydia trachomatis and Trichomonas vaginalis by polymerase chain reaction in introital specimens from pregnant women. Am J Obstet Gynecol. 1996; 175(1):165-167.
- 44. Forna F, Gülmezoglu AM. Interventions for treating trichomoniasis in women. *Cochrane Database Syst Rev.* 2003;(2):CD000218.
- 45. Gülmezoglu AM. Interventions for trichomoniasis in pregnancy. *Cochrane Database Syst Rev.* 2002;(3):CD000220.
- Corsello S, Spinillo A, Osnengo G, et al. An epidemiological survey of vulvovaginal candidiasis in Italy. *Eur J Obstet Gynecol Reprod Biol.* 2003;110(1):66-72.
- 47. Nurbhai M, Grimshaw J, Watson M, Bond C, Mollison J, Ludbrook A. Oral versus intra-vaginal imidazole and triazole anti-fungal treatment of uncomplicated vulvovaginal candidiasis (thrush). *Cochrane Database Syst Rev.* 2007;(4):CD002845.
- Pappas PG, Kauffman CA, Andes D, et al.; Infectious Diseases Society of America. Clinical practice guidelines for the management of candidiasis: 2009 update by the Infectious Diseases Society of America. *Clin Infect Dis.* 2009;48(5):503-535.
- Suckling JA, Lethaby A, Kennedy R. Local oestrogen for vaginal atrophy in postmenopausal women. *Cochrane Database Syst Rev.* 2006; (4):CD001500.