

Vaginitis: Diagnosis and Treatment

Heather L. Paladine, MD, MEd, and Urmi A. Desai, MD, MS

Columbia University Irving Medical Center, New York, New York

Vaginitis is defined as any condition with symptoms of abnormal vaginal discharge, odor, irritation, itching, or burning. The most common causes of vaginitis are bacterial vaginosis, vulvovaginal candidiasis, and trichomoniasis. Bacterial vaginosis is implicated in 40% to 50% of cases when a cause is identified, with vulvovaginal candidiasis accounting for 20% to 25% and trichomoniasis for 15% to 20% of cases. Noninfectious causes, including atrophic, irritant, allergic, and inflammatory vaginitis, are less common and account for 5% to 10% of vaginitis cases. Diagnosis is made using a combination of symptoms, physical examination findings, and office-based or laboratory testing. Bacterial vaginosis is traditionally diagnosed with Amsel criteria, although Gram stain is the diagnostic standard. Newer laboratory tests that detect *Gardnerella vaginalis* DNA or vaginal fluid sialidase activity have similar sensitivity and specificity to Gram stain. Bacterial vaginosis is treated with oral metronidazole, intravaginal metronidazole, or intravaginal clindamycin. The diagnosis of vulvovaginal candidiasis is made using a combination of clinical signs and symptoms with potassium hydroxide microscopy; DNA probe testing is also available. Culture can be helpful for the diagnosis of complicated vulvovaginal candidiasis by identifying nonalbicans strains of *Candida*. Treatment of vulvovaginal candidiasis involves oral fluconazole or topical azoles, although only topical azoles are recommended during pregnancy. The Centers for Disease Control and Prevention recommends nucleic acid amplification testing for the diagnosis of trichomoniasis in symptomatic or high-risk women. Trichomoniasis is treated with oral metronidazole or tinidazole, and patients' sex partners should be treated as well. Treatment of noninfectious vaginitis should be directed at the underlying cause. Atrophic vaginitis is treated with hormonal and nonhormonal therapies. Inflammatory vaginitis may improve with topical clindamycin as well as steroid application. (*Am Fam Physician*. 2018;97(5):321-329. Copyright © 2018 American Academy of Family Physicians.)

Vaginitis is characterized by vaginal symptoms, including discharge, odor, itching, irritation, or burning.¹ Most women have at least one episode of vaginitis during their lives,² making it the most common gynecologic diagnosis in primary care. Studies have shown a negative effect on quality of life in women with vaginitis, with some women expressing anxiety, shame, and concerns about hygiene, particularly in those with recurrent symptoms.³⁻⁸

The most common causes of vaginitis are bacterial vaginosis, vulvovaginal candidiasis, and trichomoniasis. Bacterial vaginosis is the cause in 40% to 50% of cases in

which a cause is identified, with vulvovaginal candidiasis accounting for 20% to 25% and trichomoniasis for 15% to 20% of cases. Noninfectious causes, including atrophic, irritant, allergic, and inflammatory vaginitis, are less common and account for 5% to 10% of vaginitis cases.⁹

WHAT IS NEW ON THIS TOPIC

Vaginitis

A 2013 meta-analysis showed that oral or topical antibiotic treatment of bacterial vaginosis in pregnancy does not prevent preterm birth, even in women with a history of preterm labor in previous pregnancies.

Newer laboratory tests such as DNA and antigen testing for bacterial vaginosis and vulvovaginal candidiasis, or vaginal fluid sialidase testing for bacterial vaginosis, may have similar or better sensitivity and specificity compared with traditional office-based testing. However, comparative cost-effectiveness has not been studied.

CME This clinical content conforms to AAFP criteria for continuing medical education (CME). See CME Quiz on page 307.

Author disclosure: No relevant financial affiliations.

Patient information: A handout on this topic, written by the authors of this article, is available at <http://www.aafp.org/afp/2018/0301/p321-s1.html>.

TABLE 1

Signs and Symptoms of Vaginitis

Diagnosis	Etiology	Symptoms	Signs	Other risks
Bacterial vaginosis	Anaerobic bacteria (<i>Prevotella</i> , <i>Mobiluncus</i> , <i>Gardnerella vaginalis</i> , <i>Ureaplasma</i> , <i>Mycoplasma</i>)	Fishy odor; thin, homogeneous discharge that may worsen after intercourse; pelvic discomfort may be present	No inflammation	Increased risk of HIV, gonorrhea, chlamydia, and herpes infections
Vulvovaginal candidiasis	<i>Candida albicans</i> , can have other <i>Candida</i> species	White, thick, cheesy, or curdy discharge; vulvar itching or burning; no odor	Vulvar erythema and edema	—
Trichomoniasis	<i>Trichomonas vaginalis</i>	Green or yellow, frothy discharge; foul odor; vaginal pain or soreness	Inflammation; strawberry cervix	Increased risk of HIV infection Increased risk of preterm labor Should be screened for other sexually transmitted infections
Atrophic vaginitis	Estrogen deficiency	Thin, clear discharge; vaginal dryness; dyspareunia; itching	Inflammation; thin, friable vaginal mucosa	—
Irritant/allergic vaginitis	Contact irritation or allergic reaction	Burning, soreness	Vulvar erythema	—
Inflammatory vaginitis	Possibly autoimmune	Purulent vaginal discharge, burning, dyspareunia	Vaginal atrophy and inflammation	Associated with low estrogen levels

HIV = human immunodeficiency virus.

Information from references 10, 14, and 15.

General Diagnostic Considerations

HISTORY AND PHYSICAL EXAMINATION

The history alone is unreliable for the diagnosis of different causes of vaginitis.¹⁰ Physical examination findings and office-based or laboratory test results should be used with the history to determine the diagnosis.¹¹⁻¹³ Characteristic clinical signs and symptoms for different causes of vaginitis are listed in *Table 1*.^{10,14,15} Risk factors for vaginitis are listed in *Table 2*.^{9,14,16}

Signs and symptoms that increase the likelihood of vulvovaginal candidiasis vs. bacterial vaginosis are a cheesy, curdy, or flocculent discharge; itching; vulvar or vaginal inflammation or redness; and lack of odor.¹⁰ A fishy odor makes candidiasis less likely.¹⁰ Normal physiologic vaginal discharge is clear to white, not malodorous, not accompanied by discomfort or pruritus, and the quantity varies during a woman's menstrual cycle.

OFFICE AND LABORATORY TESTING

Office-based tests include microscopy, measurement of vaginal pH, and whiff test. A speculum is not required for collecting

TABLE 2

Risk Factors Contributing to Vaginitis

Type of vaginitis	Risk factors
Bacterial vaginosis	Low socioeconomic status, vaginal douching, smoking, new or multiple sex partners, unprotected intercourse, women who have sex with women
Vulvovaginal candidiasis	Recent antibiotic use, pregnancy, uncontrolled diabetes mellitus, AIDS, corticosteroid use, other immunosuppression
Trichomoniasis	Low socioeconomic status, multiple sex partners, other sexually transmitted infections, unprotected intercourse, drug use, smoking
Atrophic or inflammatory vaginitis	Menopause, lactation, oophorectomy, radiation therapy, chemotherapy, immunologic disorders, premature ovarian failure, endocrine disorders, antiestrogen medications
Irritant vaginitis	Soaps, tampons, contraceptive devices such as condoms or diaphragms, sex toys, pessaries, topical products, douching, fastidious cleansing, medications, clothing
Allergic vaginitis	Sperm, douching, latex condoms or diaphragms, tampons, topical products, medications, clothing, atopic history

Adapted with permission from Hainer BL, Gibson MV. Vaginitis. *Am Fam Physician*. 2011;83(7):809, with additional information from references 9 and 16.

TABLE 3

Amsel Diagnostic Criteria for Bacterial Vaginosis*

Thin, homogenous discharge

Positive whiff test (i.e., the amine odor produced by mixing 10% potassium hydroxide solution with vaginal discharge sample)

Clue cells present on microscopy† (Figure 1)

Vaginal pH > 4.5

*—Three out of four criteria must be met; establishes accurate diagnosis of bacterial vaginosis in 90% of affected women.

†—Highly significant criterion.

Information from references 14 and 23.

vaginal fluid samples for these tests.¹⁷ Several studies have demonstrated a strong correlation between samples from patient self-collected swabs and those collected by clinicians for the diagnosis of bacterial vaginosis, with sensitivities of 70% to 100% and specificities of 97% to 100%.^{18–22} It is reasonable to conclude that samples for office-based microscopy and laboratory testing for other causes of vaginitis can be collected by patients as well. Patients should be instructed to insert the swab at least one inch into the vagina.

BACTERIAL VAGINOSIS

Although Gram stain is considered the diagnostic standard, bacterial vaginosis is traditionally diagnosed using the Amsel criteria (Table 3).^{14,23} Criteria include thin, homogenous discharge; a positive whiff test; the presence of clue cells on microscopy (Figure 1¹⁴); and a vaginal pH greater than 4.5. Three out of four criteria are required to make the diagnosis, with sensitivity ranging from 70% to 97% and specificity from 90% to 94%, compared with Gram stain.^{24,25} Newer methods of laboratory testing with DNA probe for *Gardnerella vaginalis* or detection of vaginal fluid sialidase activity have sensitivities of 92% to 100% and specificities of 92% to 98% compared with Gram stain.^{26–28}

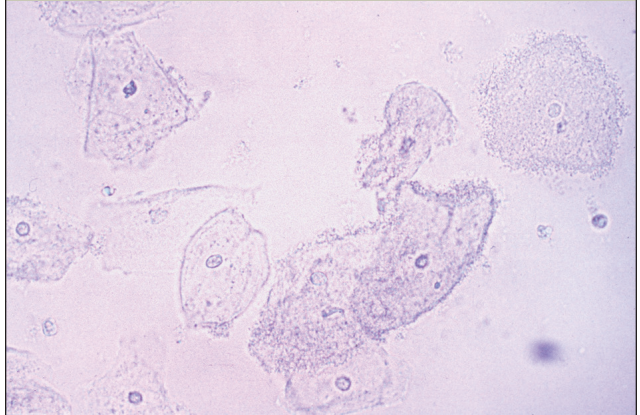
Vaginal culture and Papanicolaou (Pap) testing are not useful for diagnosing bacterial vaginosis because it is a polymicrobial infection.⁹ Group B streptococcus may be found on culture and has been associated with vaginitis symptoms, although there is no expert consensus on treatment in nonpregnant women.²⁹

Recurrence of bacterial vaginosis is common. Women should be advised to return for treatment if symptoms recur. Routine testing in asymptomatic women and retesting (test of cure) are not recommended because these bacteria can be part of normal flora.

VULVOVAGINAL CANDIDIASIS

Vulvovaginal candidiasis can be diagnosed by visualization of yeast hyphae on potassium hydroxide preparation

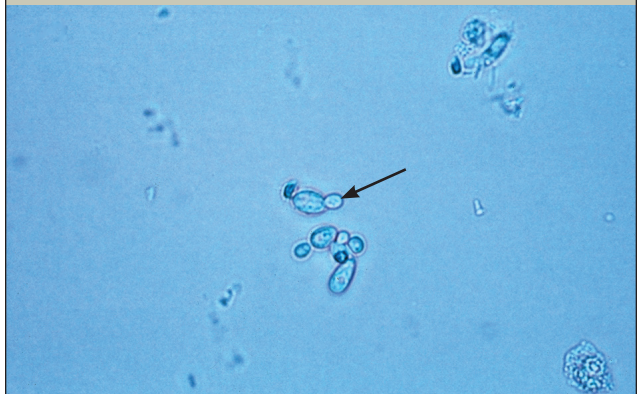
FIGURE 1



Clue cells (400 X). Vaginal epithelial cells with borders obscured by adherent coccobacilli visible on saline wet-mount preparation.

Reprinted with permission from Hainer BL, Gibson MV. Vaginitis. *Am Fam Physician.* 2011;83(7):809.

FIGURE 2



Candida species (400 X). Budding yeast visible (arrow).

Reprinted with permission from Hainer BL, Gibson MV. Vaginitis. *Am Fam Physician.* 2011;83(7):814.

(Figure 2¹⁴) in a woman with typical symptoms. It can also be diagnosed using antigen or DNA probe testing, with sensitivities of 77% to 97% and specificities of 77% to 99%, compared with culture as the diagnostic standard.^{30–32} Women with vulvovaginal candidiasis have a normal acidic vaginal pH.

Complicated vulvovaginal candidiasis is defined as recurrent (four or more episodes in one year) or severe infections, or infections that occur in a patient who is immunocompromised, such as someone with AIDS or poorly controlled diabetes mellitus. Culture is particularly important for the diagnosis and treatment of complicated vulvovaginal candidiasis, because patients are more likely to have an infection with nonalbicans

TABLE 4

Treatment Regimens for the Most Common Causes of Vaginitis

Initial regimens	Alternative regimens	Pregnancy	Recurrence	Treatment of sex partners
Bacterial vaginosis				
Metronidazole (Flagyl), 500 mg orally twice daily for seven days*	Tinidazole (Tindamax), 2 g orally once daily for two days	Metronidazole, 500 mg orally twice daily for seven days	First recurrence: Retrial of same regimen or Trial of alternative initial regimen	Routine treatment of sex partners is not recommended
or	or			
Metronidazole 0.75% gel (Metrogel), one full applicator (5 g) intravaginally daily for five days	Tinidazole, 1 g orally once daily for five days		Multiple recurrences: Metronidazole 0.75% gel, intravaginally twice weekly for four to six months	
or	or			
Clindamycin 2% cream, one full applicator (5 g) intravaginally at bedtime for seven days†	Clindamycin, 300 mg orally twice daily for seven days			
	or			
	Clindamycin (Cleocin Ovules), 100 mg intravaginally at bedtime for three days			
Vulvovaginal candidiasis				
Topical azole therapy‡ (Table 5)	—	Topical azole therapy applied intravaginally for seven days	To achieve mycologic cure§: Topical azole therapy for seven to 14 days or Fluconazole, 150 mg orally every third day for three doses	Routine treatment of sex partners is not recommended unless the partner is symptomatic
or			For maintenance: Oral fluconazole (100 mg, 150 mg, or 200 mg) weekly for six months; consider topical treatment if oral is not feasible	
Fluconazole (Diflucan), 150 mg orally, single dose				
Trichomoniasis				
Metronidazole, 2 g orally, single or divided dose on the same day	Metronidazole, 500 mg orally twice daily for seven days	Metronidazole, 2 g orally, single dose in any stage of pregnancy	Differentiate persistent or recurrent infection from reinfection If metronidazole, 2-g single dose fails: Trial of metronidazole, 500 mg twice daily for seven days	Concurrent treatment of sex partners is recommended
or			If metronidazole, 500 mg twice daily for seven days fails: Trial of metronidazole, 2 g daily for seven days	Advise refraining from intercourse until partners are treated and symptom-free
Tinidazole, 2 g orally, single dose			If above regimens fail: Consider susceptibility testing (contact Centers for Disease Control and Prevention)	

*—Because of disulfiram-like reaction, alcohol should be avoided for at least 24 hours after completing oral regimen.

†—Clindamycin cream is oil-based and can weaken latex condoms and diaphragms for at least five days after use.

‡—Topical azole creams and suppositories may be oil-based and can weaken latex condoms and diaphragms.

§—For *Candida albicans* infection. Consider culture to exclude nonalbicans infection. If nonalbicans infection is present, consider first-line therapy with seven to 14 days of a nonfluconazole azole agent. If infection recurs, prescribe 600 mg of boric acid in a gelatin capsule intravaginally once daily for two weeks. Boric acid may also be used with initial induction therapy followed by monthly maintenance therapy for recurrent albicans infection per the Society of Obstetricians and Gynaecologists of Canada recommendations.¹⁶

||—Follow-up with retesting as early as two weeks but within three months is recommended because rates of reinfection are high.

Information from references 9 and 16.

strains of *Candida*, which may require different treatment (Table 4).^{9,16} Candidal species and anaerobes can be normal flora in asymptomatic women, so retesting (test of cure) is not recommended in the absence of symptoms.

TRICHOMONIASIS

Trichomoniasis is a sexually transmitted infection that should be considered in women at risk who present with vaginitis symptoms (Table 2^{9,14,16}). It can be diagnosed when motile, flagellated protozoa are observed on saline microscopy (Figure 3¹⁴). However, the Centers for Disease Control and Prevention (CDC) recommends nucleic acid amplification testing for the diagnosis of trichomoniasis in symptomatic or high-risk women. These tests have a higher sensitivity than saline microscopy (95% to 100% vs. 51% to 65%)⁹ and can be performed on endocervical, vaginal, or urine specimens, or on liquid-based Pap test samples. Vaginal culture also has a high sensitivity for identifying *Trichomonas*, but it has largely been replaced by nucleic acid amplification testing because of the longer time (up to one week) needed for results.⁹

Because trichomoniasis is sexually transmitted and has a high rate of recurrence, the CDC recommends testing for reinfection three months after treatment.⁹

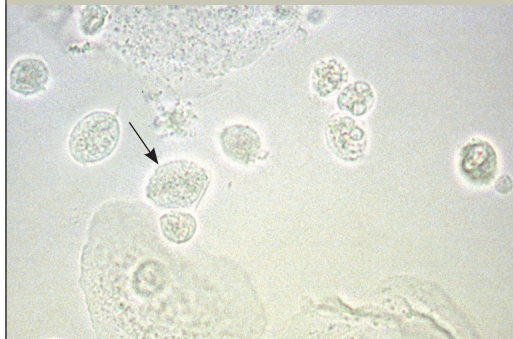
NEWER LABORATORY TESTS

Some data show that newer laboratory tests such as DNA and antigen testing for bacterial vaginosis and vulvovaginal candidiasis, or vaginal fluid sialidase testing for bacterial vaginosis, may have similar or better sensitivity and specificity compared with office-based testing.³³ However, more comparisons with diagnostic standard testing (i.e., Gram stain for bacterial vaginosis and culture for vulvovaginal candidiasis) are needed. The costs of these tests range from approximately \$17 for vaginal fluid sialidase testing to \$27 to \$49 for DNA testing, compared with \$5 to \$10 for office-based microscopy. No studies were found on the cost-effectiveness of these newer tests compared with office-based testing.

PAPANICOLAOU TESTS

When clue cells or hyphae are present on a Pap test, treatment depends on symptoms. Both can be normal findings

FIGURE 3



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

Reprinted with permission from Hainer BL, Gibson MV. Vaginitis. *Am Fam Physician*. 2011;83(7):813.

in asymptomatic women.⁹ Also, physicians should treat trichomoniasis when found on a Pap test, but a normal result does not rule out infection.³⁴ When seeking a diagnosis for vaginal symptoms at the same time as performing a Pap test, physicians can order testing on the liquid-based Pap fluid. These tests have similar sensitivity and specificity to vaginal samples.

OTHER CAUSES OF VAGINITIS

There is no cause of vaginitis identified in up to 30% of women. These women may have a range of conditions, including irritant or allergic vaginitis, atrophic vaginitis, or physiologic discharge.³⁵ Inflammatory vaginitis is an uncommon condition

characterized by purulent discharge, burning, and dyspareunia, and should be considered in patients with these symptoms if no infectious cause is found. Inflammatory vaginitis is associated with low estrogen levels, such as in menopausal or perimenopausal women.³⁶

Treatment

First-line and alternative treatment regimens for vaginitis are presented in Table 4 with suggestions for recurrent infection, treatment during pregnancy, and treatment of sex partners.^{9,16} Additional considerations for the treatment of bacterial vaginosis, vulvovaginal candidiasis, and trichomoniasis are discussed.

BACTERIAL VAGINOSIS

Treatment of bacterial vaginosis is recommended for resolving symptoms, as well as reducing the risk of *Chlamydia trachomatis*, *Neisseria gonorrhoeae*, *Trichomonas vaginalis*, human immunodeficiency virus (HIV), and herpes simplex virus type 2 infections.³⁷ Shifts in vaginal flora have been associated with increased risk of these infections, leading researchers to conclude that treatment of bacterial vaginosis may decrease susceptibility to these infections.^{38,39}

First-line therapy includes seven-day courses of oral metronidazole (Flagyl), intravaginal metronidazole (Metrogel), or intravaginal clindamycin.⁹ No significant difference in effectiveness has been demonstrated among these regimens.⁴⁰ Patient preference should be considered

TABLE 5

Recommended Topical Treatment Regimens for Vulvovaginal Candidiasis

Over-the-counter intravaginal agents

Clotrimazole 1% cream, 5 g intravaginally daily for seven to 14 days
Clotrimazole 2% cream, 5 g intravaginally daily for three days
Miconazole 2% cream, 5 g intravaginally daily for seven days
Miconazole 4% cream, 5 g intravaginally daily for three days
Miconazole 100-mg vaginal suppository, one suppository daily for seven days
Miconazole 200-mg vaginal suppository, one suppository daily for three days
Miconazole 1,200-mg vaginal suppository, one suppository for one day
Tioconazole 6.5% ointment, 5 g intravaginally in a single application

Prescription intravaginal agents

Butoconazole 2% cream (Gynazole-1, single-dose bioadhesive product), 5 g intravaginally in a single application
Terconazole 0.4% cream, 5 g intravaginally daily for seven days
Terconazole 0.8% cream, 5 g intravaginally daily for three days
Terconazole 80-mg vaginal suppository, one suppository daily for three days

Adapted from Workowski KA, Bolan GA; Centers for Disease Control and Prevention. Sexually transmitted diseases treatment guidelines, 2015 [published correction appears in MMWR Recomm Rep. 2015;64(33):924]. MMWR Recomm Rep. 2015;64(RR-03):76, with additional information from reference 16.

when choosing an agent. Physicians should explain potential adverse effects with each regimen, including a possible disulfiram-like reaction with alcohol consumption or gastrointestinal symptoms in persons taking oral metronidazole, or possible weakening of latex condoms with the use of topical therapies containing oil-based preparations.⁹

The U.S. Food and Drug Administration recently approved a single-dose oral therapy for bacterial vaginosis, secnidazole (SoloSec), which will be available in 2018.⁴¹ A randomized controlled trial demonstrated similar effectiveness and outcomes compared with metronidazole for the treatment of bacterial vaginosis.⁴² The dosing involves one-time oral administration of a 2-g packet of granules mixed into applesauce, yogurt, or pudding. A primary adverse effect of this regimen is vulvovaginal candidiasis. Data are insufficient on the safety of secnidazole use in pregnancy, and use is not recommended with breastfeeding.⁴³

Bacterial Vaginosis in Pregnancy. In the past, treatment for bacterial vaginosis during pregnancy was recommended to prevent preterm births. Further review of the evidence has demonstrated that antibiotic treatment does not prevent preterm birth for women with symptomatic or asymptomatic bacterial vaginosis.^{44,45} Although a previous meta-analysis demonstrated a possible reduction in preterm labor with treatment of bacterial vaginosis, particularly in early pregnancy (before 20 weeks' gestation),⁴⁶ a more recent meta-analysis of 21 studies found that antibiotic treatment—regardless of route of administration (oral or topical), the timing in pregnancy, or history of preterm labor in previous pregnancies—does not prevent preterm birth for women with symptomatic or asymptomatic bacterial vaginosis.⁴⁵ Two studies cited in this meta-analysis that included the presence of abnormal vaginal flora as well as bacterial vaginosis showed a possible reduction in preterm labor before 37 weeks' gestation with treatment; therefore, further investigation may provide more information about the role of abnormal bacterial flora and its treatment in pregnancy.⁴⁵ Regardless, treatment of bacterial vaginosis is generally recommended for symptomatic relief, and adverse effects of metronidazole in pregnancy have not been demonstrated.⁴⁵

VULVOVAGINAL CANDIDIASIS

Treatment of candidal infection is aimed at reducing symptoms. There are several topical azole preparations and regimens available, as well as oral fluconazole (Diflucan)

in a single 150-mg dose. All topical treatments listed in *Table 5*,^{9,16} as well as oral fluconazole, are recommended by the CDC as first-line therapy for vulvovaginal candidiasis.⁹ The choice of treatment should be made in conjunction with patient preference and previous experience with these agents. In nonpregnant patients, there is no definitive advantage of one treatment over another in terms of clinical or mycologic cure, with all treatment options having about an 80% cure rate.^{47,48} Limited data demonstrate that patients may prefer oral regimens.⁴⁹⁻⁵¹

There are several considerations when choosing between topical and oral treatment.^{52,53} Topical preparations may provide more immediate relief because of the soothing nature of topical treatment or, conversely, may cause local hypersensitivity reactions resulting in itching or burning. Oral fluconazole offers the advantage of one-dose convenience without messy creams or suppositories. Oral medications may cause systemic adverse effects, particularly gastrointestinal effects and toxicity in addition to potential medication interactions. An additional factor to consider is that topical azole creams and suppositories may be oil-based and can weaken latex condoms.⁹

Topical antifungals are available over the counter, and many patients self-diagnose and treat presumed vulvovaginal candidiasis with these products. However, studies have shown that, regardless of whether they have a history of vulvovaginal candidiasis, women are not able to accurately

SORT: KEY RECOMMENDATIONS FOR PRACTICE

Clinical recommendation	Evidence rating	References
Symptoms alone cannot differentiate between the causes of vaginitis. Office-based or laboratory testing should be used with the history and physical examination findings to make the diagnosis.	C	10-12
Do not obtain culture for the diagnosis of bacterial vaginosis because it represents a polymicrobial infection.	C	9
Nucleic acid amplification testing is recommended for the diagnosis of trichomoniasis in symptomatic or high-risk women.	C	9
Treatment of bacterial vaginosis during pregnancy improves symptoms but does not reduce the risk of preterm birth.	A	44, 45
In nonpregnant women, oral and vaginal treatment options for uncomplicated vulvovaginal candidiasis have similar clinical cure rates.	B	47

A = consistent, good-quality patient-oriented evidence; **B** = inconsistent or limited-quality 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>.

Topical imidazoles (i.e., econazole, clotrimazole, miconazole, and ketoconazole) are more effective in eradication.⁵⁹ In a small study, topical terconazole was also shown to relieve symptoms and achieve mycologic cure for about 50% of patients.⁶⁰ For patients with nonalbicans infections who do not respond to these treatments, a regimen of 600-mg vaginal boric acid capsules daily for 14 days has been shown to be effective.⁶¹

Other nonpharmacologic regimens have been proposed for recurrent vulvovaginal candidiasis. A meta-analysis did not demonstrate clear evidence for probiotics in the treatment of candidal vaginitis; however, more studies are needed because of small study size and varied probiotic regimens.⁶²

self-diagnose the condition, often using these products for causes of vaginitis other than candidal infections.⁵⁴

Vulvovaginal Candidiasis in Pregnancy. Vulvovaginal candidiasis is common during pregnancy. The CDC recommends that only topical azole therapies, applied for seven days, be used in pregnant women with vulvovaginal candidiasis.⁹ Although more data are needed, oral therapy may be associated with increased risk of miscarriage or fetal malformations, particularly at high doses.^{55,56}

Complicated Vulvovaginal Candidiasis. Patients with complicated vulvovaginal candidiasis require more aggressive therapy. To guide treatment, it is helpful to consider whether a patient has recurrent infections and whether the etiology may be a nonalbicans species of *Candida*.

For patients with severe vulvovaginal candidiasis, a second dose of fluconazole given three days after the first dose has been shown to achieve significant improvement in short-term symptoms as well as prevent recurrence at 35 days. A second dose did not have significant effects for recurrent vulvovaginal candidiasis.⁵⁷

For recurrent vulvovaginal candidiasis caused by *Candida albicans*, initial intensive therapy with fluconazole for seven to 14 days (i.e., fluconazole, 150 mg, every three days for three doses) followed by weekly treatment with 150-mg fluconazole for six months has been shown to achieve symptomatic relief at one year for most patients.⁵⁸

If severe or recurrent vulvovaginal candidiasis does not respond to initial treatment, culture may guide therapy when nonalbicans species are identified. Infections with nonalbicans species are less responsive to fluconazole.

TRICHOMONIASIS

Treatment of trichomoniasis can decrease symptoms and reduce transmission to partners. In addition, in persons with HIV infection, treatment of trichomoniasis may also decrease HIV transmission rates to partners.⁶³

Although first-line therapy for trichomoniasis in pregnant and nonpregnant women is a single 2-g dose of metronidazole, patients with HIV infection treated with a seven-day course of metronidazole had lower rates of infection at test of cure and lower rates of reinfection at three months.^{9,64} Tinidazole (Tindamax) in a single 2-g dose is also a first-line treatment for trichomoniasis; however, it is more expensive. Metronidazole-resistant trichomoniasis, which would require tinidazole, is rare.⁶⁵

Trichomoniasis in Pregnancy. Trichomoniasis has been associated with adverse pregnancy outcomes, including low birth weight and preterm birth.⁶⁶ Symptomatic pregnant women, regardless of pregnancy stage, should be tested and considered for treatment.

Noninfectious Vaginitis

Treatment of noninfectious vaginitis should be directed at the underlying cause. Atrophic vaginitis is treated with hormonal and nonhormonal therapies. Among hormonal therapies, low-dose vaginal estrogen preparations are available in creams, tablets, and rings. Systemic estrogen therapies are also available for patients with vasomotor symptoms.⁶⁷ First-line nonhormonal treatment recommendations include vaginal lubricants and moisturizers; continued sexual activity should be encouraged.⁶⁸

More research is needed to better characterize the cause and treatment of inflammatory vaginitis. Some studies have demonstrated improvement in symptoms with application of topical clindamycin or steroids; however, the ideal duration of treatment and superiority of one agent over the other have not been established.^{35,36}

This article updates previous articles on this topic by Hainer and Gibson,¹⁴ Owen and Clenney,⁶⁹ and Egan and Lipsky.⁷⁰

Data Sources: We searched PubMed, the Cochrane database, and the National Guideline Clearinghouse for vaginitis topics, and retrieved relevant references from review articles and clinical guidelines. Search terms included vaginitis, bacterial vaginosis, *Candida*, and *Trichomonas*. We also included the literature review from Essential Evidence Plus. Search dates: October 2016 and April 2017.

The Authors

HEATHER L. PALADINE, MD, MEd, is an assistant professor of medicine in the Center for Family and Community Medicine and director of the Family Medicine Residency Program at Columbia University Irving Medical Center, New York, NY.

URMI A. DESAI, MD, MS, is an assistant professor of medicine in the Center for Family and Community Medicine at Columbia University Irving Medical Center.

Address correspondence to Heather L. Paladine, MD, MEd, Columbia University Irving Medical Center, 610 W. 158th St., New York, NY 10032 (e-mail: hlp222@gmail.com). Reprints are not available from the authors.

References

- ACOG Practice Bulletin. Clinical management guidelines for obstetrician-gynecologists, number 72, May 2006: vaginitis. *Obstet Gynecol*. 2006;107(5):1195-1206.
- Sobel JD. Vaginitis. *N Engl J Med*. 1997;337(26):1896-1903.
- Bilardi JE, Walker S, Temple-Smith M, et al. The burden of bacterial vaginosis: women's experience of the physical, emotional, sexual and social impact of living with recurrent bacterial vaginosis. *PLoS One*. 2013;8(9):e74378.
- Karasz A, Anderson M. The vaginitis monologues: women's experiences of vaginal complaints in a primary care setting. *Soc Sci Med*. 2003;56(5):1013-1021.
- Payne SC, Cromer PR, Stanek MK, Palmer AA. Evidence of African-American women's frustrations with chronic recurrent bacterial vaginosis. *J Am Acad Nurse Pract*. 2010;22(2):101-108.
- Irving G, Miller D, Robinson A, Reynolds S, Copas AJ. Psychological factors associated with recurrent vaginal candidiasis: a preliminary study. *Sex Transm Infect*. 1998;74(5):334-338.
- Zhu YX, Li T, Fan SR, Liu XP, Liang YH, Liu P. Health-related quality of life as measured with the Short-Form 36 (SF-36) questionnaire in patients with recurrent vulvovaginal candidiasis. *Health Qual Life Outcomes*. 2016;14:65.
- Ehrström S, Kornfeld D, Rylander E. Perceived stress in women with recurrent vulvovaginal candidiasis. *J Psychosom Obstet Gynaecol*. 2007;28(3):169-176.
- Workowski KA, Bolan GA; Centers for Disease Control and Prevention. Sexually transmitted diseases treatment guidelines, 2015 [published correction appears in *MMWR Recomm Rep*. 2015;64(33):924]. *MMWR Recomm Rep*. 2015;64(RR-03):1-137.
- Anderson MR, Klink K, Cochrssen A. Evaluation of vaginal complaints. *JAMA*. 2004;291(11):1368-1379.
- Schaaf VM, Perez-Stable EJ, Borchardt K. The limited value of symptoms and signs in the diagnosis of vaginal infections. *Arch Intern Med*. 1990;150(9):1929-1933.
- Bornstein J, Lakovsky Y, Lavi I, Bar-Am A, Abramovici H. The classic approach to diagnosis of vulvovaginitis: a critical analysis. *Infect Dis Obstet Gynecol*. 2001;9(2):105-111.
- Nwankwo TO, Aniebue UU, Umeh UA. Syndromic diagnosis in evaluation of women with symptoms of vaginitis. *Curr Infect Dis Rep*. 2017;19(1):3.
- Hainer BL, Gibson MV. Vaginitis. *Am Fam Physician*. 2011;83(7):807-815.
- Farage MA, Miller KW, Ledger WJ. Determining the cause of vulvovaginal symptoms. *Obstet Gynecol Surv*. 2008;63(7):445-464.
- van Schalkwyk J, Yudin MH; Infectious Disease Committee. Vulvovaginitis: screening for and management of trichomoniasis, vulvovaginal candidiasis, and bacterial vaginosis. *J Obstet Gynaecol Can*. 2015;37(3):266-274.
- Blake DR, Duggan A, Quinn T, Zenilman J, Joffe A. Evaluation of vaginal infections in adolescent women: can it be done without a speculum? *Pediatrics*. 1998;102(4 pt 1):939-944.
- Nelson DB, Bellamy S, Gray TS, Nachamkin I. Self-collected versus provider-collected vaginal swabs for the diagnosis of bacterial vaginosis: an assessment of validity and reliability. *J Clin Epidemiol*. 2003;56(9):862-866.
- Strauss RA, Eucker B, Savitz DA, Thorp JM Jr. Diagnosis of bacterial vaginosis from self-obtained vaginal swabs. *Infect Dis Obstet Gynecol*. 2005;13(1):31-35.
- Kashyap B, Singh R, Bhalla P, Arora R, Aggarwal A. Reliability of self-collected versus provider-collected vaginal swabs for the diagnosis of bacterial vaginosis. *Int J STD AIDS*. 2008;19(8):510-513.
- Morgan DJ, Aboud CJ, McCaffrey IM, Bhide SA, Lamont RF, Taylor-Robinson D. Comparison of Gram-stained smears prepared from blind vaginal swabs with those obtained at speculum examination for the assessment of vaginal flora. *Br J Obstet Gynaecol*. 1996;103(11):1105-1108.
- Tanksale VS, Sahasrabhojane M, Patel V, Nevrekar P, Menezes S, Mabey D. The reliability of a structured examination protocol and self administered vaginal swabs: a pilot study of gynaecological outpatients in Goa, India. *Sex Transm Infect*. 2003;79(3):251-253.
- 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.
- Simoes JA, Discacciati MG, Brolazo EM, Portugal PM, Dini DV, Dantas MC. Clinical diagnosis of bacterial vaginosis. *Int J Gynaecol Obstet*. 2006;94(1):28-32.
- Schwebke JR, Hillier SL, Sobel JD, McGregor JA, Sweet RL. Validity of the vaginal gram stain for the diagnosis of bacterial vaginosis. *Obstet Gynecol*. 1996;88(4 pt 1):573-576.
- Menard JP, Mazouni C, Fenollar F, Raoult D, Boubli L, Bretelle F. Diagnostic accuracy of quantitative real-time PCR assay versus clinical and Gram stain identification of bacterial vaginosis. *Eur J Clin Microbiol Infect Dis*. 2010;29(12):1547-1552.
- Myziuk L, Romanowski B, Johnson SC. BVBlue test for diagnosis of bacterial vaginosis. *J Clin Microbiol*. 2003;41(5):1925-1928.
- Kampan NC, Suffian SS, Ithnin NS, Muhammad M, Zakaria SZ, Jamil MA. Evaluation of BV(®) Blue Test Kit for the diagnosis of bacterial vaginosis. *Sex Reprod Healthc*. 2011;2(1):1-5.
- Leclair CM, Hart AE, Goetsch MF, Carpentier H, Jensen JT. Group B streptococcus: prevalence in a non-obstetric population. *J Low Genit Tract Dis*. 2010;14(3):162-166.

VAGINITIS

30. 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.e1-309.e4.
31. Marot-Leblond A, Nail-Billaud S, Pilon F, Beucher B, Poulain D, Robert R. Efficient diagnosis of vulvovaginal candidiasis by use of a new rapid immunochromatography test. *J Clin Microbiol.* 2009;47(12):3821-3825.
32. Dan M, Leshem Y, Yeshaya A. Performance of a rapid yeast test in detecting *Candida* spp. in the vagina. *Diagn Microbiol Infect Dis.* 2010;67(1):52-55.
33. 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.
34. 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.
35. Nyirjesy P. Management of persistent vaginitis. *Obstet Gynecol.* 2014;124(6):1135-1146.
36. Sobel JD, Reichman O, Misra D, Yoo W. Prognosis and treatment of desquamative inflammatory vaginitis. *Obstet Gynecol.* 2011;117(4):850-855.
37. Schwebke JR, Desmond R. A randomized trial of metronidazole in asymptomatic bacterial vaginosis to prevent the acquisition of sexually transmitted diseases. *Am J Obstet Gynecol.* 2007;196(6):517.e1-517.e6.
38. 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.
39. Wiesenfeld HC, Hillier SL, Krohn MA, Landers DV, Sweet RL. Bacterial vaginosis is a strong predictor of *Neisseria gonorrhoeae* and *Chlamydia trachomatis* infection. *Clin Infect Dis.* 2003;36(5):663-668.
40. Ferris DG, Litaker MS, Woodward L, Mathis D, Hendrich J. Treatment of bacterial vaginosis: a comparison of oral metronidazole, metronidazole vaginal gel, and clindamycin vaginal cream. *J Fam Pract.* 1995;41(5):443-449.
41. Symbiomix Therapeutics. FDA approves Symbiomix Therapeutics' Solosec (secnidazole) oral granules for the treatment of bacterial vaginosis in adult women. September 18, 2017. <https://symbiomix.com/fda-approves-symbiomix-therapeutics-solosec-secnidazole-oral-granules-treatment-bacterial-vaginosis-adult-women/>. Accessed October 28, 2017.
42. Bohbot JM, Vicaut E, Fagnen D, Brauman M. Treatment of bacterial vaginosis: a multicenter, double-blind, double-dummy, randomised phase III study comparing secnidazole and metronidazole. *Infect Dis Obstet Gynecol.* 2010;2010.
43. Solosec (secnidazole) [prescribing information]. Newark, N.J.: Symbiomix Therapeutics LLC; September 2017. https://www.accessdata.fda.gov/drugsatfda_docs/label/2017/209363s000lbl.pdf. Accessed October 28, 2017.
44. Koss CA, Baras DC, Lane SD, et al. Investigation of metronidazole use during pregnancy and adverse birth outcomes. *Antimicrob Agents Chemother.* 2012;56(9):4800-4805.
45. Brocklehurst P, Gordon A, Heatley E, Milan SJ. Antibiotics for treating bacterial vaginosis in pregnancy. *Cochrane Database Syst Rev.* 2013;(1):CD000262.
46. Leitch H, Bodner-Adler B, Brunbauer M, Kaidler 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.
47. Sobel JD, Brooker D, Stein GE, et al. Single oral dose fluconazole compared with conventional clotrimazole topical therapy of *Candida* vaginitis. Fluconazole Vaginitis Study Group. *Am J Obstet Gynecol.* 1995;172(4 pt 1):1263-1268.
48. Watson MC, Grimshaw JM, Bond CM, Mollison J, Ludbrook A. Oral versus intra-vaginal imidazole and triazole anti-fungal agents for the treatment of uncomplicated vulvovaginal candidiasis (thrush): a systematic review. *BJOG.* 2002;109(1):85-95.
49. 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.
50. Merkus JM. Treatment of vaginal candidiasis: orally or vaginally? *J Am Acad Dermatol.* 1990;23(3 pt 2):568-572.
51. Reef SE, Levine WC, McNeil MM, et al. Treatment options for vulvovaginal candidiasis, 1993. *Clin Infect Dis.* 1995;20(suppl 1):S80-S90.
52. Sobel JD. Factors involved in patient choice of oral or vaginal treatment for vulvovaginal candidiasis. *Patient Prefer Adherence.* 2013;8:31-34.
53. Sobel JD, Faro S, Force RW, et al. Vulvovaginal candidiasis: epidemiologic, diagnostic, and therapeutic considerations. *Am J Obstet Gynecol.* 1998;178(2):203-211.
54. Ferris DG, Nyirjesy P, Sobel JD, Soper D, Pavletic A, Litaker MS. Over-the-counter antifungal drug misuse associated with patient-diagnosed vulvovaginal candidiasis. *Obstet Gynecol.* 2002;99(3):419-425.
55. Mølgaard-Nielsen D, Svanström H, Melbye M, Hviid A, Pasternak B. Association between use of oral fluconazole during pregnancy and risk of spontaneous abortion and stillbirth. *JAMA.* 2016;315(1):58-67.
56. Howley MM, Carter TC, Browne ML, Romitti PA, Cunniff CM, Druschel CM. Fluconazole use and birth defects in the National Birth Defects Prevention Study. *Am J Obstet Gynecol.* 2016;214(5):657.e1-657.e9.
57. Sobel JD, Kapernick PS, Zervos M, et al. Treatment of complicated *Candida* vaginitis: comparison of single and sequential doses of fluconazole. *Am J Obstet Gynecol.* 2001;185(2):363-369.
58. Sobel JD, Wiesenfeld HC, Martens M, et al. Maintenance fluconazole therapy for recurrent vulvovaginal candidiasis. *N Engl J Med.* 2004;351(9):876-883.
59. Richter SS, Galask RP, Messer SA, Hollis RJ, Diekema DJ, Pfaller MA. Antifungal susceptibilities of *Candida* species causing vulvovaginitis and epidemiology of recurrent cases. *J Clin Microbiol.* 2005;43(5):2155-2162.
60. Sood G, Nyirjesy P, Weitz MV, Chatwani A. Terconazole cream for non-*Candida albicans* fungal vaginitis: results of a retrospective analysis. *Infect Dis Obstet Gynecol.* 2000;8(5-6):240-243.
61. Sobel JD, Chaim W, Nagappan V, Leaman D. Treatment of vaginitis caused by *Candida glabrata*: use of topical boric acid and flucytosine. *Am J Obstet Gynecol.* 2003;189(5):1297-1300.
62. Falagas ME, Betsi GI, Athanasiou S. Probiotics for prevention of recurrent vulvovaginal candidiasis: a review. *J Antimicrob Chemother.* 2006;58(2):266-272.
63. Wang CC, McClelland RS, Reilly M, et al. The effect of treatment of vaginal infections on shedding of human immunodeficiency virus type 1. *J Infect Dis.* 2001;183(7):1017-1022.
64. Kissinger P, Mena L, Levison J, et al. A randomized treatment trial: single versus 7-day dose of metronidazole for the treatment of *Trichomonas vaginalis* among HIV-infected women. *J Acquir Immune Defic Syndr.* 2010;55(5):565-571.
65. Johnson GL. Tinidazole (Tindamax) for trichomoniasis and bacterial vaginosis. *Am Fam Physician.* 2009;79(2):102-105.
66. Cotch MF, Pastorek JG II, Nugent RP, et al. *Trichomonas vaginalis* associated with low birth weight and preterm delivery. The Vaginal Infections and Prematurity Study Group. *Sex Transm Dis.* 1997;24(6):353-360.
67. Lynch C. Vaginal estrogen therapy for the treatment of atrophic vaginitis. *J Womens Health (Larchmt).* 2009;18(10):1595-1606.
68. Management of symptomatic vulvovaginal atrophy: 2013 position statement of The North American Menopause Society. *Menopause.* 2013;20(9):888-902.
69. Owen MK, Clenney TL. Management of vaginitis. *Am Fam Physician.* 2004;70(11):2125-2132.
70. Egan ME, Lipsky MS. Diagnosis of vaginitis. *Am Fam Physician.* 2000;62(5):1095-1104.