Challenging Vaginal Discharge, Lactobacillosis and **Cytolytic Vaginitis**

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Abstract

In women's health, clinicians are often faced with diagnosing and administering treatment to patients who present with an increased vaginal discharge without much other information to work with. It's only once a patient's background is reviewed and lab work is completed that two common reasons behind vaginal discharge with similar symptoms, Lactobacillosis and Cytolytic Vaginitis, can be differentiated and a proper diagnosis can be reached.

Now, we understand that Lactobacillosis and Cytolytic Vaginitis are symptomatically confused by patients as well as inexperienced clinicians. Lactobacillosis involves the presence of abnormal, extremely long Lactobacilli, a keystone species for vaginal health, while Cytolytic Vaginitis mainly involves, increased Lactobacilli, pH between 3.5 and 4.5, presence of cytolysis, and a lowered white blood cell count.

Both present with similar discharge, leading to their conflation. A microscopic examination of the discharge using a wet mount with normal saline can provide 80% diagnostic sensitivity until further testing can be completed.

Keywords: Lactobacillus; Microbiota; Vaginitis; Vaginal Discharge

Introduction

Increased vaginal discharge is the most common reason for women to seek health care advice (1). Establishing a correct diagnosis for symptomatic patients complaining about increased vaginal discharge is a challenge when based only on symptoms and without a proper workup. Analysis of increased vaginal discharge may be completed after obtaining a thorough medical history, review of hygiene practices, a problem-focused physical examination, and targeted laboratory testing (2, 3).

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The collection, preparation, identification, and culture of vaginal specimens are necessary components for successful diagnosis and treatment. Properly performed vaginal wet mount microscopy can be achieved with 80% diagnostic sensitivity in office settings (3). However, two challenging vaginal discharges that puzzle the clinician: Lactobacillosis and Cytolytic Vaginitis.

Physiologic Vaginal Environment

The vaginal microbiome of healthy childbearing age women is approximately 90% composed of *Lactobacillus* species. It is important to recognize that women will normally produce between 1.6 g to 4.8 g of vaginal discharge per day. This discharge includes vaginal epithelial cells that



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provide *Lactobacilli* with a physiologic environment to thrive. Vaginal transudate also prevents dryness, helps to protect against infections, and includes cervical mucous (with immune globulins), Skene, and Bartholin Gland secretions required for normal vaginal function during the reproductive years (2-4).

Lactobacilli shield the vaginal ecosystem from bacterial pathogens, but also partially from fungi and viruses. Lactobacilli achieve these functions through the production of lactic acid, hydrogen peroxide, bacteriocins, bio-surfactants, hydroxyl radicals, organic acids, and arginine deaminases (5). The number of Lactobacilli present in the vagina can be estimated utilizing the Spiegal scale: that runs from 0 to 4 (none = 0, rare = 1, few = 2, moderate = 3, many = 4) (2-4).

The specific mechanisms causing dysfunction in the vaginal ecosystem, are complex. Amongst the factors involved in vaginal dysbiosis, pH, and balanced female hormone levels seem to be critical. The vaginal pH is acidic in most women and is strongly dependent on the presence of functionally normal *Lactobacilli* that will maintain a pH between 3.8 and 4.2. The vaginal fluid of women with healthy vaginas normally indicates a prevalence of *Lactobacilli* microflora composed of *L. crispatus*, *L. gasseri*, *L. jensenii*, and *L. iners* (5) (Figure 1).

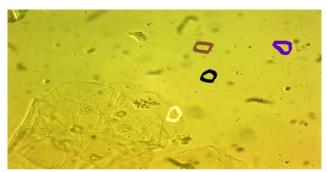


Figure 1: Differentiation of *Lactobacilli*. Black: *L. gasseri*, Red: *L. crispatus*, White: *L. jensenii*, and Blue: *L. iners*

Vaginal Lactobacillosis

Vaginal Lactobacillosis (VL) is a cyclical clinical condition that was first reported by Horowitz *et al.* in 1994. VL presents as an annoying and copious white vaginal discharge leading to an impression by the patient of unremittingly having wet underwear (6). After prospectively studying 30 controls and 37 patients with a clinical condition characterized by profuse and bothersome vaginal discharge, Horowitz *et al.* called the condition VL. Horowitz concluded

that it was associated with extremely long *Lactobacilli* seen in the microscope but reported that the cause of the morphologic transformation observed in the *Lactobacilli* was unknown. Horowitz suggested that the condition could be effectively treated with antibiotics (6). In 1994, Kaufmann *et al.* demonstrated that *Leptothrix* associated with VL are nonbranching, segmented, and gram-positive anaerobic rods that are longer than *Lactobacilli* but shorter than *Candida* filaments (7).

Feo *et al.* had classified them before in the *Lactobacillus* genus based on their biochemical and cultural properties. In vaginal wet mount preparations, the microscope examination of vaginal discharge from patients with VL is characterized by the presence of plentiful, normal, and far longer segmented *Lactobacilli* chains (identified as *Leptothrix*), ranging from $40~\mu m$ to $75~\mu m$ in length (7).

Normal *Lactobacilli* length is between 5 μ m and 15 μ m and the total number of *Lactobacilli* observed in a microscope at 400x magnification is between 15 and 25 per image (Figure 2). In patients with VL, there are two to several times more *Lactobacilli* observed per image. Women experiencing VL also exhibit *Lactobacilli* which are up to eight times longer than normal with many presenting bizarre forms (5, 6, 8).



Figure 2: Vaginal Lactobacillosis: abundant Lactobacilli with bizarre forms

The etiology of VL is still under investigation but it has been linked to other conditions, including antifungal topical medications misuse, Diabetes Mellitus, and Vulvodynia. VL is diagnosed in approximately 15% of patients complaining about copious vaginal discharge (9).

The treatment of VL consists of eliminating the abnormal *Lactobacilli* with oral amoxicillin-clavulanate,

using doxycycline as an alternative (6, 9). The replenishment of normal Lactobacilli then could be achieved by taking oral live Lactobacilli and acidifying the perineum with mild acidic solutions. Lactobacilli have shed some genes to acquired vaginotropism that assists them in vaginal colonization.

Cytolytic Vaginosis

In 1991, Cibley et al. described patients presenting with a condition named "Cytolytic Vaginosis" who complained of a white cheesy vaginal discharge, pruritus, vulvar dyspareunia, and a recurring increase in symptoms more noticeable in the luteal phase (10, 11).

Cibley's diagnostic criteria included the presence of discharge, as well as exclusion of Trichomonas, Gardnerella, and Candida on the wet mount (10). Cibley also observed an increased number of Lactobacilli, a scarcity of white blood cells, evidence of cytolysis, and a pH between 3.5 and 4.5 (Figure 3). Cibley suggested a treatment using sodium bicarbonate douches (10). Douching is no longer recommended and current suggested treatments include topical metronidazole and/or topical steroid creams (10).

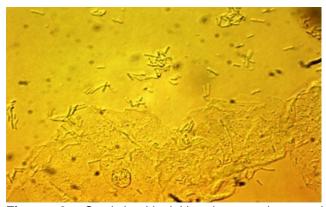


Figure 3: Cytolytic Vaginitis: increased normal Lactobacilli with abundant cytolysis

Controversies

We can now understand that Lactobacillosis and Cytologic Vaginitis are often symptomatically confused by patients and inexperienced clinicians. Many women therefore incorrectly self-diagnose or are incorrectly diagnosed with fungal infection or colonization (12). As a result, many women try overthe-counter topical antifungal medications or are prescribed oral antifungals only to get partial or no relief. The microscopic wet mount examination with an 80% sensitivity provides the patient with a more targeted treatment until confirmation testing is

complete (13).

Evidently, we are in front of two different conditions. Abundant cytolysis is characteristic of Cytolytic Vaginitis, while the clear presence of Leptothrix is the "sine qua non" of Lactobacillosis. Also, should we refer to Cytolytic Vaginitis as Desquamative Vaginitis or as Cytolytic Vaginosis, considering that these conditions may coexist (Figure 4)?

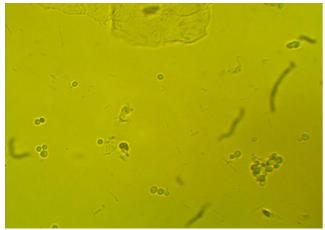


Figure 4: Lactobacillosis and Candida dubliniensis

Knowing how to properly recognize the images of these two symptomatically similar conditions, at the wet mount microscopy with normal saline, will certainly assist in successful differentiation during the patient's visit. This can provide the patient with an 80% diagnostic sensitivity and therefore an improved targeted treatment until confirmation testing is complete.

Conclusion

The correct diagnosis of symptomatic patients with discharge vaginal is challenging. increased Differentiation may be established using patient history, physical examination, and proper laboratory testing. Correctly performed vaginal wet mount microscopy with normal saline can be achieved with an 80% diagnostic sensitivity in an office setting providing the patient with targeted treatment.

Conflict of Interests

Authors have no conflict of interests.

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