

Hidradenitis suppurativa: An underdiagnosed skin problem of women

Gynecologists are uniquely positioned to diagnose this common skin problem



Robert L. Barbieri, MD

Editor in Chief, OBG MANAGEMENT Chair, Obstetrics and Gynecology Brigham and Women's Hospital, Boston, Massachusetts Kate Macy Ladd Professor of Obstetrics, Gynecology and Reproductive Biology Harvard Medical School, Boston

n recent decades the practice of medicine has drifted away from the performance of a physical examination during most patient encounters and evolved toward the more intensive use of history, imaging, and laboratory studies to guide management decisions. For example, it is common for a woman to present to an emergency department with abdominal or pelvic pain and undergo a computerized tomography scan before an abdominal and pelvic examination is performed. Some authorities believe that the trend to reduce the importance of the physical examination has gone way too far and resulted in a reduction in the quality of health care.^{1,2}

Many skin diseases only can be diagnosed by having the patient

Instant Poll

Do you think that the trend to not perform a physical examination has adversely impacted patient care? Without violating HIPAA provisions, are you aware of a case example of how not performing a physical examination adversely impacted patient care?

Tell us at rbarbieri@frontlinemedcom.com Please include your name and city and state.

disrobe and examining the skin. Gynecologists are uniquely positioned to diagnose important skin diseases because, while performing a reproductive health examination, they may be the first clinicians to directly examine the anogenital area and inner thighs. Skin diseases that are prevalent and can be diagnosed while performing an examination of the anogenital region include lichen sclerosus (LS) and hidradenitis suppurativa (HS). The prevalence of each of these conditions is in the range of 1% to 4% of women.³⁻⁵

Failure to examine the anogenital area and insufficient attention to the early signs of LS and HS may result in a long delay in the diagnosis.⁶ In 1 survey, of 517 patients with HS, there was a 7-year interval between the

onset of the disease and the diagnosis by a clinician.⁷ Delay in diagnosis results in increased scarring, which makes it more difficult to effectively treat the disease. In this editorial, I will focus on the pathogenesis, diagnosis, and treatment of HS.

Diagnosis, presentation, and staging

Hidradenitis suppurativa (from the Greek, hidros means sweat and aden means glands) is a painful, chronic, relapsing, inflammatory skin disorder affecting the follicular unit. It is manifested by nodules, pustules, sinus tracts, and scars, usually in intertriginous areas. The diagnosis is made by history and physical examination. The 3 cardinal features of HS are 1) deepseated nodules, comedones, and fibrosis; 2) typical anatomic location of the lesions in the axillae, inguinocrural, and anogenital regions, and 3) chronic relapsing course.⁸

Disease severity is often assessed using the Hurley staging system:

- stage I: abscess formation without sinus tracts or scarring (FIGURE, page 10)
- stage II: recurrent abscesses with tract formation and scarring, widely separated lesions

CONTINUED ON PAGE 10



FIGURE Multiple inflammatory nodules in the genital area without sinus tracts or fistulas, classified as Hurley stage I disease. Image courtesy of Cosmetic Dermatology.

2011;24:226–238. ©2011, Frontline Medical Communications Inc.

• stage III: diffuse or near-diffuse involvement or multiple interconnected tracts and abscesses.

In one report, stage I, II, and III disease was diagnosed in 65%, 31%, and 4% of cases, respectively, indicating that most HS is diagnosed in stage I and suitable for treatment by a gynecologist.⁹

HS typically presents after puberty and women are more commonly affected than men. In one case series including 232 women with HS the regions most commonly affected were: axillae, inguinofemoral, urogenital, and buttocks in 79%, 77%, 51%, and 40% of cases, respectively.¹⁰ Risk factors for HS include obesity, cigarette smoking, tight fitting clothing, and chronic friction across the affected skin area.⁵

Pathogenesis

The pathophysiology of HS is thought to begin with occlusion of the follicle, resulting in follicle rupture deep in the dermis, thereby triggering inflammation, bacterial infection, and scarring. Dermal areas affected by HS have high concentrations of cytokines, including tumor necrosis factor (TNF)-alpha, interleukin (IL)-1-beta, IL-23, and IL-32.^{11,12} Once HS becomes an established process, it is difficult to treat because the dermal inflammatory process and scarring provides a microenvironment that facilitates disease progression. Hence early detection and treatment may result in optimal long-term outcomes.

Treatment

Many recommended treatments for HS have not been formally tested in large randomized trials. A recent Cochrane review identified only 12 high-quality trials and the median number of participants was 27 per trial.13 Consequently, most treatment recommendations are based on expert opinion. Recommended treatments include smoking cessation, weight loss, topical and systemic antibiotics, antiandrogens, antiinflammatory biologics (adalimumab and infliximab), and surgery. Smoking cessation and weight loss are strongly recommended in the initial treatment of HS. Bariatric surgery and significant postprocedure weight loss has been reported to cause a reduction in disease activity.14

Stage I management. For the initial treatment of stage I HS, **clindamycin gel 1%** applied twice daily to affected areas is recommended.¹⁵ Recommended oral antibiotic treatments include **tetracycline 500 mg twice daily for 12 weeks**¹⁶ or **doxycycline 100 mg or 200 mg given daily for 10 weeks** or **clindamycin 300 mg twice daily plus rifampicin 600 mg once daily for 10 weeks**.^{17,18} These antibiotics have both antimicrobial and anti-inflammatory effects.

Hormonal interventions that

suppress androgen production or action may help reduce HS disease activity. For women with HS who also need contraception, an estrogenprogestin contraceptive may help reduce HS disease activity in up to 50% of individuals.¹⁹ The 5-alpha reductase inhibitor finasteride, at high doses (5 to 15 mg daily), has been reported to reduce HS disease activity.20,21 Finasteride is a teratogen, and the FDA strongly recommends against its use by women. Spironolactone, an antimineralocorticoid and antiandrogen, at a dose of 100 mg daily has been reported to reduce disease activity in about 50% of treated individuals and is FDA approved for use in women.²² Among reproductiveage women, spironolactone, which is a teratogen, only should be prescribed to women using an effective form of contraceptive. HS is often associated with obesity and insulin resistance. Metformin 500 mg three times daily has been reported to decrease disease activity.23,24

Stage II or III management. For Hurley stage II or III HS, referral to a dermatologist is warranted. There is evidence that too few people with HS are referred to a dermatologist.25 For severe HS resistant to oral medications, anti-TNF monoclonal antibody treatment with adalimumab (Humira) or infliximab (Remicade) is effective. Adalimumab is administered by subcutaneous injection and is US Food and Drug Administration (FDA)-approved to treat HS. Following a loading dose, adalimumab is administered weekly at a dose of 40 mg.²⁶ Infliximab, which is not FDA approved to treat HS, is administered by intravenous infusion at a dose of 5 mg/kg at weeks 0, 2, and 6, and then every 8 weeks.27

Surgical management. HS is sometimes treated surgically with



laser destruction of lesions, punch debridement, or wide excision of diseased tissue.28,29 There are no high quality clinical trials of surgical treatment of HS. Punch debridement can be performed using a 5- to 7-mm circular skin punch to deeply excise the inflamed follicle. Wide excision can be followed by wound closure with advancement flaps or split-thickness skin grafting. Wound closure by secondary intention is possible but requires many weeks or months of burdensome dressing changes to complete the healing process. Recurrence is common following surgical therapy and ranges from 30% with deroofing or laser treatment to 6% following wide excision and skin graft closure of the wound.³⁰

Physical examination vital to early diagnosis

Delay in diagnosis of an active disease process has many causes, including nonperformance of a physical examination. In a webbased survey of physicians' experiences with oversights related to the physical examination, 3 problems frequently reported were: nonperformance of any portion of the physical examination, failure to undress the patient to examine the skin, and failure to examine the abdomen and anogenital region in a patient with abdominal or pelvic pain.31 Oversights in the physical examination frequently caused a delay in diagnosis and treatment. With both LS and HS, patients may not recognize that they have a skin disease, or they may be embarrassed to show a clinician a skin change they have noticed. Early diagnosis and treatment are essential to achieving a good outcome and make a tremendous difference in the quality of life for the patient. Physical examination is a skill we have learned through diligent study and experience in practice. We can use these skills to greatly improve the lives of our patients.

RBARBIERI@FRONTLINEMEDCOM.COM

Dr. Barbieri reports no financial relationships relevant to this article.

References

- 1. Jauhar S. The demise of the physical examination. N Engl J Med. 2006;354(6):548–551.
- Feddock CA. The lost art of clinical skills. Am J Med. 2007;120(4):374–378.
- Goldstein AT, Marinoff SC, Christopher K, Srodon M. Prevalence of vulvar lichen sclerosus in a general gynecology practice. J Reprod Med. 2005;50(7):477-480.
- Jemec GB, Heidenheim M, Nielsen NH. The prevalence of hidradenitis suppurativa and its potential precursor lesions. J Am Acad Dermatol. 1996;35(2 pt 1):191–194.
- Revuz JE, Canoui-Poitrine F, Wolkenstein P, et al. Prevalence and factors associated with hidradenitis suppurativa: results from two case-control studies. J Am Acad Dermatol. 2008;59(4):596–601.
- Cooper SM, Gao XH, Powell JJ, Wojnarowska F. Does treatment of vulvar lichen sclerosus influence its prognosis? Arch Dermatol. 2004;140(6):702–706.
- Saunte DM, Boer J, Stratigos A, et al. Diagnostic delay in hidradenitis suppurativa is a global problem. Br J Dermatol. 2015;173(6):1546–1549.
- Zouboulis CC, Del Marmol V, Mrowietz U, et al. Hidradenitis suppurativa/acne inversa: criteria for diagnosis, severity assessment, classification and disease evaluation. Dermatology. 2015:231(2):184–190.
- Revuz J. Hidradenitis suppurativa. J Eur Acad Dermatol Venereol. 2009;23(9):985–998.
- Canoui-Poitrine F, Revuz JE, Wolkenstein P, et al. Clinical characteristics of a series of 302 French patients with hidradenitis suppurativa, with an analysis of factors associated with disease severity. J Am Acad Dermatol. 2009;61(1):51-57.
- Shah A, Alhusayen R, Amini-Nik S. The critical role of macrophages in the pathogenesis of hidradenitis suppurativa. Inflamm Res. 2017;66(11):931-945.
- 12. Thomi R, Yerly D, Yawalkar N, Simon D,

Schlapbach C, Hunger RE. Interleukin-32 is highly expressed in lesions of hidradenitis suppurativa. Br J Dermatol. 2017;177(5):1358–1366.

- Ingram JR, Woo PN, Chua SL, et al. Interventions for hidradenitis suppurativa: a Cochrane systematic review incorporating GRADE assessment of evidence quality. Br J Dermatol. 2016;174(5):970– 978.
- Gallagher C, Kirthi S, Burke T, O'Shea D, Tobin AM. Remission of hidradenitis suppurativa after bariatric surgery. JAAD Case Rep. 2017;3(5):436–437.
- Clemmensen OJ. Topical treatment of hidradenitis suppurativa with clindamycin. Int J Dermatol. 1983;22(5):325-328.
- Jemec GB, Wendelboe P. Topical clindamycin versus systemic tetracycline in the treatment of hidradenitis suppurativa. J Am Acad Dermatol 1998;39(6):971-974.
- Gener G, Canoui-Poitrine F, Revuz JE, et al. Combination therapy with clindamycin and rifampicin for hidradenitis suppurativa: a series of 116 consecutive patients. Dermatology. 2009;219(2):148–154.
- Saunte DM, Jemec GB. Hidradenitis suppurativa: advances in diagnosis and treatment. JAMA. 2017;318(20):2019–2032.
- Mortimer PS, Dawber RP, Gales MA, Moore RA. A double-blind controlled cross-over trial of cyproterone acetate in females with hidradenitis suppurativa. Br J Dermatol. 1986;115(3):263–268.
- Joseph MA, Jayaseelan E, Ganapathi B, Stephen J. Hidradenitis suppurativa treated with finasteride. J Dermatolog Treat. 2005;16(2):263–268.
- Randhawa HK, Hamilton J, Pope E. Finasteride for the treatment of hidradenitis suppurativa in children and adolescents. JAMA Dermatol. 2013;149(6):732-735.
- Lee A, Fischer G. A case series of 20 women with hidradenitis suppurativa treated with spironolactone. Australas J Dermatol. 2015;56(3):192–196.
- 23. Arun B, Loffeld A. Long-standing hidradenitis

suppurativa treated effectively with metformin. Clin Exp Dermatol. 2009;34(8):920–921.

- Verdolini R, Clayton N, Smith A, Alwash N, Mannello B. Metformin for the treatment of hidradenitis suppurativa: a little help along the way. J Eur Acad Dermatol Venereol. 2013;27(9):1101–1108.
- Garg A, Lavian J, Strunk A. Low utilization of the dermatology ambulatory encounter among patients with hidradenitis suppurativa: a population-based retrospective cohort analysis in the USA [published online ahead of print September 28, 2017]. Dermatology. doi:10.1159/000480379.
- Kimball AB, Okun MM, Williams DA, et al. Two phase 3 trials of adalimumab for hidradenitis suppurativa. N Engl J Med. 2016;375(5):422–434.
- Grant A, Gonzalez T, Montgomery MO, Cardenas V, Kerdel FA. Infliximab therapy for patients with moderate to severe hidradenitis suppurativa: a randomized, double-blind, placebocontrolled crossover trial. J Am Acad Dermatol. 2010;62(2):205–217.
- Mikkelsen PR, Dufour DN, Zarchi K, Jemec GB. Recurrence rate and patient satisfaction of CO2 laser evaporation of lesions in patients with hidradenitis suppurativa: a retrospective study. Dermatol Surg. 2015;41(2):255–260.
- van der Zee HH, Prens EP, Boer J. Deroofing: a tissue-saving surgical technique for the treatment of mild to moderate hidradenitis suppurativa lesions. J Am Acad Dermatol. 2010;63(3):475–480.
- Mehdizadeh A, Hazen PG, Bechara FG, et al. Recurrence of hidradenitis suppurativa after surgical management: a systematic review and metaanalysis. J Am Acad Dermatol. 2015;73(5 suppl 1):S70–S77.
- Verghese A, Charlton B, Kassirer JP, Ramsey M, Ioannidis JP. Inadequacies of physical examination as a cause of medical errors and adverse events: a collection of vignettes. Am J Med. 2015;128(12):1322-1324.e3