

V is for Vagina, Vulva, Vestibule

and all things amazing

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Let's start with a normal vagina...

- The vagina is a dynamic ecosystem made up of 10^9 bacterial colony forming units
- It discharges – that's normal!
 - Clear, white, odorless
- pH 3.5-4.5
 - pH blood ~ 7.4
 - pH semen ~ 7.5
- No two look the same...this too is normal

Vaginitis

*Vaginal discharge, vulvar pruritus, irritation, and odor

Bacterial Vaginosis 40-50%

- Off white/gray, thin, malodorous discharge
- No dyspareunia
- pH >4.5
- Positive Amine test (70-80% of patients)
- Clue cells (>20% of epithelial cells)
- pH also elevated in trich, atrophy, and DIV

Vulvovaginal Candidiasis 20-25%

- Thick, white, clumpy discharge
- Pruritus, soreness, dyspareunia
- pH 4.0-4.5
- Negative Amine test
- Pseudohyphae (40% of patients), budding yeast for nonalbicans Candida
- Also consider contact irritant dermatitis or vulvodynia

Trichomoniasis 15-20%

- Thin, green-yellow, malodorous discharge
- Burning, *postcoital bleeding*, dyspareunia, *dysuria*
- pH 5.0-6.0
- Amine test often positive
- Motile trichomonads (60% of patients)
- Also consider DIV, atrophic vaginitis, erosive lichen planus

BV

- Most common cause of vaginal discharge in women of childbearing age (40-50% of cases)
- Overgrowth of bacteria which is normal vaginal flora
- Absence of inflammation
 - *Vaginosis vs Vaginitis*
- Sexual Activity:
 - Risk factor
 - Not classified as an STI
 - 25-50% prevalence in WSW
 - Increased risk with presence of other STIs
- Other risk factors:
 - African American
 - Cigarette smoking
 - Douching
 - Possible genetic component
 - Not associated with immunosuppressive states
- Reported more frequently after vaginal intercourse and after menstrual cycle
- Symptoms may resolve spontaneously

BV linked to:

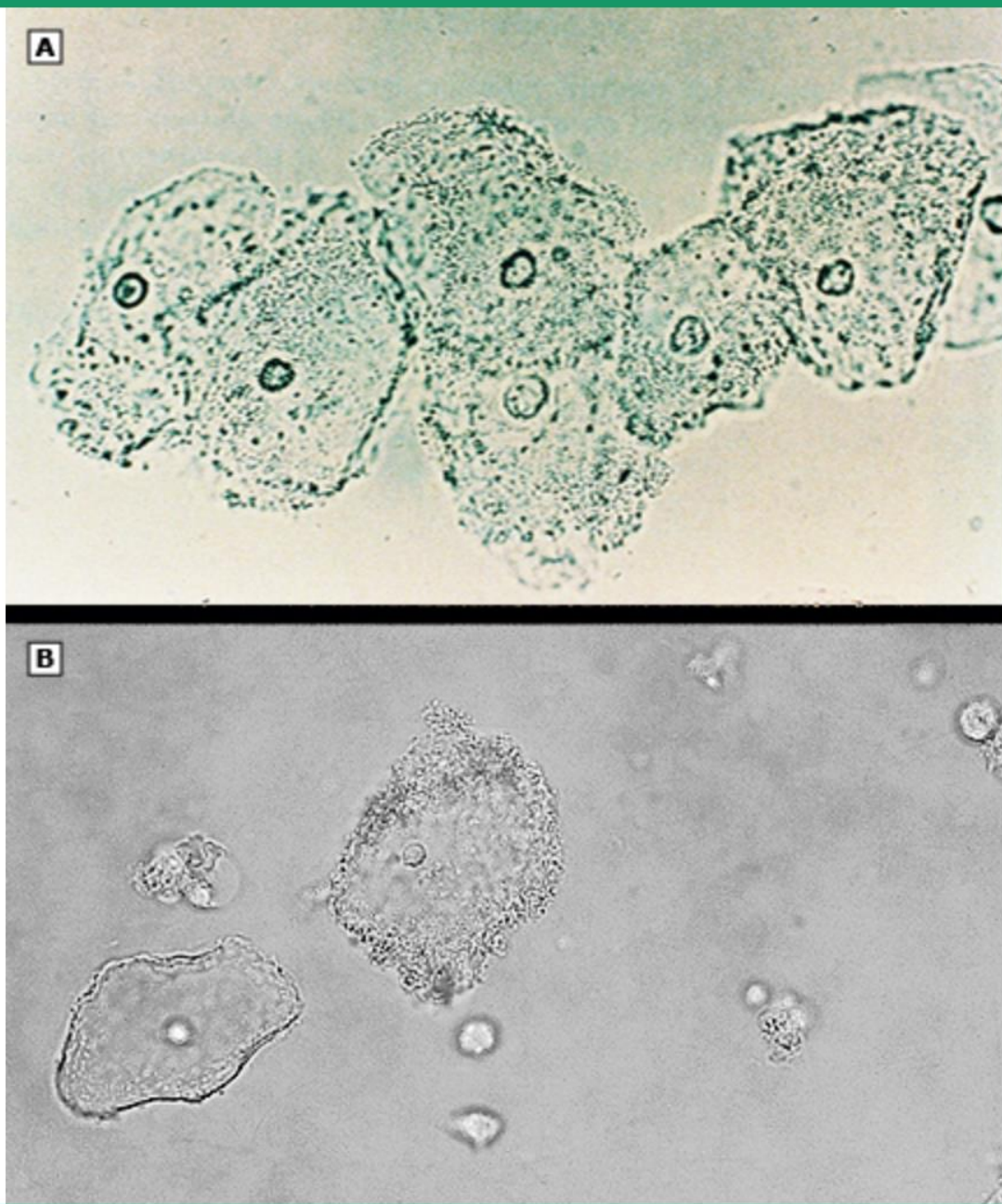
- PROM
- Preterm delivery and low birth weight
- HIV, N. gonorrhoea, C. trachomatis, and HSV-2
- PID
- Post op infections after Gyn procedures
- Recurrence of BV

BV Diagnosis

- Amsel Criteria – clinical criteria
 - Characteristic vaginal discharge: homogenous, thin, gray-white that smoothly coats the vaginal walls
 - pH >4.5
 - Positive whiff amine test
 - Clue cells on saline wet mount (>20% of the epithelial cells)

****At least 3 of the 4 criteria must be met for diagnosis****

- Gram stain
 - Gold standard
- Culture
- DNA probe



Clue Cells

(A) Wet mount showing characteristic clue cells. Note that the epithelial cells are so heavily covered by bacteria as to obscure the margins.

(B) A clue cell. The vaginal epithelial cell on the right has shaggy borders obscured by coccobacilli (1003 magnification). The more normal appearing epithelial cell on the left has sharper borders.

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BV Treatment

- Oral or vaginal
- Treat symptomatic patients (40-50%)
- Metronidazole
 - *500mg po BID x 7d*
 - *1 applicatorful (5g) PV qhs x 5d*
- Clindamycin
 - *2% cream 1 applicatorful (100mg) PV qhs x 7d (may use x 3d if not pregnant)*
 - *300mg po BID x 7d*
 - *Ovules 100mg PV qd x 3d*

Treatment in Pregnancy

- *Only treat if symptomatic*
- Metronidazole 500mg PO BID x 7d
- Metronidazole 250mg PO BID x 7d
- Clindamycin 300mg PO BID x 7d

Vulvovaginal Candidiasis

- 2nd most common cause of vaginitis symptoms (after BV)
- Candida sp. also normal vaginal flora
- 75% of women will have at least 1 episode, 40-45% will have 2 or more
- Prevalence increases with age up to menopause
- 80-92% of episode caused by Candida albicans (remainder mostly C. glabrata)
- Risk factors:
 - DM
 - Antibiotic use
 - Increased estrogen levels
 - Immunosuppression
 - Contraceptive devices
 - Sexual activity – not an STI, but sexually relevant

Candidiasis Diagnosis

- pH 4 to 4.5
- Wet mount
- Culture **IF** necessary (gold standard)
 - Taken from vaginal wall (not cervix)
- Self diagnosis should be discouraged
 - Even with a history of confirmed VVC, women are accurate in self diagnosis only 35% of the time

Candida hyphae

True hyphae of *Candida albicans*



True hyphae (as opposed to pseudohyphae) elongate through a process of apical synthesis that does not involve budding. Since buds are not present at the hyphal tips, the hyphae do not exhibit periodic constrictions associated with the budding process.

Courtesy of Wiley Schell, MS.

Candidiasis Treatment

- Treat symptomatic infection
- May take 2 days or 2 weeks to respond to treatment, depending on severity
- No need to treat sexual partners, may want to abstain during treatment due to inflammation and discomfort
- Oral and topical preparations have similar cure rates, but most women prefer convenience of oral treatment
- Fluconazole (Diflucan) 150mg x 1, may repeat in 72 hours if necessary.
 - Complicated infection could require 2-3 doses, one every 72 hours –OR- 7-14 days of topical azole therapy (vs 1-3 days for uncomplicated infection)
- Ibrexafungerp (Brexafemme) 150mg 2 po BID x 1 day
 - Fungicidal
- *C. glabrata*, **if symptomatic**, treated with boric acid 600mgPV q hs x 2 wks, flucytosine cream PV, and/or amphotericin B cream 4-10% 5g qhs x 2 wks

Treatment in Pregnancy

- 7 day topical regimens are recommended - OTC
- Fluconazole is not recommended

Trichomoniasis

- Most common non-viral STI worldwide
- Estimated 1 million cases annually in the US (\$24 million)
- Often asymptomatic (~50-70%)
- Affects women > men
- Co-infection with BV 60-80%
- Can cause post coital bleeding
- Strawberry cervix (2%)
- Untreated may last months to years and may progress to urethritis or cystitis

Trich linked to:

- PROM
- Preterm delivery
- Cervical neoplasia
- Post-hysterectomy cuff cellulitis or abscess
- PID in women with HIV
- Infertility

Trichomonas Vaginalis Diagnosis

- Wet mount shows mobile trichomonads (60-70% of the time)
 - Motion is jerky and spinning
 - Remain motile for 10-20 minutes after sample collection
- NAAT (gold standard)
- Culture if NAAT not available

Trichomonas Vaginalis Treatment

- Treat symptomatic AND asymptomatic infection
- Curative treatment with metronidazole or tinidazole
- Abstain from intercourse during treatment (or at least 7 days after last antibiotic if using a 1 day course)
- Expedited partner treatment
- Single 2g dose (500mg po x4)
- Alt: Metronidazole 500mg PO BID x 7d
- *Do not consume alcohol for 24 hours after metronidazole tx and for 72 hours after tinidazole tx
- Retest all women 2 wks– 3 months after completing treatment

Treatment in Pregnancy

- Treat all symptomatic women, regardless of gestational age

Trichomonads

Trichomonas vaginalis



High power microscopy revealing *Trichomonas vaginalis* with easily identified flagella.

Courtesy of Jack D. Sobel, MD and William E. Secor.

Recurrent infections

- Defined as ≥ 4 episodes of symptomatic infection in 1 yr
- Speciate candida in refractory or recurrent cases
- **BV** – metronidazole gel twice weekly for 4-6 months
- **Candida albicans**– oral fluconazole 150mg every 72 hours x 3 doses, then once weekly x 6-12 months
- **Candida glabrata** – Boric acid 600mg daily x 3 wks with flucytosine 17% crm
- **Trich** – metronidazole 500mg BID x 7d OR tinidazole 2g PO x 1
 - If that also fails, consider tinidazole or metronidazole 2g PO qd x 5d
- Be sure to address sexual dysfunction and relationship discord that may occur with recurrent vaginitis
- Probiotics unproven for vaginal health
 - RepHresh ProB

Desquamative Inflammatory Vaginitis

- Unknown etiology
- Non-infectious
- More common in peri-menopause
- Pain (dyspareunia, vestibular pain, burning)
- Copious exudative vaginitis
 - Requiring change of pad or underwear several times a day
- Profuse vaginal discharge
 - Yellow, gray or green

DIV Diagnosis

*ALL of the following required:

- At least one of the following:
 - Vaginal discharge
 - Dyspareunia
 - Pruritus
 - Burning
 - Irritation
- Vaginal inflammation (spotted ecchymotic rash, erythema, focal or linear erosion)
- pH > 4.5
- Saline microscopy showing increased numbers of parabasal and inflammatory cells (Leukocyte to epithelial cell ratio is > 1:1)

DIV Differential Diagnosis

- Severe atrophic vaginitis
- Erosive lichen planus
- Pemphigus vulgaris
- Cicatricial pemphigoid

DIV Treatment

- Clindamycin 2% crm 4-5g PV qhs x 4-6 wks
- Hydrocortisone 10% crm 3-5g PV qhs x 4-6 wks
- Estradiol if postmenopausal
- For mild disease:
 - Hydrocortisone 25mg rectal supp ½ or 1 placed in the vagina BID
- Treatment continued until complete remission
 - Based on symptoms on microscopy
- Follow monthly for several months after remission to ensure stability
- If relapse occurs, utilize drug not used for initial treatment until remission, then taper

Vulvar dystrophies

- Vulvoscopy!!!
- Biopsy if there is a lesion – use a 4mm punch biopsy and suture
 - Only exception would be in pediatrics
- Send all biopsies to a dermatopathologist – give them as much background info as possible – even a picture of the lesion if possible
- Do NOT empirically treat with steroids!
- All Lichens are not created equal – educate your patients that they need to know the entire name

Lichen Sclerosus

- Likely autoimmune
 - Consider associated auto-immune disorders with diagnosis – thyroid, alopecia areata, vitiligo, pernicious anemia, lichen planus, diabetes
- Potential Koebner phenomenon
 - May be precipitated by trauma, injury, sexual abuse in certain genetically predisposed women
- Prevalence
 - 1 in 70 women
- Females – any age
 - Highest incidence in premenarchal and postmenopausal ages
- 85-98% of LS is found in the anogenital region

LS

- Lack of estrogen can worsen symptoms, but estrogen therapy is not curative
 - Childhood LS does not resolve at puberty due to estrogen, but symptoms may decrease
- Often misdiagnosed as yeast, HSV, or vitiligo
- 3-6% risk of developing squamous cell carcinoma
- Women with LS are likely to be less sexually active due to pain or embarrassment (vaginal intercourse, oral intercourse, masturbation)

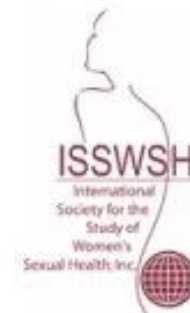
Signs and Symptoms

- Symptoms
 - Pruritus
 - Burning
 - Dyspareunia
 - Dysuria
- Signs
 - “Cigarette paper” crinkling
 - Fissures
 - Waxy look
- Scarring and architectural changes
 - Fusion of clitoral hood
 - Phimosis of clitoris
 - Resorption of labia minora
 - Narrowing of introitus caused by recurrent tearing

Lichen Sclerosus



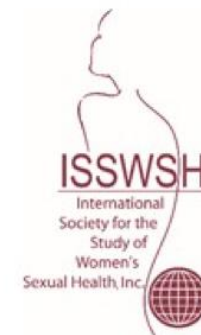
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Lichen Sclerosus



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Lichen Sclerosus



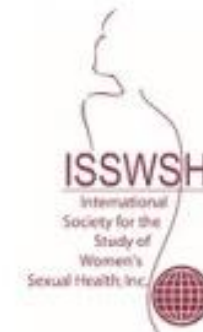
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Lichen Sclerosus Confined to Perineum



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Lichen Sclerosus: Classic Presentation



- LS is identified by loss of pigmentation, texture changes, ecchymosis, alterations in labial architecture and fissuring

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Lichen Sclerosus on Abdomen and Breast



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LS Treatment

- Clobetasol 0.05% ointment qhs after soaking
 - Taper frequency to 1-2x/wk and/or taper potency when all LS and symptoms have resolved (not just when symptoms have resolved)
 - Will likely take 4-6 weeks – re-evaluate 4 wks after starting topical steroid
- Testosterone does not work better than petrolatum ointment alone
- Vaginal or systemic estradiol is not therapeutic for LS
- Current research focused on using topical macrolide immunosuppressants

LS Treatment

- Surgery may be considered if necessary for narrowing of the introitus
- May also surgically correct clitoral phimosis
 - Only after all active disease has resolved

Lichen Planus

- Erosive, papular, hypertrophic lesions of the vulva
 - May affect vagina as well
- Affects 0.5-2% of the population
 - Vulvar LP is a subtype of LP
- Most common in women ages 50-60
- Can co-exist with LS
 - LS NEVER extends above the hymen
- May resemble LS
 - Particularly when late agglutination and architectural distortion occurs
- Always biopsy to confirm diagnosis
- Can affect skin, mucosa, nails, and scalp

Signs and Symptoms

*consistent or intermittent

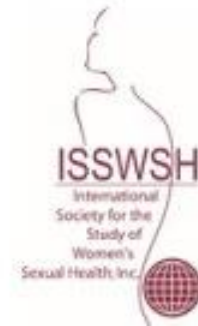
- Symptoms
 - Burning pain
 - Severe dyspareunia
 - Pruritus
 - Post coital bleeding
 - Vaginal stenosis
 - Sticky yellow discharge
- Signs
 - Red plaques on mucous membranes (vulva, vagina, mouth)
 - White "lacey" edges or violaceous borders

Lichen Planus: Classic Presentation



- LP with painful vulvar erosion and irregular white lacy border (Wickham Striae)

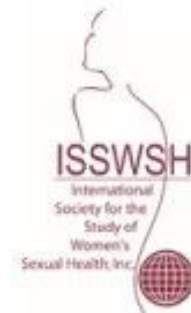
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Erosive Lichen Planus



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Erosive Lichen Planus



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Erosive vulvar LP



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Erosive oral LP



Buccal mucosa



Gingiva

Erosive vulvar LP



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Cutaneous LP on volar wrists and ankles



Fair skin



Dark skin

LP Treatment

- Ultrapotent corticosteroids (Clobetasol)
- Tacrolimus or pimecrolimus (caution – absorbed from vagina)
- Systemic steroids, retinoids, cyclosporine, methotrexate
- Vaginal dilators essential to prevent vaginal stenosis from synechiae

Lichen Simplex Chronicus: Clinical Presentation

- End stage of itch-scratch cycle
- May be superinfected with yeast or bacteria
- Symptoms:
 - Intense pruritus
 - Temporary relief with scratching
- Signs
 - Thick, lichenified skin
 - Erythema – namely labia majora
 - Potential erosions, fissuring, broken hairs, alopecia, exaggerated skin marking

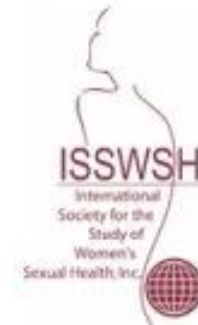
Lichen Simplex Chronicus

- AKA: vulvar eczema, vulvar dermatitis, atopic dermatitis, neurodermatitis
- Triggered by:
 - Irritants
 - Allergens
 - Infections
 - Vulvar intraepithelial neoplasia
 - Mast cell/histamine mediated

Lichen Simplex Chronicus



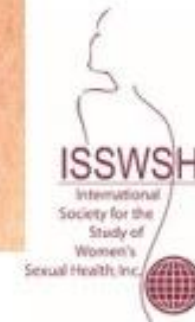
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Lichen Simplex Chronicus



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Loss of architecture



The distinction between the labia majora and minora is lost, and the clitoris becomes buried under the fused prepuce.

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<http://www.blackwell-science.com>.

Vulvar lichen simplex chronicus



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UpToDate®

LSC Treatment

- Discontinue any irritants
 - Soaps, detergents, douches, lotions, etc.
- Sitz baths
 - Warm water 1x/d for 1—15 minutes
- High potency topical corticosteroid ointment
 - Rub in for 1-2 minutes
- Night time routine
 - Amitriptyline 10-50mg qhs
 - Ice packs (frozen peas) at bedtime to help prevent scratching during sleep
- Treat underlying infection
 - Amox/clavulanic acid + fluconazole
- Current research: Pimecrolimus vs corticosteroid

Contact dermatitis vulva



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Genitourinary Syndrome of Menopause

- AKA: Vulvovaginal atrophy, atrophic vaginitis, urogenital atrophy
- Can occur in any low estrogen state
 - Postpartum
 - Breastfeeding
 - Antiestrogenic medications
 - Hypothalamic amenorrhea
 - Pre-/peri-menopausal
- Multiple surveys from 2008 to present on women's views of the impact of GSM
 - Consistent finding of negative impact of GSM on sexual health
 - Barriers to treatment

Numerous barriers

- 70% of women with GSM have not discussed it with the HCP
- 30% of women with GSM had not spoken to anyone
 - Embarrassment, private, doesn't concern others, just part of growing old, others don't want to hear about vaginal problems
- 31% prefer that HCP initiate the conversation
- Ageism, sexism, lack of awareness, cultural factors

Prevalence of GSM

- Estrogen maintains thick, robust, rugated vaginal tissue which is rich in glycogen – ultimately maintains acidic vaginal environment
- Prevalence of vaginal dryness:

Reproductive age	Early menopausal transition	Later menopausal transition	3 yrs post menopause
3%	4%	21%	47%

Estrogens!

E1

- Estrone

E2

- Estradiol

E3

- Estriol

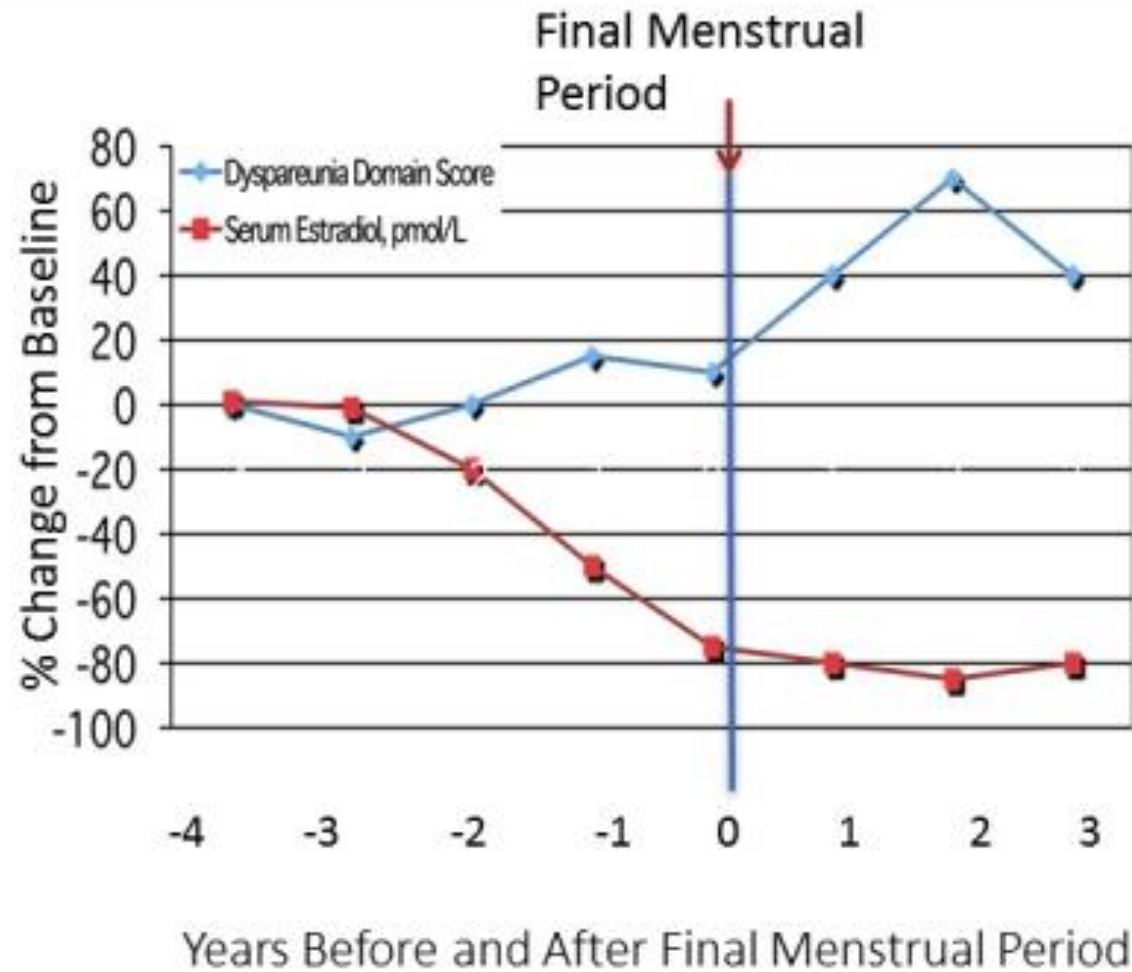
Atrophic vaginitis



External genitalia show scarce pubic hair, diminished elasticity and turgor of the vulvar skin, decreased introital moisture, and fusion of the labia minora.

Courtesy of Aron Schuftan, MD.

Estrogen Decline and Dyspareunia



Diagnosis

- Made clinically based on signs and symptoms
- Assess degree of introital stenosis with a gloved finger prior to attempting to insert pediatric speculum
- Labs available, but typically not necessary
- Vaginal Maturation Index
 - Assess quantity and type of vaginal epithelium cells present on smear per 100
 - Premenopausal:
 - 40-70 intermediate cells
 - 30-60 superficial cells
 - 0 parabasal cells

Vaginal Maturation Index¹⁻³

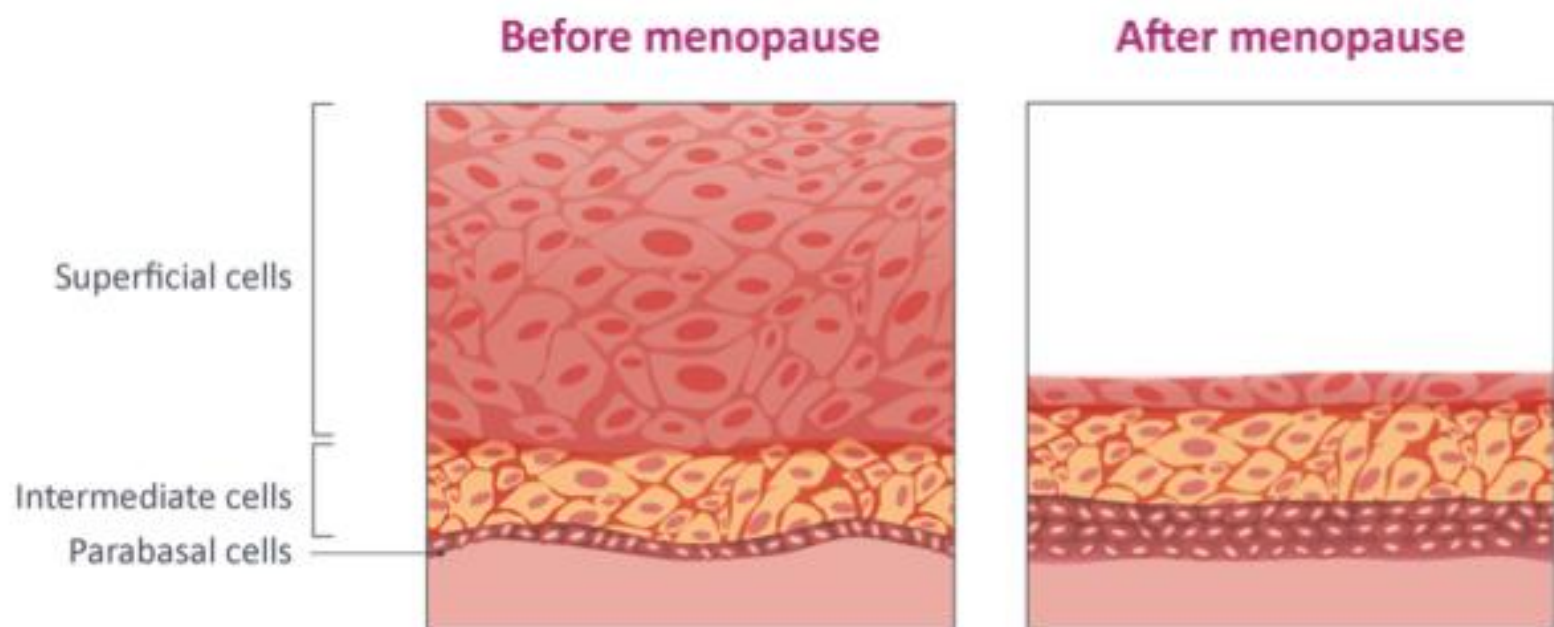
Postmenopausal vaginal epithelium:



% parabasal cells



% superficial cells



GSM Signs and Symptoms

- Dryness and insufficient moisture
- Dyspareunia
- Itching
- Burning
- Soreness
- Tightness
- Loss of elasticity
- Recurrent UTIs
- Thinning of vaginal tissue
- Vaginal discharge
- pH > 5.0
- Mucosal defects (petechiae, microfissures, ulcerations, inflammation)
- Shortening, fibrosis, obliteration of vaginal vault
- Narrowing of introitus
- Smoothing of fornix, flattening of vaginal rugae
- Diminished blood flow
- Erythema early, pallor later

Non-prescription therapies for GSM

- **Moisturization**
 - **Lubrication**
 - **Dilation**
 - **Massage**
 - **Vibration**

Prescription therapies for GSM

- Local therapies vs oral therapies
- Vaginal estrogen
- Vaginal testosterone (compounded)
- Vaginal DHEA
- Ospemifene (Osphena)

Vaginal estrogen preparations

- Low dose local treatment
 - $\leq 50\text{mcg}$ estradiol or $\leq 0.3\text{mg}$ conjugated estrogen/ 0.5mg cream
- Estradiol 10mcg tablets (*Vagifem*) qhs x 2 wks, then 2-3 nights a week
- Estradiol 7.5mcg ring (*Estring*) inserted and replaced every 90 days
- Estradiol $100\text{mcg}/1\text{g}$ crm (*Estrace*) 2-4g PV qhs x 2 wks, then 1g 2-3 nights a week
- Conjugated estrogen $0.625\text{mg}/1\text{g}$ crm (*Premarin vaginal*) 0.5-2g PV qhs x 2 wks, then taper to maintenance dose 2-3 nights a week
- Protection of the uterine lining with a progestin for **low dose** vaginal estrogen therapy is not necessary
- Estradiol 50-100mcg ring (*Femring*) – administered vaginally but this is a **SYSTEMIC** therapy!!!

Systemic absorption of vaginal estrogens

Table 1. Estradiol Preparations and Maximum Annual Delivered Dose

Product name	Route/Type of administration	Typical regimen	Nominal daily delivery rate or administered lowest approved dose (mg/day)	Typical serum level (pg/mL)	Maximum annual delivered dose (mg) ¹
Vaginal estradiol					
Vagifem	Vaginal tablet	1 Tablet daily × 14 then 2 × weekly	10 µg	4.6	1.14
Estring	Vaginal ring	1 Ring vaginally q 3 months	7.5 µg	8.0	2.74
Estrace	Vaginal cream	1 g cream vaginally q week ²	variable ²	NA	7.1
FemRing	Vaginal ring	1 Ring vaginally q 3 months	0.05 mg	40.6	18.25
Oral estradiol					
Estrace tablets and generics	Oral tablet	1 Tablet p.o. qd	0.5 mg	55.4	182.5
Transdermal estradiol					
Divigel ³	Gel	0.25 mg packet qd	0.003	9.8	1.09
Estrogel	Gel	0.75 mg/pump qd	0.035	28.3	12.78
Evamist ³	Spray	1.53 mg spray qd	0.021	19.6	7.67
Climara ⁴	Patch	1 Patch weekly	0.025	22	9.13
Menostar	Patch	1 Patch weekly	0.014	13.7	5.11
Vivelle-Dot ⁵	Patch	1 Patch twice weekly	0.0375	34	12.78

Ospemefine

- Selective Estrogen Receptor Modulator (SERM)
- 60mg PO qd with a light snack
- Estrogen agonist in vagina
- Appears to have no clinically significant estrogenic effect on endometrium or breast
- Side effects
 - Hot flashes!
 - Potential risk of VTE

Vaginal DHEA

- Prasterone (Intrarosa) 6.5mg inserts PV q hs
- Approved 11/16
- Precursor of sex steroids
 - Converted intracellularly into active androgens and/or estrogens
 - Specifically testosterone and estradiol
- Systemic levels remain similar to that of unsupplemented postmenopausal women at 52 weeks
- Did note a small but significant increase in estrone (not estradiol) levels systemically – but still within normal postmenopausal levels
- As with low dose estrogen preparations – unknown effect with small increases in systemic levels

Vulvodynia

Definition:

- Vulvar pain of at least 3 months duration, without clear identifiable cause, which may have potential associated factors (2015 Consensus terminology)
- Localized, Generalized, or Mixed
- Provoked, Spontaneous, or Mixed
- Onset – Primary or Secondary
- Temporal pattern – intermittent, persistent, constant, immediate, delayed

Associated factors

- Co-morbid with other pain syndromes (BPS, fibromyalgia, IBS, TMJ)
- Genetics
- Hormonal factors
- Inflammation
- Musculoskeletal (overactive pelvic floor)
- Neurologic (Central or Peripheral)
- Psychosocial
- Structural defects

Persistent Vulvar Pain: Diagnostic and Treatment Algorithm

ANDREW T. GOLDSTEIN, MD FACOG IF
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A proposed diagnostic and treatment algorithm for vulvodynia, vulvar pain, and dyspareunia. Comments and criticisms are always welcome. It is essential that clinicians and researchers differentiate between different causes of vulvar pain to improve treatment and research.

INTROITAL DYSPAREUNIA & VULVAR PAIN: A diagnostic and treatment algorithm

VESTIBULODYNIA

TENDERNESS THROUGHOUT THE ENTIRE VESTIBULE

HORMONALLY MEDIATED VESTIBULODYNIA
PE: Gland cells are erythematous, mucosal pallor with overlying erythema, decreased size of labia minora and clitoris
LABS: High SHBG, low free testosterone
CAUSES: hormonal contraceptives, spironolactone, Tamoxifen, Aromatase inhibitors, oophorectomy, amenorrhea, lactation
TREATMENT: Stop medications, topical estradiol combined with topical testosterone. Topical estradiol 0.01%/testosterone 0.1% in a hydroalcoholic base QD. May substitute with 0.02% or the estradiol in women with severe atrophy/hypertrophy/agenesis.

INFLAMMATORY VESTIBULODYNIA
HC: chronic infections, allergic reactions, copious yellowish discharge
PE: erythema, leukorrhea, induration, vaginal mucosal tenderness, cervicitis/ectropion
CAUSES: desquamative inflammatory vaginitis, chronic candidiasis (see below), latex allergy/ semen allergy
TREATMENT: interferon 1.5 million units SQ TID for 12 doses (if within 6 months), Singular, Neogen, Topical Clonidine, SQ Tramadol, Capsaicin 0.025% 20 minutes QHS for 12 weeks, gabapentin 4% cream, Vulvar vestibulodynia if failed conservative treatment

NEUROPROLIFERATION

CONGENITAL NEUROPROLIFERATIVE VESTIBULODYNIA
HC: Pain since first tampon use, speculum insertion, and coitus. No pain free sex. Late colesche > 25 years old.
PE: tenderness of the entire vestibule from Herta line to the hymen, often with erythema that worsens after touch with cotton swab. Unilateral hypersensitivity in approximately 62% of women.
LABS: increased density of c-epineph receptors if using 5-100 of PGP 9.5
TREATMENT: VULVAR VESTIBULECTOMY

ACQUIRED NEUROPROLIFERATIVE VESTIBULODYNIA
HC: allergic reaction
PE: chronic yeast infection
LABS: polymorphisms in IL1RA, MBL, IL1B associated with vulvitis, lichen, sensitive skin
TREATMENT: interferon 1.5 million units SQ TID for 12 doses (if within 6 months), Singular, Neogen, Topical Clonidine, SQ Tramadol, Capsaicin 0.025% 20 minutes QHS for 12 weeks, gabapentin 4% cream, Vulvar vestibulodynia if failed conservative treatment

DESQUAMATIVE INFLAMMATORY VAGINITIS
HC: Copious yellow vaginal discharge that turns underwear or requires a pantyliner, vulvar pruritus where discharge dries
PE: Copious leukorrhea, vaginal mucosa erythema, cervicitis, cervical ectropion
CAUSES: Unknown but current hypotheses include: infection pathogen, immune lichen planus, vulvovaginal atrophy, genital ectropion
TREATMENT: intravaginal boric acid/undecylenol cream, hydrocortisone 1 significant estrogen

RECURRENT CANDIDIASIS
PE: erythema, induration, thin fissures, perianal erythema, Discharge is often thin and yellow, not "white cottage cheese"
LABS: Hyphae and increased WBCs on wet mount. Positive cultures
CAUSES: Diet high in simple sugars, antibiotics, OCPs
TREATMENT: Decrease dietary sugars and take probiotics (Probiecta), Oral Nystatin 500,000 units TID for three months + Fluconazole 150mg QD days 1-4 doses 1x a week for 3 months.

HYPERTONIC PELVIC FLOOR MUSCLE DYSFUNCTION
-Pain at 4.5 o'clock if hypertonus of pubococcygeus
-Pain at 6 o'clock if hypertonus of puborectalis
-urinary symptoms if it involves coccygeus (frequency, sensation of incomplete emptying, hesitancy)
-constipation, rectal fissures, hemorrhoids if it involved puborectalis
-associated with ANXIETY, low back pain, scoliosis, hip pain, "holding urine", excessive core strengthening exercises
TREATMENT: PELVIC FLOOR PHYSICAL THERAPY, DIAZEPAM SUPPOSITORIES, VAGINAL DILATORS, HOME PELVIC FLOOR EXERCISES, BOTOX INJECTION

PAIN EXTENDS OUTSIDE THE VESTIBULE (physical exam only, not subjective)

PAIN CONFINED TO THE POSTERIOR VESTIBULE

PUDENDAL NEURALGIA
-PN tender at ischial spine
-unilateral or significantly greater on one side
-history of coccyx trauma
-history of hip pain or labral tear
-better with lying prone-standing, worse with sitting
-pain improved temporarily with PN block
TREATMENT: SERIAL PN BLOCKS, GABAPENTIN, LYRICA, PUDENDAL NERVE NEUROMODULATION

PERSISTENT GENITAL AROUSAL DISORDER
Causes: Pudendal neuralgia, Tarlov cyst, pelvic varicocele, mass along dorsal nerve of clitoris, change in psychotropic medicine, EDS
Dx: tenderness at ischial spine, MRI, pudendal nerve block, dorsal clitoral nerve block

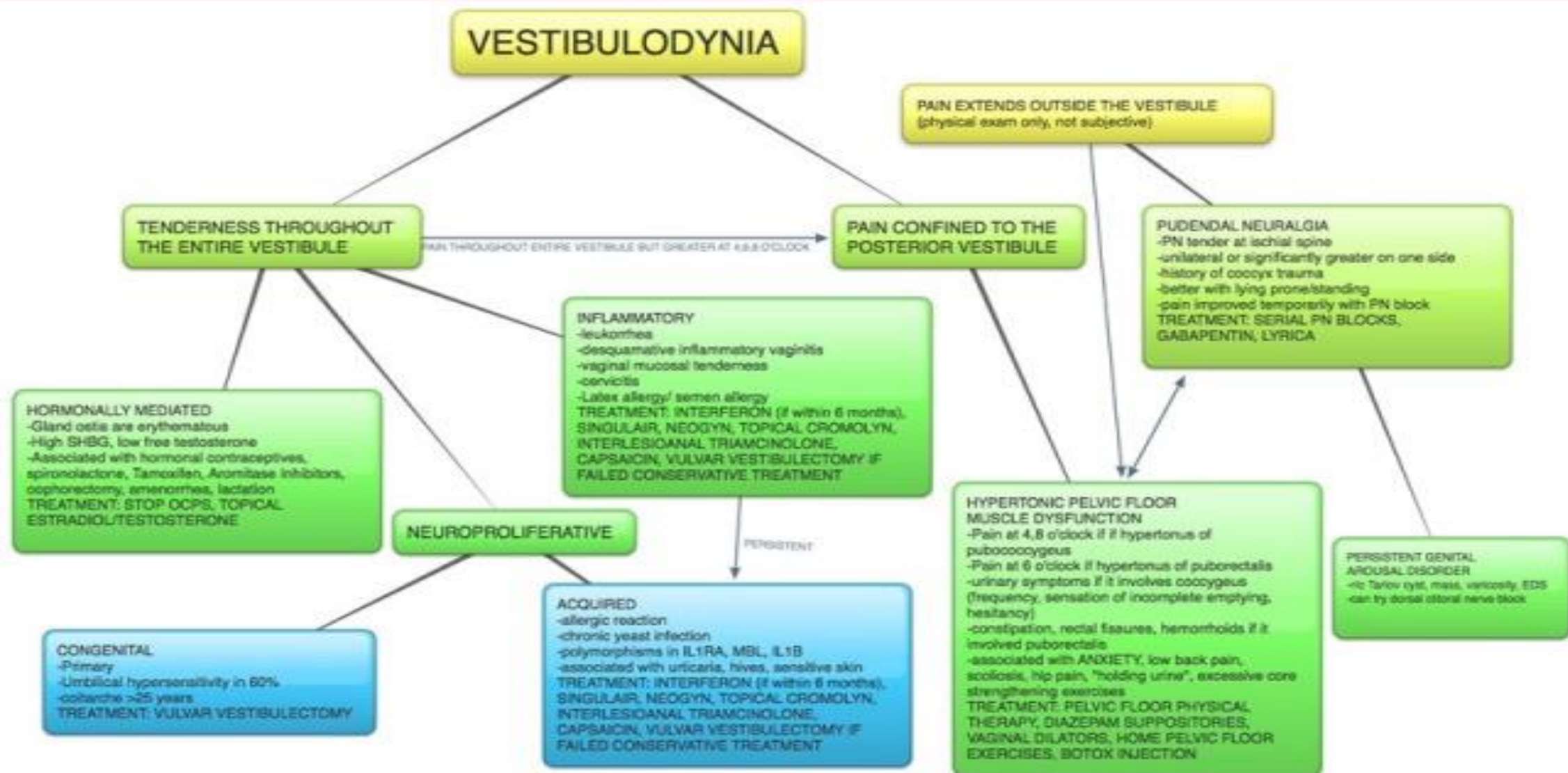
LICHENIFICATION, ULCERATION, RESORPTION OF THE LABIA MINORA, CLITORAL PHIMOSIS, NARROWING OF THE INTROITUS WITH EVIDENCE OF FISSURING

LICHEN SCLEROSIS
Anogenital in a "figure 8" distribution but does not go inside the vagina.
AFFECTS 1/60 WOMEN
3-5% MALIGNANT TRANSFORMATION (VULVOSCOPY NECESSARY)
BIOPSY BEFORE TREATMENT
TREATMENT: CLOBETASOL OINTMENT, SQ TRAMADOLONE, SURGERY FOR PHIMOSIS OR RECURRENT TEARING (VULVAR GRANULOMA FISURATUM)

LICHEN PLANUS
Affects the squamous epithelium of the vulva and causes ulceration in the vestibule (Wickham's stria)
Affects mucous membrane of the mouth and vagina. Can cause synechiae/scarring of the vagina.
PREMALIGNANT
TREATMENT: CLOBETASOL, ELIDEL, PROTOPIC, NEED TO TREAT VAGINA- USE MEDS ON VAGINAL DILATORS, SYSTEMIC STEROIDS OR OTHER IMMUNOSUPPRESSANTS MAY BE NEEDED



Vestibulodynia: Common Etiological Pathways





Vulvodynia physical exam

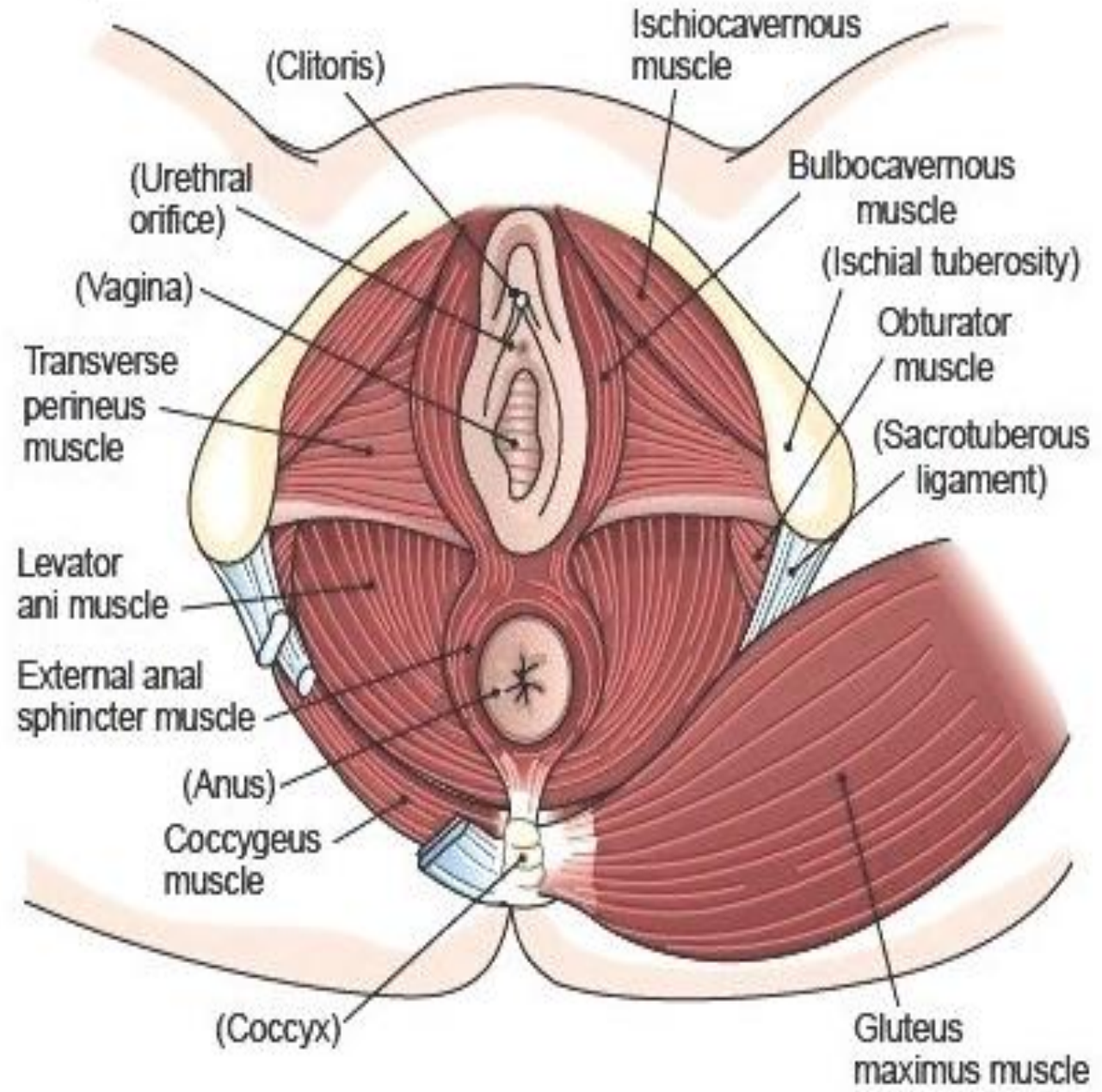
- Let the patient know she is in charge
- Ask her to let you know if she needs a break or if she would like you to stop the exam at any time
- NEVER say the word “relax”
- Let her know when and where you plan to touch her
- Demonstrate the pressure you will use on another body part
- Significantly important with a history of abuse
- Ask if she does yoga – recommend yoga breaths
 - If not, walk her through some breathing techniques

Q tip test

- Current standard assessment for evaluating for provoked vulvodynia
- 1:00, 3:00, 5:00, 6:00, 7:00, 9:00, 11:00 all tested randomly
- Pain rating scale x/10 at each location
- Make sure your system of mapping out vestibular regions is reproducible
- Wet cotton swab
- Vulvalgesiometer
 - Research purposes
 - Standardized pressures

Overactive pelvic floor

- Holding urine or stool
- Postural abnormalities
- Prolonged lack of motion
- Leg length discrepancies
- Gait abnormalities
- Pelvic girdle abnormalities
- Pregnancy
- Labor and delivery
- Running
- Gynecologic surgery
- Anxiety
- Sexual abuse history



Symptoms of overactive pelvic floor

Vulvar Pain

Vulvodynia

Dyspareunia

Superficial or deep

Vaginismus

Pain with tampon use

Pelvic Pain

Abdominal pain

Hip pain

Low back pain

Coccydynia

Rectal pain/fullness

Vulvovaginal burning

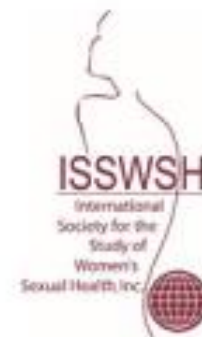
Urinary urgency/frequency

Vestibulodynia: Exam and Lab Findings



- Diffuse vestibular tenderness
- Ostia of glands are frequently erythematous
- Vestibule may have diffuse pallor with superimposed erythema
- Low estradiol, low free testosterone, very high SHBG

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Hormonally Mediated Vestibulodynia

Hormonally-Associated Persistent Vulvar Pain



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- Commonly caused by hormonal contraceptives
- Other causes include:
 - Menopause
 - Oophorectomy
 - Hormonal control of endometriosis or hirsutism
 - Breast-feeding
 - Infertility treatments
 - Treatment of breast cancer

Hormonally-Medicated Vestibulodynia



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- Stop hormonal contraceptives
- Consider topical estradiol 0.03%/testosterone 0.01% in base (e.g., versabase) twice daily to vestibule
- Improvement- none expected at 6 weeks, 50% at 12 weeks
- Consider topical estradiol 0.03%/testosterone 0.01% in base (e.g. methylcellulose) twice daily to vestibule
- Clinical experience suggests estradiol 0.01%/testosterone 0.01% may also be effective.

Congenital Neuroproliferative Vulvodynia



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- Increased density of C-afferent nociceptors in the vestibular mucosa.
- Nociceptors extend into the superficial dermis
- Primary: congenital neuronal hyperplasia in the primitive urogenital endoderm.
 - Umbilical hypersensitivity

Acquired Neuroproliferative Vulvodynia



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- Women report onset of symptoms after severe or recurrent candidiasis or allergic reaction
- Polymorphism in genes coding for IL-1ra, IL-1b
- Decreased INF-a
- Elevated TNF, IL-1b, IL-6, IL-8, Heparinase
- Increased mast cells in mucosa
- Persistent inflammation can lead to a proliferation of C-afferent nociceptors

VIN

- Potential precursor to vulvar cancer
- No routine screening for this
- HPV detected in 86.7% of VIN in a review of > 2,000 specimens
 - HPV 16 most commonly
 - Followed by 33 and 18
- Study in 2000 showed incidence of VIN 3 was 2.86 per 100,000 women
- Low-grade squamous intraepithelial lesion (LSIL) of the vulva (flat condyloma, HPV effect)
 - Not a precancerous lesion – no tx necessary unless symptomatic
- High-grade squamous intraepithelial lesion (HSIL) of the vulva (VIN unusual type)
- VIN differentiated type (dVIN)

VIN

- Average age at diagnosis is 46
- Risk factors:
 - HPV
 - Cigarette smoking
 - Immunodeficiency
- dVIN is associated with vulvar LS
- Typically asymptomatic
- Surgical excision
 - May consider ablative therapy or pharmacologic treatment for some

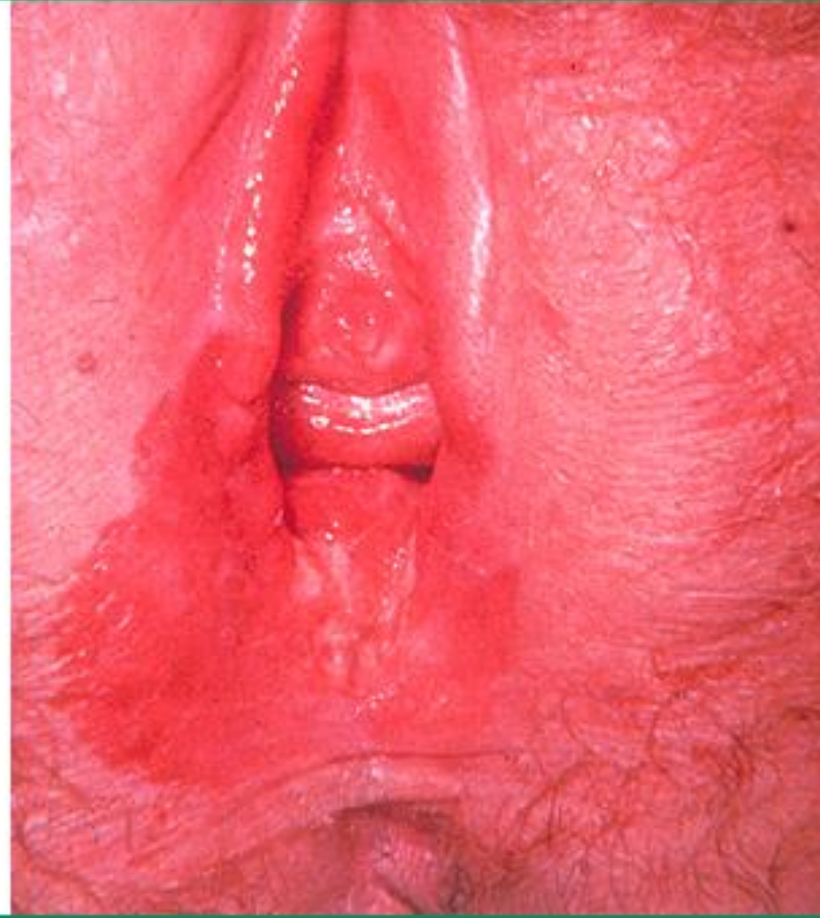
White plaques of vulvar high-grade squamous intraepithelial lesions (HSIL)



Raised whitish plaques as a manifestation of high-grade squamous intraepithelial lesions of the vulva (HSIL).

Courtesy of Christine Holschneider, MD.

Red macular high-grade squamous intraepithelial lesion (HSIL) of vulva



Red macular lesion as a manifestation of a high-grade squamous intraepithelial lesion (HSIL) of the vulva.

Courtesy of Christine Holschneider, MD.

Brown macular high-grade squamous intraepithelial lesion (HSIL) of vulva



High grade squamous intraepithelial lesion (HSIL) manifesting as a brown macular lesion of the vulva.

Courtesy of Christine Holschneider, MD.

Vulvar cancer

- Vulvar cancer is the 4th most common gyn malignancy in the US
 - Uterine, ovarian, and cervical top 3
- Squamous cell carcinoma is most common (75%)
 - Melanoma, basal cell carcinoma, Bartholin gland adenocarcinoma, sarcome, and Paget disease are others
- 6,000 cases and 1,000 deaths from vulvar cancer each year
- 0.3% lifetime risk of vulvar cancer in US
- Average age of diagnosis in US – 68 yrs
- 5 yr survival rate in US – 72.1%

Vulvar cancer risk factors

- Vulvar intraepithelial neoplasia
- Cervical intraepithelial neoplasia
- h/o cervical cancer
- Cigarette smoking
- Vulvar LS
- Immunodeficiency syndromes
- Northern European ancestry
- High risk HPV

Vulvar cancer

- Typical presentation – vulvar lesion
- Typically unifocal vulvar plaque, ulcer, or mass – fleshy, warty, or nodular
- Many are asymptomatic, some present with vulvar pruritus or bleeding
- Dysuria, dyschezia, rectal bleeding, enlarged lymph node in groin, LE edema – all suggestive of advanced disease
- Often associated with a 2nd malignancy – most commonly cervical neoplasia
- Biopsy, vulvoscopy

Vulvar squamous cell cancer and lichen sclerosus



Reproduced with permission from Lynne J Margesson, MD.

Vaginal Intraepithelial Neoplasia (VaIN)

- Squamous cell atypia without invasion
- Carcinoma in situ (considered VaIN) encompasses full thickness epithelium – considered VaIN 3
- Estimated 0.1-0.3 cases per 100,000 women in the US
- Average patient is 43-60 yrs
- HPV common risk factor
 - Also immunosuppression
- Associated with prior or concurrent neoplasia elsewhere in GU tract 50-90% of the time

VaIN diagnosis

Typically asymptomatic

Occasional postcoital spotting or vaginal discharge

Digital palpation to assess for thickening of the vaginal wall

Colposcopy with biopsy if indicated

Majority of lesions upper 1/3 of vagina

2-8% will progress to invasive vaginal carcinoma

Treatment

- Excision, ablation, topical therapy, radiation of lesions

VaIN 1 lesions often regress spontaneously, and no established malignant potential

f/u with cytology every 6 months for 1-2 years, then annually

Vaginal Carcinoma

3% of all malignant neoplasms of the female genital tract

- 4,000 cases and 900 deaths annually in US

Mainly SCC

- Also melanoma, sarcoma, adenocarcinoma, and others

Metastatic spread to vagina is not uncommon

- From endometrium, cervix, vulva, ovary, breast, rectum, and kidney

Mean age at diagnosis 60 yrs

Same risk factors as cervical neoplasia

- Multiple sexual partners
- Early age at coitarche
- Current smoker

Vaginal Carcinoma

- Many are asymptomatic
- Vaginal bleeding is the most common presentation
 - Post coital or post meno
- Diagnosis:
 - History – complete GU
 - Pelvic exam – may note a mass, plaque, or ulcer
 - Vaginal cytology
 - Vaginal biopsy
 - Examine inguinal canal for nodes



We still think of a powerful man as a
born leader and a powerful woman
as an anomaly.

— *Margaret Atwood* —

AZ QUOTES