

#### FEMALE GENITAL SYSTEM PATHOLOGY Lec. 1

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#### **Case presentation**

A 65-year-old woman, without any history of significant medical or family medical history, **presented** with history of vulvar and perianal itching with white patches and vulvar swelling. It started as erythematous macule, expanding gradually, and finally turned out to be a white patch.

Examination revealed a well-demarcated, mild atrophic white patch with a violaceous periphery over vulvar and perianal area, systemic examination was normal.

\* Histopathologic examination of the lesion showed:-

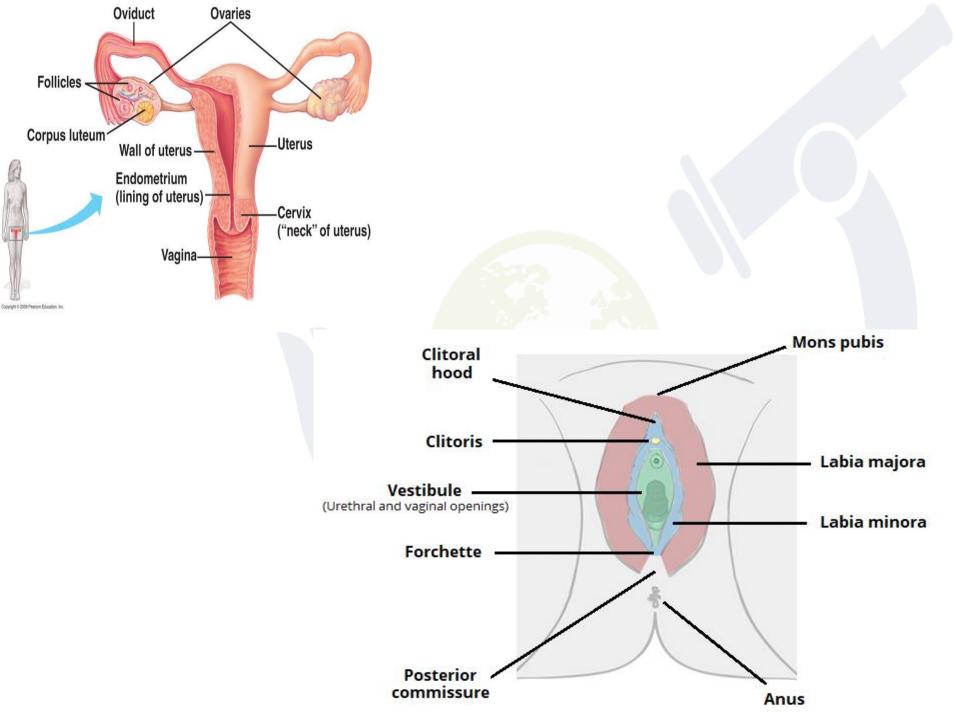
Superficial hyperkeratosis, Epidermal thinning, and dermal fibrosis with

a mononuclear perivascular infiltrate.



## Objectives of this lecture At the end of the lecture

- Anatomy and normal histology of vulva (rapid revision)
- Classification of vulvar pathological disorders
- Non-neoplastic epithelial disorders of vulva
- Intraepithelial neoplasia
- Invasive disease (vulvar cancer)
- Types
- Presentation
- Histopathological finding



#### **Disorders of the Vulva**

#### Classification

- 1. Nonneoplastic epithelial disorders of vulva
- Lichen sclerosus
- Squamous cell hyperplasia (lichen simplex chronicus)
- Other dermatoses (e.g. psoriasis, lichen planus)
- 2. Intraepithelial neoplasia
- Squamous vulvar intraepithelial neoplasia (VIN)
- Non-squamous Glandular intraepithelial neoplasia
  - Extramammary Paget's disease
- 3. Invasive disease (vulval cancer)

#### NON NEOPLASTIC LESIONS

#### 1- LICHEN SCLEROSUS ET ATROPHICUS



Inflammatory dermatosis that mostly affects anogenital area. Lesions begin as papules or macules that eventually coalescent into smooth, white parchment-like areas.

#### Pathogenesis is uncertain:-

- Autoimmunity: autoantibody against ECMP1
- Genetic: high correlation of it has been reported between twins and family members
- Infection: spirochetes(borrelia burgdoferi), or viruses (HPV, Hepatites C)
- Local skin changes: trauma, oxidative stress DNA and Protein damage
- Hormonal: Since LS is primarily found in women with a low **estrogen** state (prepubertal and postmenopausal women). To date though, very little evidence has been found to support this theory

# **Clinical Findings**





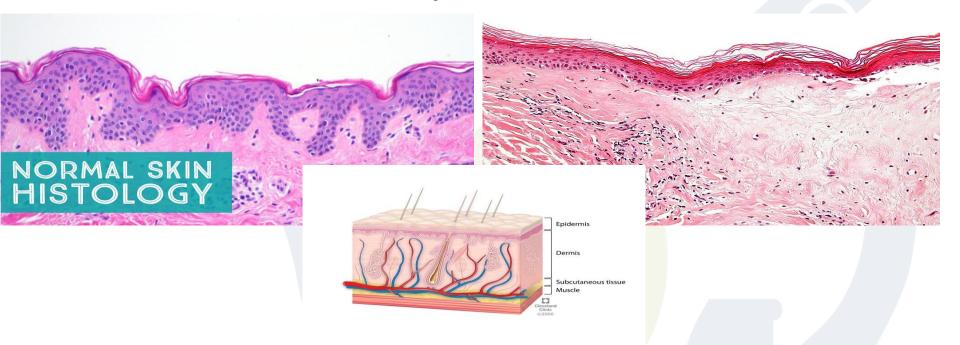
- Age: Occur in all age group but Most common in menopausal women
- Site: It may be appeared elsewhere on the skin
- Symptoms: Commonly present with vulval and perianal itching, Pain may occur, but is usually secondary to skin trauma from scratching, Dysuria as urine comes into contact with the split skin. Narrowing of introitus from the scarring effect causing dyspareunia

#### • Signs:

- Skin is thin, inelastic, white, crinkled "tissue paper"
- Fissures and erosions might occur from local trauma.
- Inflammatory adhesions.

#### Histopathology

 Superficial hyperkeratosis ,Epidermal thinning, and dermal fibrosis with a mononuclear perivascular infiltrate.



**Complications and Cancer risk.** 

Main issues are the <u>scarring complications</u> of the disease.

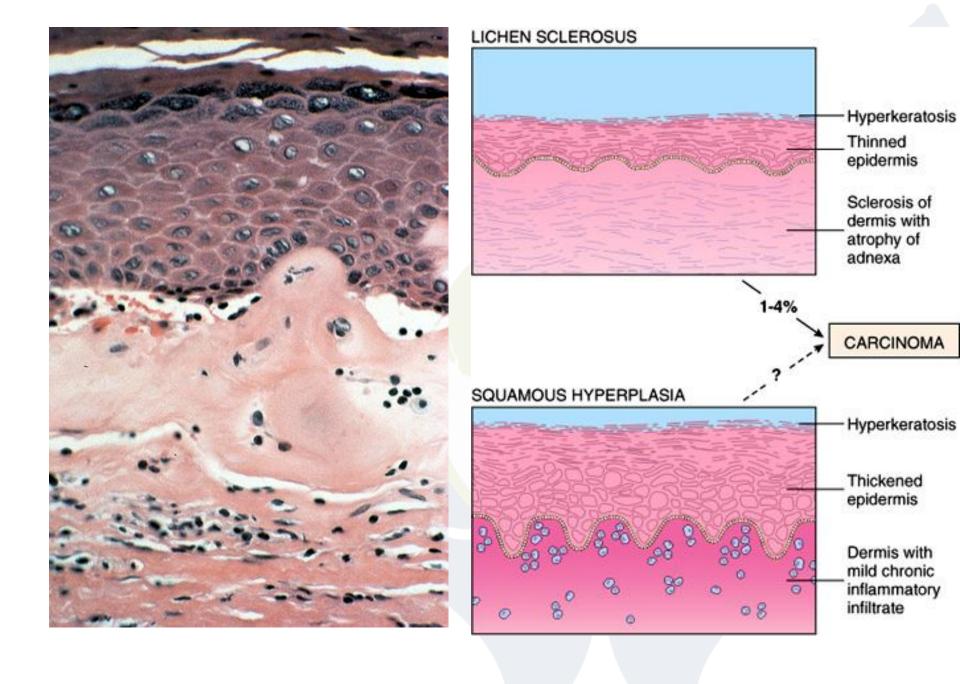
The labia can become atrophic and stiffened, with constriction of the vaginal orifice

 Increased risk of squamous cell cancer of vulva in women with lichen sclerosus (2-4%).

#### NON NEOPLASTIC LESIONS

### 2- Lichen simplex chronicus (Squamous Cell Hyperplasia)

This is a non-specific response to recurrent scratching to relieve pruritus (recurrent trauma) Clinically:- It is characterized by white plaques Histologically:- marked epidermal thickening (d.t increased mitotic activity in basal & pickle cell layer) with significant surface hyperkeratosis. Although lichen simplex chronicus does not exhibit epithelial atypia and there is no increased predisposition to malignancy BUT It is often present at the margins of vulvar carcinoma.



#### **NON NEOPLASTIC LESIONS**

#### 3- Benign Exophytic Lesions:-

#### **Condyloma Acuminatum:**

These are verrucous lesions on the vulva, perineum, vagina, and (rarely) cervix that are sexually transmitted by HPV types 6 or 11. **Condyloma acuminatum is not considered pre-cancerous.** 

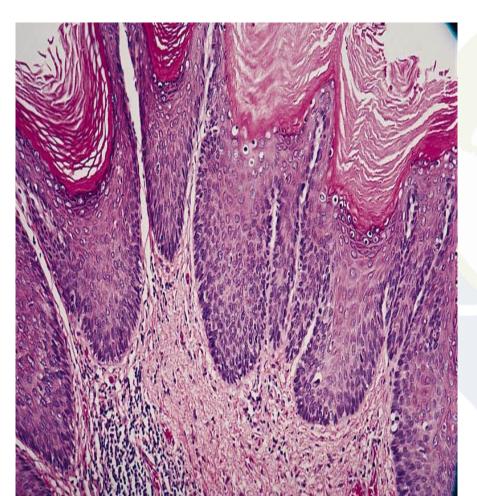
These appear as elevated warty localized red-pink to white often multiple lesions that measure up to several centimeters in diameter.

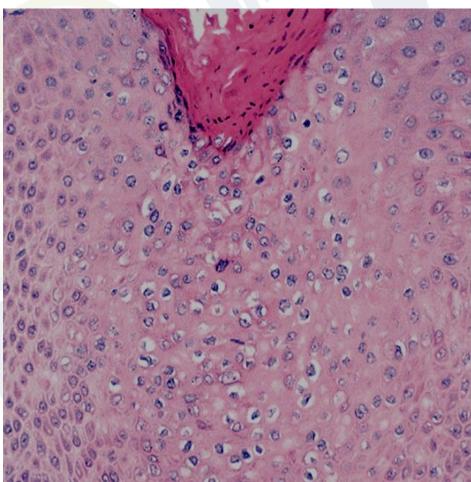


#### **Condyloma acuminatum**

#### They comprise:-

- Branching epithelial proliferations of stratified squamous epithelium;
- Thickening of the epidermis ,hyperkeratosis
- Mature superficial cells exhibit characteristic perinuclear cytoplasmic clearing with nuclear pleomorphism (koilocytotic atypia) and cytoplasmic vacuolation (koilocytosis).





#### **Vulval Intraepithelial Neoplasia**

#### **Squamous Intraepithelial Neoplastic Lesions**

- Dysplasia with varying degrees of atypia that has not invaded the basement membrane but has invasive potential.
- The current classification for VIN :
- Classic (VIN) (previously designated carcinoma in situ or Bowen disease)
  - Multifocal
  - Graded to mild , moderate & sever acc.
     To thickness involved
  - Premenopausal women
  - Most are positive for HPV 16 and are often associated with vaginal and/or cervical HPV-related lesions, smoking and immunodeficiency
  - May have similar pathophysiology to & associated with cervical intraepithelial neoplasia CIN.
  - Associated with developing into the warty and basaloid type carcinoma.

- Differentiated VIN
- Unifocal
- Not graded
- Postmenopausal women
- Associated with lichen sclerosis and usually have a non viral etiology
- Not classically associated with cervical intraepithelial neoplasia CIN.
- It is associated with atypia and then carcinoma of the squamous epithelium.

#### **Clinical Manifestations**

- Approximately 50% of cases asymptomatic.
- Classic VIN lesions manifest as discrete, hyperkeratotic, fleshcolored or pigmented, slightly raised plaques.



- Pruritus
- Dyspareunia
- The cervix and perianal area to be examined to exclude CIN and anal intraepithelial neoplasia.

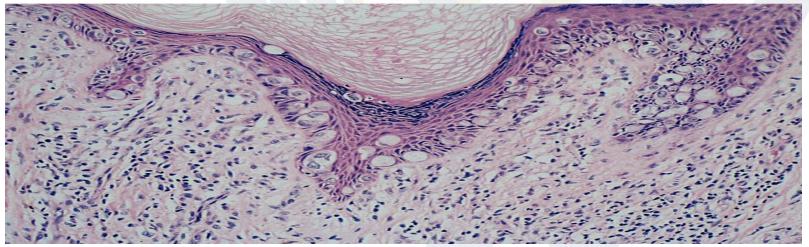
#### **Glandular Intraepithelial Neoplastic Lesions**

#### **Extramammary Paget Disease:-**

• This malignant lesion appears as a red, crusted, sharply demarcated, map-like area, seen in <u>postmenopausal women</u>, Symptom is <u>mainly pruritus</u>, Associated with an <u>underlying</u> adenocarcinoma(gastrointestinal, urinary tracts and the breasts) should be checked.

Paget's disease of vulva in 87-year-old woman. Disease involves labia majora and labia minora. Note red, crusted, sharply demarcated, map-like area





There are scattered large, clear tumor cells within the squamous epithelium

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