

# CLINICAL STOMATOLOGY CONFERENCE

DNSC D9910.00

September 12, 2007

## White lesions

### Overview

- Squamous papilloma
- Smokeless tobacco keratosis
- Leukoplakia
- Squamous cell carcinoma
- Verrucous carcinoma

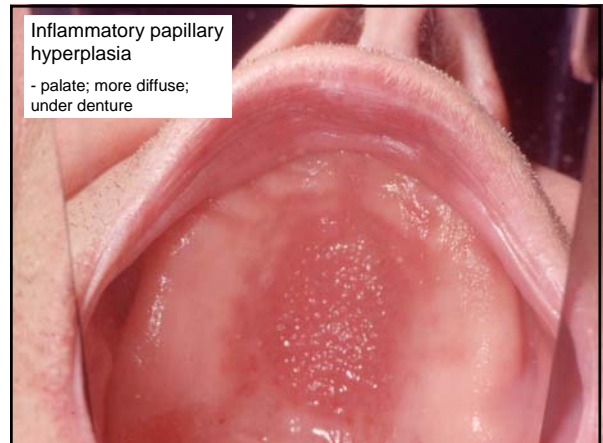
### Squamous papilloma

- Benign proliferation of *squamous cells*
- Etiology: Presumably human papillomavirus (HPV)  
HPV types 6 and 11 have been identified
- Gender: No gender predilection
- Age: Any age
- Site: Any oral site  
Mostly tongue, lips, soft palate
- Clinical features:  
White to pink  
Soft, painless  
Finger-like projections – “cauliflower-like”



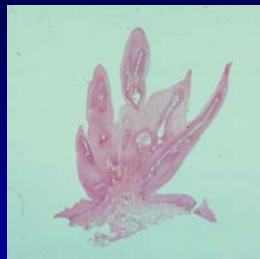
## Squamous papilloma

- Differential diagnosis:
  - 1) Verruca vulgaris (common wart)
    - \* At times, papillomas and warts can appear identical \*
  - 2) Condyloma acuminatum
    - broader base
  - 3) Inflammatory papillary hyperplasia
    - under ill-fitting denture
  - 4) Verrucous carcinoma



## Squamous papilloma

- Histology:
  - keratinized, stratified squamous epithelium
  - papillary configuration
  - fibrovascular connective tissue
  - variable inflammation



- Treatment: Conservative surgical excision

## Smokeless tobacco keratosis

- Etiology:
  - Habit of chewing or holding tobacco in oral cavity
  - allows absorption of nicotine and other carcinogens through oral mucosa
- Age and gender:
  - Older and young males
  - In some populations, females predominate

## Smokeless tobacco keratosis

- Clinical features:

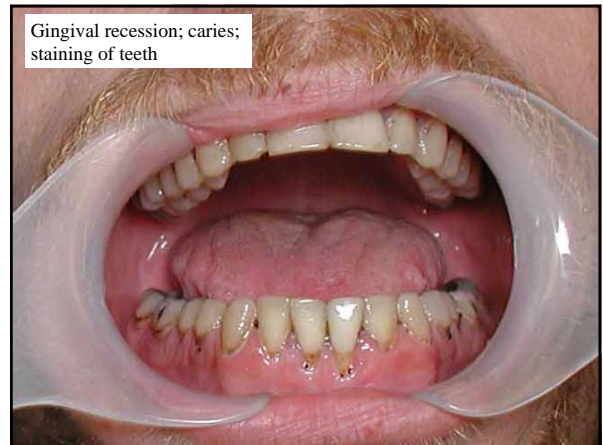
Thin, gray-white plaque

Velvety

Fissured, rippled

Older lesions: leathery or nodular

\* *Gingival recession; dental caries; staining \**



## Smokeless tobacco keratosis

- Differential diagnosis:

1) Frictional hyperkeratosis

Cheek/lip biting

2) Chemical injury / burn

3) True leukoplakia

4) Lichen planus





## Smokeless tobacco keratosis

- Histology:
  - hyperkeratosis
  - parakeratin chevrons
  - thickened spinous layer (acanthosis)
  - amorphous material in connective tissue
  - ± dysplasia, CIS, SCC



## Smokeless tobacco keratosis

- Treatment:  
Depends on histologic diagnosis  
Risk of developing:
  - 1) Squamous cell carcinoma
    - risk 4x greater than non-smokeless tobacco users
    - if combined with *betel quid*, risk 10x greater
  - 2) Verrucous carcinoma

\* 98% of mild smokeless tobacco keratoses will resolve on cessation of habit \*

## Leukoplakia

- Definition:  
“A **white**, plaque-like lesion which cannot be wiped off AND cannot be clinically diagnosed as any other disease entity”
- Clinical term, not related to *histopathology*

## Leukoplakia

- Diagnosis of exclusion  
Must exclude *clinically distinct* entities
  - 1) Frictional (traumatic) keratosis
    - e.g. chronic cheek/lip biting
  - 2) Lichen planus
  - 3) Leukoedema
    - bilateral buccal mucosa; translucent
  - 4) Nicotine stomatitis
    - hard palate; mostly related to pipe smoking

## Leukoplakia

- 5-25% of leukoplakias are diagnosed as **dysplasia** after microscopic examination
- 4% diagnosed as **squamous cell carcinoma (SCC)**



Therefore, true leukoplakias are considered to be **pre-malignant lesions**

## Leukoplakia

- Proposed etiologies:

- Tobacco**

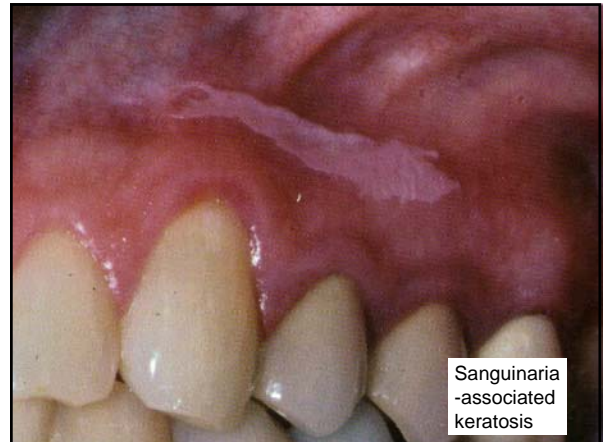
- large proportion of leukoplakias in smokers
- disappear after 1<sup>st</sup> year of smoking cessation

- Smokeless tobacco** (tobacco pouch keratosis)

- Sanguinaria** (sanguinaria-associated keratosis)

- Viadent products
- location: maxillary vestibule or alveolar mucosa
- may not disappear after cessation of sanguinaria use

**\*\*Frictional keratosis** – not true leukoplakias



## Leukoplakia

- Gender:** Male predilection
- Age:** Older than 40 yo
- Site:** Lip vermilion, bu mucosa, gingiva  
Tongue, floor of mouth, lip vermilion  
- *high risk areas for dysplasia*
- Clinical features:**  
Gray to white plaques  
Flat; slightly elevated; fissured
- Clinical variants:**  
Thin; thick; granular/nodular; verrucous;  
proliferative verrucous leukoplakia (PVL)

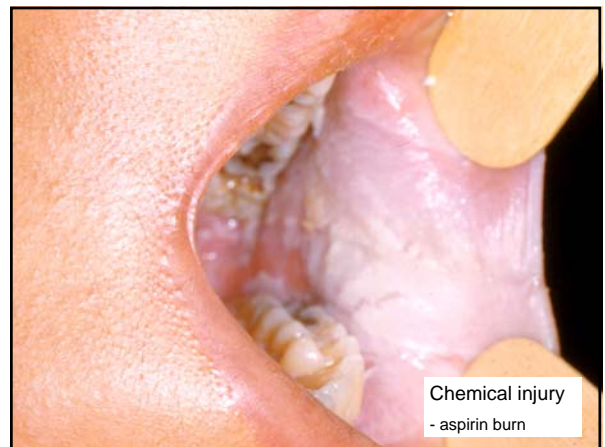




## Leukoplakia

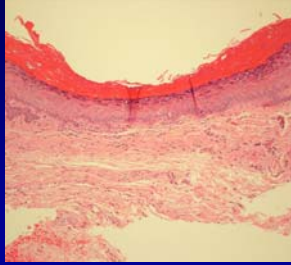
- Differential diagnosis:
  - 1) Frictional (traumatic) hyperkeratosis
  - 2) Linea alba
    - bilateral lines on buccal mucosa; level of occlusal plane
  - 3) Lichen planus
    - look for striations
  - 4) Chemical injuries
    - e.g. aspirin, hydrogen peroxide

Frictional Hyperkeratosis



## Leukoplakia

- Histology:
  - hyperkeratosis
  - acanthosis
  - ± dysplasia; mild, moderate, severe
  - ± carcinoma-in-situ (CIS)
  - ± squamous cell carcinoma



## Leukoplakia

- Transformation to dysplasia rates:
  - Thin leukoplakia – seldom (80% hyperkeratoses)
  - Thick leukoplakia – 1-7%
  - Granular leukoplakia – 4-15%
- Transformation to SCC rates:
  - Moderate dysplasia – 4-11%
  - Severe dysplasia – 20-35%
  - Usually 2-4 years after onset of leukoplakia
  - Risk increased if *persistent lesion, female patient, nonsmoker, FOM or ventral tongue site*

## Leukoplakia

- Treatment:

Depends of histologic diagnosis

  - 1) If *hyperkeratosis or mild dysplasia:*
    - ⇒ Clinical follow-up every 6 months
    - ⇒ Complete removal (surgical, laser, etc)
  - 2) If *moderate dysplasia or worse:*

If high-risk area (*floor of mouth, ventral or lateral tongue*):

    - ⇒ Complete removal (surgical, laser, etc)

Try to preserve specimen for histological exam

## Leukoplakia

Treatment: (cont'd)

**\*\* Careful long-term follow-up \*\***

## Squamous cell carcinoma

- Malignant neoplasm of *squamous cells*
- Incidence:

Oral cancer < 3% of all cancers in U.S.  
94% of oral cancers are SCC  
21,000 new cases diagnosed annually
- Groups at risk:

Caucasian males, >65 yo; *decreasing*  
AA males, middle aged; *increasing*

## Squamous cell carcinoma

- Etiology:

**Multifactorial disease**

Most preceded by *leukoplakia* or *erythroplakia*

  1. **Tobacco smoking**
  2. **Smokeless (spit) tobacco**
  3. **Betel quid (paan)**
  4. **[Alcohol]**
  5. **Iron deficiency**
  6. **Immunosuppression**

## Squamous cell carcinoma

- pipe and cigar smoking, *greater risk*
- more cigarettes, *greater risk*
- reverse smoking, *greater risk*
- **SCC that occurs in non-smokers**
  - female
  - cancer of oral cavity (esp. tongue) vs. pharynx, larynx
  - young age

## Squamous cell carcinoma

- Gender: M:F = 3:1
- Age: Older adults
- Site:
  - Intraoral** – tongue; posterior lateral border, ventral
    - FOM > soft palate > gingiva > bu mucosa > la mucosa > hard palate
  - \* **FOM SCC** – high risk of second primary malignancy
  - Oropharyngeal** – tonsil, posterior soft palate, base of tongue

## Squamous cell carcinoma

- Site:
  - Lip vermillion**
    - *Risk factors* – light-skinned
      - long-term exposure to sunlight
      - outdoor occupation
    - *Site* – lower lip

## Squamous cell carcinoma

- Clinical features:
  - May be painless
  - Varied clinical appearance
  - 1. **Growth pattern**
    - Exophytic, endophytic
  - 2. **Surface characteristics**
    - Intact, ulcerated
    - Granular, papillary
  - 3. **Color**
    - White, red, mixed









### Squamous cell carcinoma

- Radiographic features:  
 May cause destruction of underlying bone  
 Ill-defined radiolucency  
 Can mimic periodontal disease

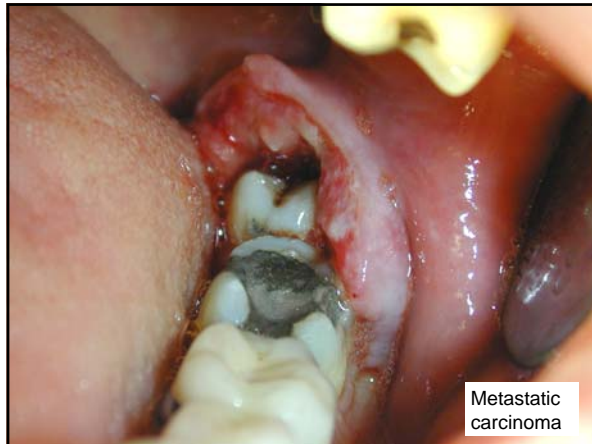
### Squamous cell carcinoma

- Differential diagnosis:

**Intraoral SCC**

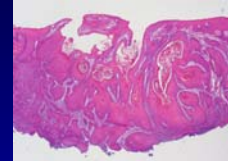
- 1) Leukoplakia/erythroplakia
- 2) Traumatic ulcer
- 3) Ulcer of infectious origin (e.g. TB, syphilis, deep fungal)
- 4) Other malignancies (e.g. primary, metastatic, lymphoma)





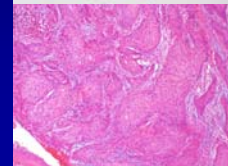
## Squamous cell carcinoma

- Histology:
  - dysplastic epithelium
  - islands of invasive squamous cells



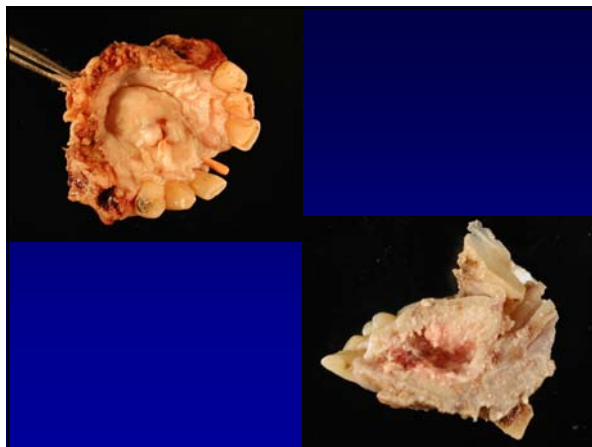
### Grading:

- 1) Well-differentiated
- 2) Moderately differentiated
- 3) Poorly differentiated



## Squamous cell carcinoma

- Treatment:
  - Depends on clinical staging (size, nodal status, distant metastasis)
  - Surgical resection ± radiation therapy ± chemotherapy
  - ± Neck dissection
  - 5-year survival rates – 9-85%
  - Risk for second primary
    - highest risk for smokers and drinkers
    - FOM location
    - usually within 3 years of initial malignancy



## Verrucous carcinoma

- Malignant neoplasm of *squamous cells*
- Low-grade variant of SCC
- Incidence:
  - 1-10% of oral SCCs
- Etiology / Risk factors:
  - Chewing tobacco use
  - Snuff use

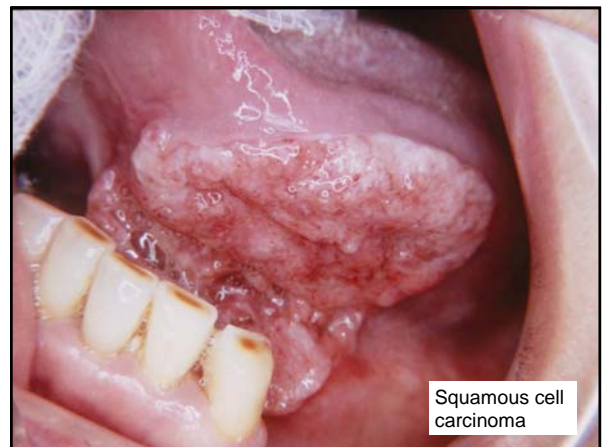
## Verrucous carcinoma

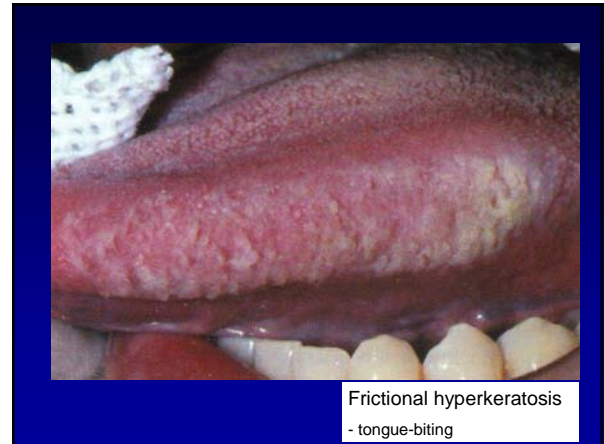
- Gender: M>F
- Age: >55 yo
- Site: Mandibular vestibule, buccal mucosa, hard palate, gingiva  
Site of tobacco placement
- Clinical features:  
White, may be pink  
Diffuse, painless  
Thick mass with papillary projections



## Verrucous carcinoma

- Differential diagnosis:
  - 1) Conventional squamous cell carcinoma
  - 2) Leukoplakia; proliferative verrucous leukoplakia (PVL)
  - 3) Traumatic lesion, factitial (self-induced) injury
  - 4) Smokeless tobacco keratosis
  - 5) Squamous papilloma, verruca vulgaris

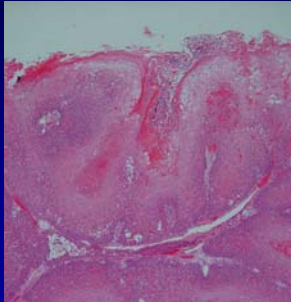




### Verrucous carcinoma

- Histology:
  - papillary surface
  - abundant keratin
  - wide, bulbous rete pegs
  - "pushing" invasion
  - little atypia

\* conventional SCC in 20% \*

A histological slide showing a cross-section of the oral mucosa. The surface is covered by a thick, keratinized layer. The underlying epithelium shows a characteristic "pushing" invasion into the underlying connective tissue, with wide, bulbous rete pegs extending downwards. The overall appearance is that of a well-differentiated squamous cell carcinoma with a verrucous surface.

### Verrucous carcinoma

- Treatment:
  - Surgical resection
  - Radiation therapy less effective