
Differential diagnosis of oral and maxillofacial diseases

19

CHAPTER

ULCERATIONS AND EROSIONS OF THE ORAL MUCOSA

Traumatic etiology

- mechanical (decubital ulcer)
- chemical, electric, thermal damage, burns
- irradiation

Precancerous and neoplastic lesions

- erosive, ulcerative lichen
- discoid lupus erythematosus
- erythroleukoplakia
- carcinoma and other malignant tumors
- multiple myeloma
- non-Hodgkin lymphoma

Recurrent aphthous stomatitis

- minor aphtha
- major aphtha
- herpetiform aphtha

Skin and oral diseases

- oral erosive, ulcerative lichen
- lichenoid reaction
- pemphigus vulgaris
- bullous pemphigoid
- cicatricial pemphigoid
- benign pemphigoid of the oral mucosa
- exudative erythema multiforme
- hereditary epidermolysis bullosa
- Lyell's syndrome (toxic epidermal necrolysis)
- herpetiform dermatitis
- linear IgA disease

Systemic diseases

Diseases of the blood and the hemopoietic organs

- sideropenia (anemia)
- aplastic anemia
- neutropenia
- agranulocytosis

leukemia

Gastroenteric diseases

- Crohn's disease
- coeliac disease
- ulcerative colitis

Immunological and other diseases

- lupus erythematosus
- Reiter's syndrome
- Wegener's granulomatosis
- midline granuloma
- mixed connective tissue disease
- aphthous stomatitis
- Behçet's disease
- Graft versus host disease

Histiocytosis x

- eosinophilic granuloma
- Letterer-Siwe disease
- Hand-Schüller-Christian syndrome

Infectious diseases

- ulcerative gingivostomatitis
- ulcerative necrotizing gangrenous stomatitis
- primary herpetic stomatitis
- labial herpes
- varicella
- herpes zoster
- hand foot and mouth disease
- herpangina
- HIV infection
- infectious mononucleosis
- cytomegalovirus infection
- tuberculosis
- syphilis
- leprosy
- leishmaniasis
- blastomycosis
- sporotrichosis

histoplasmosis
 mucormycosis
 paracoccidioidomycosis

Drugs

cytostatic and other drugs
 indomethacin
 methyldopa
 gold

Other

lingual infarction (ischemic necrosis)
 pyostomatitis vegetans
 sialometaplasia

ACUTE MULTIPLE ULCERS

herpetiform aphthous ulcer
 primary herpes simplex infection
 secondary herpes simplex infection
 herpangina
 hand foot and mouth disease
 herpes zoster
 varicella
 erythema multiforme

WHITE LESIONS

CONGENITAL

pachonychia congenita
 oral mucosal exfoliative leukoedema
 (white sponge nevus)
 congenital dyskeratosis
 focal palmoplantar and oral mucosa hyperkeratosis
 syndrome
 Darier's disease

AQUIRED

Infectious

candidiasis (acute pseudomembranous candidiasis)
 Candida leukoplakia
 chronic mucocutaneous candidiasis
 hairy leukoplakia (AIDS)
 Koplik's spots
 mucosal symptoms of syphilis
 verruca vulgaris
 ulceronecrotic stomatitis
 papilloma

Non-infectious

materia alba
 combustion (e.g. pizza burn)
 stomatitis nicotina palatine
 morsicatio buccarum et labiorum (cheek and lip biting)
 friction keratosis
 smoker's keratosis
 migratory glossitis
 allergic stomatitis (fibrinoid pseudomembrane)
 pyostomatitis vegetans

uremia
 psoriasis vulgaris
 leukoedema
 linea alba

Fox-Fordyce papules (yellow-white)
 xanthoma (yellow-white)

Precancerous lesions, conditions and tumors

keratoses (leukoplakias)
 submucosal fibrosis
 lichen planus
 lupus erythematosus
 epidermoid carcinoma
 verrucous carcinoma (Ackerman-tumor)

Other

drugs and chemicals (aspirin, phenol, trichloroacetic acid,
 silver nitrate, etc.)
 skin transplant in the mouth
 hairy leukoplakia (non-HIV infected)

RED LESIONS

traumatic lesion
 allergic reaction (contact)
 drug-related enanthema
 herpetic stomatitis
 sporotrichosis
 gonococcal infection
 infectious mononucleosis
 irradiation mucositis
 scurvy
 planocellular cancer
 anemia (iron deficiency)
 pernicious anemia
 folic acid deficiency
 neutropenia
 leukemia
 hemorrhagic purpura
 capillary hemangioma
 soft tissue plasmacytoma
 porphyria
 Sjögren's syndrome
 Reiter's syndrome
 acute-chronic atrophic candidiasis
 lichen planus (atrophic)
 erythroplakia
 glossitis rhombica mediana
 pemphigus-pemphigoid
 angular cheilitis

RED AND WHITE LESIONS

migratory erythema
 morsicatio
 morsicatio buccarum et labiorum (cheek and lip biting)
 acute pseudomembranous candidiasis
 lichen planus
 stomatitis nicotina palate
 lupus erythematosus

MOST COMMON IMMUNOLOGICAL DISEASES*Atopic allergic diseases – triggered by foreign antigen*

contact dermatitis and mucositis
eczema, urticaria, asthma
food allergy
drug allergy
angioneurotic edema – anaphylactic shock

Autoimmune diseases (endogen, antigen)

systemic lupus erythematosus
systemic sclerosis (scleroderma)
Sjögren's syndrome
rheumatoid arthritis
polyarteritis nodosa (periarteritis)
dermatomyositis
mixed connective tissue disease

Autoimmune diseases with specific autoantigens

pernicious anemia
hemolytic anemia

Addison's disease
hypothyroidism (Hashimoto's thyroiditis)
hyperthyroidism
idiopathic hypoparathyroidism
idiopathic and drug induced thrombocytopenic purpura
drug associated leukopenia
pemphigus vulgaris
mucosal pemphigoid
aphtha
epidermolysis bullosa acquisita
lichen planus
multiform erythema
Kawasaki's disease (mucocutaneous lymph node syndrome)
herpetiform dermatitis
Behcet's syndrome
reactive arthritis (Reiter's syndrome)
Papillon-Lefevre syndrome
Wegener's granulomatosis
Melkersson-Rosenthal syndrome

DIFFERENTIAL DIAGNOSIS OF COMMON VESICOBULLOUS SKIN AND ORAL DISEASES

	Pemphigus	Benign mucosal pemphigoid	Exudative erythema multiforme	Herpetiform dermatitis	Lichen planus
Age	middle-aged	elderly	young adult	middle-aged	middle-aged
Sex	women	women	man	man	women
Predisposing factors	–	–	drugs, infections, herpes simplex	gluten sensitive enteropathy	drugs
Oral mucosal symptoms	erosions, blisters for a short amount of time, Nikolsky's sign is positive	blisters (sometimes filled with blood), Nikolsky's sign might be positive	swollen lips with yellowish brown serosanguineous exudates, large erosions in the first third of the mouth or rarely blisters, round iris pattern	blisters, ulcers	white lesions, red lesions, erosions, rarely blisters
Skin symptoms	loose blisters	rare, mild		itching vesicles	itching brownish bluish red papulae with shiny surface
Other sites in the body	all over the body	conjunctiva	on flexor side of the forearm, on back of the hands, on palm of the hands, on mucosae (conjunctiva, anogenital region)	on extensor side of the extremities, on the elbow, on the abdomen, on scalp	on flexor side of the wrist, on extensor side of the shin

Vesicobullous diseases

herpetic stomatitis (primary)
labial herpes (recurring)
herpangina
hand foot and mouth disease
herpes zoster
varicella
pemphigus vulgaris
bullous pemphigoid

cicatricial pemphigoid
paraneoplastic pemphigus
epidermolysis bullosa acquisita
linear IgA disease
herpetiform dermatitis
bullous lichen
erythema multiforme
stomatitis venenata (allergic)
stomatitis medicamentosa (drug – systemic)
toxic epidermal necrolysis (Lyell's syndrome)

GINGIVAL SWELLINGS*Drug associated*

Cyclosporine
Phenytoin
Antihypertensive drugs (calcium antagonists)
Nifedipine

OTHER LOCALIZED AND GENERALIZED SWELLINGS

gestational gingivitis
chronic 'hyperplastic' gingivitis
Wegener's granulomatosis
orofacial granulomatosis
sarcoidosis
amyloidosis
scurvy
leukemic infiltration
exostosis (multiple)
gingival cyst (neonate – adult)
eruption cyst
odontogenic cysts

gestational granuloma
fibroepithelial polyp
granuloma fissuratum
verruca vulgaris
Focal epithelial hyperplasia (Heck's disease)
Accumulated condyloma
epulis
carcinoma
non-Hodgkin lymphoma
Burkitt's lymphoma
multiple myeloma
tuberous sclerosis
Cowden's syndrome
congenital gingival fibromatosis
Melkersson-Rosenthal syndrome
Crohn's disease
Paget's disease
fibrous dysplasia
acromegalia
sialolithiasis
mucocele
ranula
exostosis (tori)

LESIONS OF THE ORAL MUCOSA ASSOCIATED WITH HIV-INFECTION

Group 1. Highly associated	Group 2. Rarely associated	Group 3. Probably associated
candidiasis	atypical ulcer	bacterial infections (gingivitis; periodontitis excluded)
acute pseudomembranous candidiasis	diseases of the salivary glands	<i>Actinomyces israelii</i> (actinomycosis)
hyperplastic candidiasis	dry mouth (decreased salivary gland function)	<i>Enterobacter coli</i>
angular cheilitis (infected with <i>Candida albicans</i>)	uni-, or bilateral salivary gland swelling	<i>Klebsiella pneumoniae</i>
hairy leukoplakia	thrombocytopenic purpura	<i>Mycobacterium tuberculosis</i>
HIV gingivitis	virus infection (other than Epstein-Barr viruses)	cat scratch disease
HIV necrotizing gingivitis	cytomegalovirus infection	adverse drug reactions (ulcerative, multiform erythema, lichenoid reaction)
HIV periodontitis	herpes simplex virus infection	exacerbation of apical periodontitis
Kaposi's sarcoma	human papilloma virus (wart-like lesions)	mycotic infections (other than candidiasis)
non-Hodgkin lymphoma	Accumulated condyloma	<i>Cryptococcus neoformans</i>
	focal epithelial hyperplasia	<i>Geotrichum candidum</i>
	verruca vulgaris	<i>Histoplasma capsulatum</i>
	herpes zoster	Mucoraceae (mucormycosis)
	varicella	<i>Aspergillus flavus</i>
		Melanotic hyperpigmentation
		Neurological diseases facial nerve paresis trigeminal nerve neuralgia
		osteomyelitis
		sinusitis
		submandibular cellulitis
		epithelial carcinoma

XEROSTOMIA (DRY MOUTH)*Physiological causes*

- long speech
- mouth breathing

*Lack of salivary glands (agenesia)**Nasal congestion*

- nasopharyngeal tonsil hyperplasia (adenoid)
- deviated nasal septum
- nasal polyps
- malocclusion (Angle II.)
- allergic rhinitis

Age of patient

- Menopause

Fever and respiratory tract diseases

- bronchitis
- bronchial asthma
- pneumonia

Anticholinergic drugs

- atropine and its derivatives
- antidepressant drugs
- antihistamines
- antiemetics
- tranquillizer

Sympatomimetics

- bronchodilators (e.g. Bricanyl)
- appetite reducers (amphetamine)

Other drugs

- lithium
- disopyramide
- cytostatics

Dehydration

- diabetes mellitus (uncontrolled)
- diabetes insipidus
- vomiting, diarrhea
- severe hemorrhage

Psychogenic causes

- distress
- depression
- hypochondria

Diseases and harms affecting organ systems and organs

- sialolithiasis
- Sjögren's syndrome (sicca syndrome)
- sarcoidosis
- irradiation
- HIV infection
- bone marrow transplant
- epidemic parotitis (transient)
- hyperparathyroidism
- uremia
- myelodysplastic syndrome
- iron deficiency
- avitaminosis
- chronic sialoadenitis
- scleroderma (systemic sclerosis)
- submucosal fibrosis
- amyloidosis

- rheumatoid arthritis
- lupus erythematosus
- hemochromatosis

OROPYROSIS (BURNING MOUTH)**GLOSSOPYROSIS (BURNING TONGUE)***Local irritation*

- allergy to denture materials (monomer)
- ill-fitting denture
- smoking
- mouth rinse (corticosteroid)

Infections

- candidiasis
- HIV infection

Deficiencies

- iron deficiency (sideropenia)
- vitamin B₁₂ deficiency (pernicious anemia)
- folic acid deficiency
- vitamin B₂ (riboflavin) deficiency
- vitamin B₃ (nicotinic acid) deficiency
- vitamin B₅ (pantothenic acid) deficiency

Psychogenic causes

- cancerophobia
- distress
- depression
- hypochondria
- vegetative neurosis

Local and systemic diseases

- diabetes mellitus
- gastric illnesses (hyper- or hypoacidity), chronic gastritis
- gastroesophageal reflux disease
- trigeminal neuralgia
- xerostomia
- periodontologic diseases
- hypothyroidism
- angioneurotic edema
- lingua fissurata
- Hunter's glossitis
- mercury poisoning
- antibiotic therapy
- galvanism
- liver and gall bladder diseases
- ulcerative colitis
- arteriosclerotic diseases
- allergy
- lymphogranulomatosis
- chronic pyelonephritis
- menopause

Drugs

- sodium lauryl sulfate
- chlorhexidine
- aspirin
- hydrogen peroxide

HYPERPIGMENTATION

CONGENITAL

racial pigmentation
 ephelis (freckle)
 Peutz-Jeghers syndrome
 xeroderma pigmentosum
 neurofibromatosis
 naevus pigmentosus
 blue naevus

ACQUIRED

Endocrine diseases

Addison's disease
 Cushing's syndrome
 hyperthyroidism

Nutritional factors

scurvy
 pellagra
 vitamin A deficiency

Internal diseases

biliary cirrhosis
 hemochromatosis

*Chronic renal failure**Chronic hepatic disease (chloasma hepaticum)**Hormone-induced*

chloasma gravidarum
 hormonal chloasma
 chloasma climacterium

Physical factors

ionizing radiation
 thermal harm (smoker's melanosis)
 UV light

Drug-induced

antimalarial drugs
 hydantoin derivatives
 contraceptives
 tetracyclines
 sulfonamides
 cytostatics
 Bleomycin
 corticosteroids (corticosteroid-induced chloasma)
 heavy metals, drugs and poisoning
 silver, bismuth, lead, mercury, arsenic

Mucocutaneous diseases

lichen ruber planus
 lupus erythematosus
 nevi
 lentigo maligna
 lentigo maligna melanoma
 malignant melanoma
 scleroderma
 herpetiform dermatitis
 colour changes of exogenous origin, tattoo
 lead
 amalgam
 ink
 carborundum

Others

AIDS
 cosmetics (cosmetic chloasma)

HYPOPIGMENTATION

GENERALIZED MELANOCYTE DEPLETION

*Albinism**Endocrinopathies*

Simmonds' disease
 hypopituitarism

WELL DEFINED FOCAL MELANOCYTE DEPLETIONS

Vitiligo

diabetes mellitus
 pernicious anemia
 Addison's disease
 hyperthyroidism
 hypoparathyroidism
 Hashimoto's thyroiditis
 tuberous sclerosis
 leprosy

HYPER- AND HYPOPIGMENTATIONS

lupus erythematosus
 dermatomyositis
 congenital dyskeratosis
 graft versus host disease

RARE DISEASES OF THE ORAL MUCOSA

Disease	Cause	Symptoms
Pyostomatitis vegetans	Intestinal diseases (ulcerative colitis)	Intraepithelial vesiculation, Abscess formation, Ulcers
Bullous epidermolysis	Autosomal recessive hereditary disease with severe collagen deficiency	Subepithelial bulla formation with scar tissue formation due to mild trauma
Reiter's disease	Reaction conjunctivitis after infection, urethritis, arthritis	Painless zig-zagged white lesions
Gonorrhoea	Neisseria gonorrhoea	Oropharyngeal erythema-ulcers
Leprosy	Mycobacterium leprae	Nodes in the oral cavity, ulcers
Paraneoplastic pemphigus	Lymphoreticular tumors, sarcomas	Vesiculo-bullous lesions, ulcers
IgA pemphigus	Autoimmune	Vesiculo-bullous lesions similar to pemphigus vulgaris
Keratosis follicularis (Darier's disease)	Autosomal dominant genetic disease	Hyperkeratosis pebble-like white spots and intraepithelial bulla formation

MANDIBULAR CYSTS

Odontogenic cysts
gingival
odontogenic keratocyst
eruption (dentigerous)

Inflammatory
radicular

Non-odontogenic cysts
nasopalatine cyst
nasolabial

Cysts without epithelial lining (pseudocyst)
aneurysmal bone cyst
solitary (simple bone cyst)

COMMON ODONTOGENIC TUMORS

Benign epithelial neoplasms
ameloblastoma

Benign connective tissue tumors
odontogenic fibroma
odontogenic myxoma

Hamartomas
odontomas

Malignant neoplasms of dermal origin
odontogenic carcinoma

SWELLINGS OF THE MANDIBULAR BONE (TUMORS)

odontogenic tumors
giant cell lesions
primary, non-odontogenic bone tumors
cysts (mostly odontogenic)
fibroosseous lesions
chronic osteomyelitis
metastatic neoplasms

NON-ODONTOGENIC TUMORS OF THE MANDIBULAR BONE

Benign – primary tumors
hemangioma (central)
central giant cell granuloma
osteoma
melanotic neuroectodermal tumor

Malignant – primary tumors
osteosarcoma
chondrosarcoma
Ewing's sarcoma
multiple myeloma
amyloidosis
Langerhans cell histiocytosis (eosinophilic granuloma, Hand-Schüller-Christian syndrome, Letterer-Siwe disease)
metastatic carcinoma

SWELLINGS OF THE SALIVARY GLANDS

Inflammatory diseases
epidemic parotitis (mumps)
bacterial sialadenitis
recurring parotitis
actinomycosis
Sjögren's syndrome
sarcoidosis
Heerfordt's syndrome (uveoparotitis)
tuberculosis
syphilis
AIDS
gonorrhoea (gonococcal parotitis)
infectious mononucleosis

Tumorous diseases
benign and malignant tumors of the salivary glands
leukemia
lymphomas
Hodgkin's disease

Non-Hodgkin's disease
lymphosarcoma

Diseases of organ systems and organs

cirrhosis (alcoholic)
uremia
diabetes mellitus
hemolytic anemia
acromegalia
myelofibrosis (parotis)

Other

sialolithiasis
Mikulicz's disease
Urbach-Wiethe disease
sialosis

Drugs

phenylbutazone
chlorhexidine
iodine preparation
thiouracil
catecholamines
sulfonamides
phenothiazine
insulin

ENLARGED LYMPH NODES

INFECTIOUS DISEASES

Virus infections

herpetic stomatitis
herpes zoster
cytomegalovirus infection
infectious mononucleosis
HIV infection
rubeola (measles)
hand, foot and mouth disease
rubella (German measles)

Bacterial infections

tooth originating tonsil, nasal, sinus, face, scalp infections
tuberculosis and atypical *Mycobacterium* infections
syphilis
gonorrhoea
tularemia

diphtheria
cat scratch disease
Lyme disease
actinomycosis
leprosy

Mycotic infections

histoplasmosis
blastomycosis
sporotrichosis
other systemic mycoses

Parasite infections

toxoplasmosis

DRUGS

phenytoin
penicillin (rare)
non-steroid anti-inflammatory drugs
antimalarial drugs
phenylbutazone

TUMORS

Primary:

Hodgkin and non-Hodgkin lymphoma
plasmocytoma
leukemia

Secondary:

carcinoma (oral, skin, salivary gland)
mesenchymal tumors (sarcomas)
Burkitt's lymphoma
Sezary's syndrome
malignant melanoma

OTHER

sarcoidosis
immune deficient diseases
connective tissue diseases
histiocytosis
hyperthyroidism
agranulocytosis
aphthous stomatitis
exudative erythema multiforme
Waldenström macroglobulinemia

LESIONS PRONE TO MALIGNANT TRANSFORMATION

Lesion	Etiology	Risk of malignant transformation
Leukoplakia (homogenous)	Not known	Low – it might recess
Leukoplakia (non-homogenous)	Not known	High
Erythroplakia	Idiopathic – tobacco, alcohol, etc.	Very high
Smoker's keratosis (pipe)	Heavy smoking, pipe	Low
Oral submucous fibrosis	Mawa-betel	High
Tertiary syphilis	<i>Treponema pallidum</i>	High
Congenital dyskeratosis	Genetic	High
Lichen planus	Idiopathic (diabetes, psychological instability, hypertension, etc.)	Low
Discoid lupus erythematosus	Autoimmune	Low for lips and skin
Chronic candidiasis	<i>Candida albicans</i>	Low
Tobacco keratosis	Smokeless tobacco	Low

**PARANEOPLASTIC DISEASES (SYNDROMES)
AND SYMPTOMS**

herpes zoster
pemphigus vulgaris (pemphigoid)
Cowden's syndrome
Peutz-Jeghers syndrome
AIDS
Keratoacanthoma
hereditary benign telangiectasia
prurigo
vitiligo
palmoplantar erythema
seborrheic verruca
palmoplantar hyperkeratosis (keratoderma)

erythema nodosum
acanthosis nigricans
eruptive angiomatosis
erythroderma
hypertrichosis
rhinophyma
vasculitis
urticaria
Sweet's syndrome
congenital dyskeratosis
celiac disease
ulcerative dermatitis (pyoderma gangrenosum)
glucagonoma syndrome
sebocystomatosis (Gardner's syndrome)

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