



# Objectives of the course

- Microscopic structure of the organs of the orofacial system
- Connections of structure and function
- Detailed understanding of developmental processes
- Understanding the background of congenital malformations

LecturesWednesday 8:30 - 10:10PracticalsWednesday 8:30 - 10:10

Invited lecture: Úterý 21. 5. 2024, 12:30 – 13:45, **B11/206** prof. Anamaria Balic

#### **Teachers:**

doc. Mgr. Jan Křivánek, Ph.D.

Marina Štruncová, DDS, Ph.D.

#### **Invited lecture**



prof. Anamaria Balic, MD, Ph.D.

University of Zurich<sup>UZH</sup>

**Center for Dental Medicine** 

Tuesday 21. 5. 2024, 12:30 – 13:45, **B11/206** 

**Mandatory participation = Instead of practices** 

#### **Research Interests:**

- Dental stem cells and stem cell niches
- Tooth development
- Tooth agenesis
- Tooth regeneration
- Craniofacial development

## Conditions to successfully pass the course

**Practicals:** 100% attendance

Successfully completed credit test (8. 5. 2024)

Individually completed all ROPOTS

Attendance at the Invited Lecture on 21 May 2024



Written test (minimally 60 % of correct answers)

30 questions +1 bonus. Multiple choice + short written answers (upgraded tests)

Successful completion of Histology I + II is no longer a prerequisite for admission to the OHE examination

During lectures there may be opportunities to earn bonus points in the final test

The exam may include **questions from presentations in practicals and lectures** (written and **orally communicated** information), from **ROPOTS** and from **discussions** during practicals and lectures.

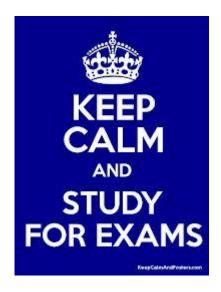
For a more detailed understanding of the presented information, the study of comprehensive literature is recommended:

Ten Cate's Oral Histology: Development, Structure, and Function. Antonio Nanci

Essentials of Oral Histology and Embryology: A clinical Approach

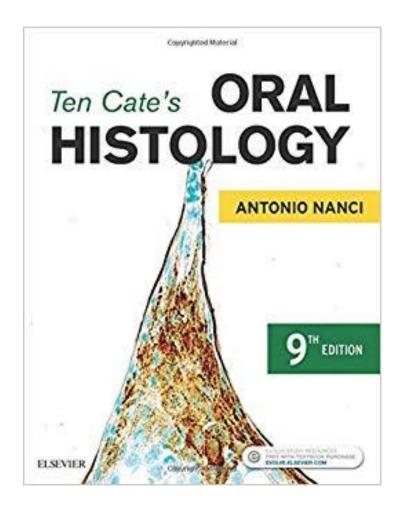
Illustrated Dental Embryology, Histology and Anatomy, Fehrenbach and Popowics

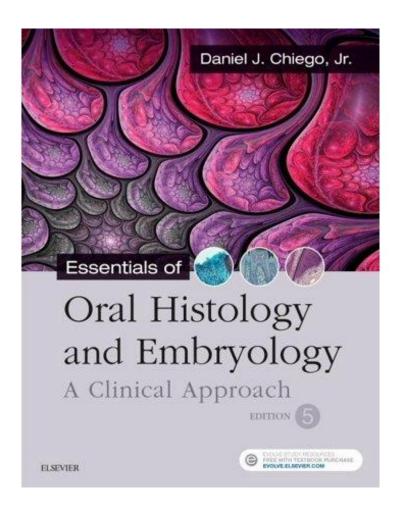
Oral Anatomy, histology and Embryology, Berkovitz, Holland, Moxham



#### Ten Cate's Oral Histology: Development, Structure, and Function.

#### **Antonio Nanci**





**Essentials of Oral Histology and Embryology: A Clinical Approach**Daniel J. Chiego



## ROPOTS

- Every 14 days, one ROPOT
- The ROPOT will be published in the "lecture" week
- It has to be completed by the end of the week in which practicals are held
- Each answer sheet consists of about 10-15 questions to be answered <u>in your own words</u>
- The answer sheets should enable to practice the knowledge acquired
- Questions from the ROPOTS may appear on the exam

#### Semester 4, Spring 2023

## embryology (aZLOH) for the 2nd year of Dentistry Mgr. Jan Křivánek, Ph.D.

doc. Mgr. Jan Křivánek, Ph.D. doc. RNDr. Petr Vaňhara, Ph.D. Marina Štruncová, DDS, Ph.D.

Timetable of lessons

Lectures (even weeks)	Practice (odd weeks)
1. 19. 2. – 23. 2. 2024	1. 19. 2. – 23. 2. 2024
Introduction, information about the completion of the course, recommended literature.	
Orofacial system, its structural components, and functions. Oral cavity - walls and contents. Structure and functions of the oral mucosa, types of mucosae. Taste buds.	
2. 26. 2. – 1. 3. 2024	2. <b>26. 2. – 1. 3. 2024</b>
	Microscopic structure and functional histology: lips, palate, cheeks, tongue.
	Samples: labium oris, palatum molle, apex linguae, papilla vallata, radix linguae.
3. 4.3. – 8.3.2024	3. 4.3. – 8.3.2024
Salivary glands, TMJ	
Microstructure and classification of salivary glands. Temporomandibular joint, microstructure and function.	
4. 11. 3. – 15. 3. 2024	4. 11. 3. – 15. 3. 2024
	Salivary glands, TMJ - microstructure.
	Samples: gl. parotis, gl. submandibularis, gl. sublingualis, gl. apicis linguae, TMJ.
5. 18. 3. – 22. 3. 2024	5. 18. 3. – 22. 3. 2024
Alveolar process, Periodontium	
Microstructure of the alveolar process and clinical aspects of its remodelling.  Microstructure of the periodontium, its function and clinical aspects. Gingiva, sulcus gingivalis.	
6. 25. 3. – 29. 3. 2024	6. 25. 3. – 29. 3. 2024
	Tonsils, Introduction to the tooth
	Samples: Tonsilla palatina, tonsilla lingualis.
7. 1.4. – 5.4.2024	7. 1.4. – 5.4.2024
Enamel, Cementum	
Enamel microstructure, function, amelogenesis and age-related changes. Microstructure of cementum, types and its clinical significance.	

Programme of lectures and practicals in Oral histology and

8. 8.4. – 12.4.2024	8. 8. 4. – 12. 4. 2024
	Dentin-pulp complex
	Dentin as living tissue. Microstructure of the dental pulp, functions.
	Samples: Tooth (ground section).
9. 15. 4. – 19. 4. 2024	9. 15. 4. – 19. 4. 2024
Development of the face, oral and nasal cavities	
Development of the face, oral and nasal cavities, palate, nasal septum, atrium of the oral cavity, upper and lower jaws.	
10. 22. 4. – 26. 4. 2024	10. 22. 4. – 26. 4. 2024
	Tooth development
	Samples: Different stages of tooth development - pig, human.
11. 29. 4. – 3. 5. 2024	11. 29. 4. – 3. 5. 2024
Development of the tongue, salivary glands, pharyngeal arches	
Tongue development, defects. Development of salivary glands. Development and features of pharyngeal arches and their derivatives.	
12. 6. 5. – 10. 5. 2024	12. 6. 5. – 10. 5. 2024
	Science and research, regenerative dental medicine
	Current focus of dental research, advances in the field of regenerative dentistry. Are we going to be able to repair or regenerate our teeth?
	Discussion.
	Credit test (the course will be credited only after attendance at the invited lecture)
13. <b>13.</b> 5. – <b>17.</b> 5. 2024	13. <b>13.</b> 5. – 17. 5. 2024
Permanent dentition, defects	
Development of permanent dentition and a time overview. Mixed dentition. Differences in the structure of primary and secondary teeth. Developmental defects of teeth.	
14. 20. 5. – 24. 5. 2024	14. 20. 5. – 24. 5. 2024
	INVITED LECTURE (COMPULSORY)

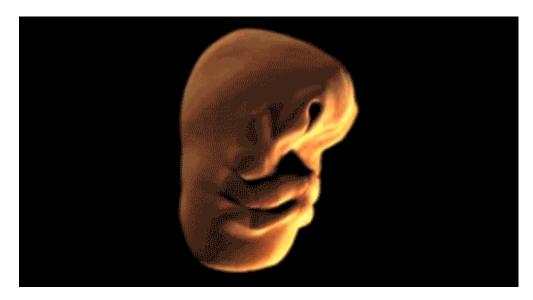
Doc. MVDr. Aleš **Hampl**, CSc. Head of Department

# Orofacial system

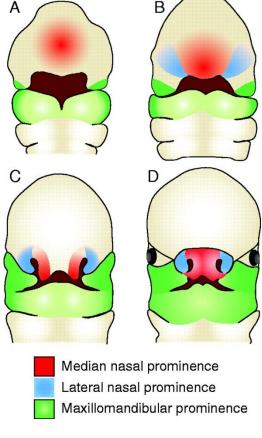
# Orofacial system

#### Structures of the head and neck which:

- Are essential for intake, grinding and processing of food
- Maintain taste and tactile sensations
- Forms an interface for social interactions (phonetic, aesthetic-physiognomic function, mimics, speak)



Development from pharyngeal arches, frontonasal prominence and maxillary and mandibular prominences

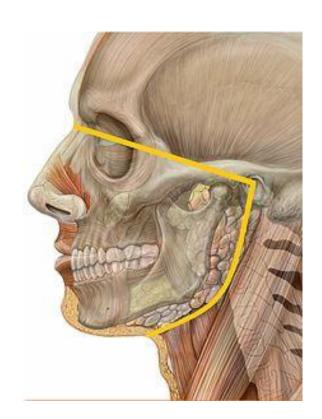


(Helms et al. 2005)

# Orofacial system

#### Orofacial system is composed of:

- **Skeleton faciei** (facial skeleton) mandible, maxilla, ossa zygomatica, os ethmoides, ossa nasalia et lacrimalia, vomer, ossa palatina, os hyoides) + art. temporomandibularis)
- Cavitas oris lingua (tongue), dentes, periodontium, salivary glands (glandulae salivariae)
- Art. temporomandibularis
- Mimic muscles and muscles of mastication
- **Soft tissues of the face** lips, cheeks
- **Hard and soft palate** (palatum durum a palatum molle)
- Isthmus of the fauces (isthmus faucium)
- Palatinal and tongue tonsils



# Oral cavity (cavitas oris)

Basic anatomy

- Oral mucosa and microscopic structure
  - Lining mucosa
  - Masticatory mucosa
  - Specialized mucosa
- Lips
- Microscopic structure of tongue
- Taste buds

# Oral cavity (cavitas oris)

#### vestibulum oris / cavitas oris propria

#### **Borders**

Lips, cheeks, hard and soft palates, caudally floor of cavity, faucial isthmus (connection to oropharynx)

#### Inside

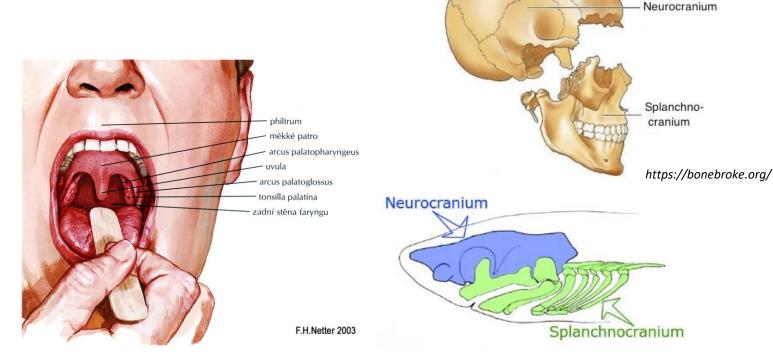
Tongue, teeth, gums, tonsilla palatina

#### Major salivary glands:

gl. submandibulatis

gl. sublingualis

gl. parotis (positioned outside)



## Oral mucosa

Except of teeth it covers all surfaces inside the oral cavity

Oral mucosa has 2 layers (epithelium + *lamina propria mucosae*)

At some places is between mucosa and the base (bone/muscles) located connective tissue - *tela submucosa* 

#### Functions of oral mucosa:

- Protective resistant to mechanical and chemical forces or effects of the bacterial flora
- **Secretory** saliva a product of small and large salivary glands
- **Sensory** contains receptors for perception of temperature, pain, touch and taste
- **Thermoregulatory** in animals (protruding tongue)
- Food processing

# B Early tailbud Mouth formation Teeth development vertebrate scheme B ECT Stomodeum ECT teeth A scheme of axolot! development Stomodeal collar Stomodeal collar Stomodeal collar CSoukup et al., Nature 2005)

#### Features of the oral mucosa:

- Forms special transitory zone inserted between the skin and the mucosa of the alimentary canal (starts in the pharynx)
- The oral mucosa differs from mucosa of the alimentary canal or mucosa other tubular organs by the origin it was developed from the ectoderm and head mesenchyme of ectodermal origin (ectomesenchyme neural crest), while elsewhere from the entoderm or mesoderm and mesenchyme of mesodermal origin.
- Thanks to these circumstances the oral mucosa shows some characteristics of the skin: keratinization of the epithelium, presence of lamina propria protrusions against the epithelium (papillae)

### Oral mucosa regeneration

Oral epithelium turnover time: 4 - 24 days

Significant local differences

Gingivo-dental <u>junctional epithelium</u> 4-6 days <u>Gingiva affixa</u> epith. (masticatory mucosa) 10 days

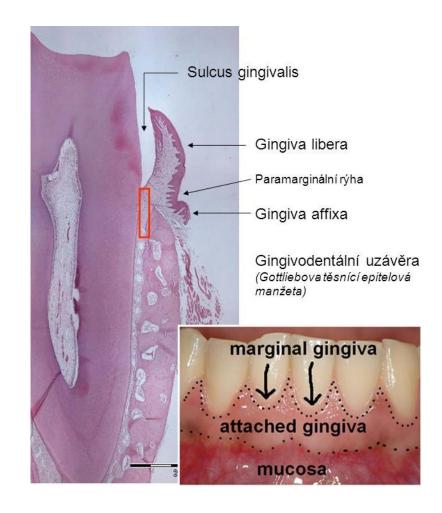
<u>Taste buds</u> 10 - 14 days

Lining epith. of <u>lips and cheeks</u> mucosa 14 days

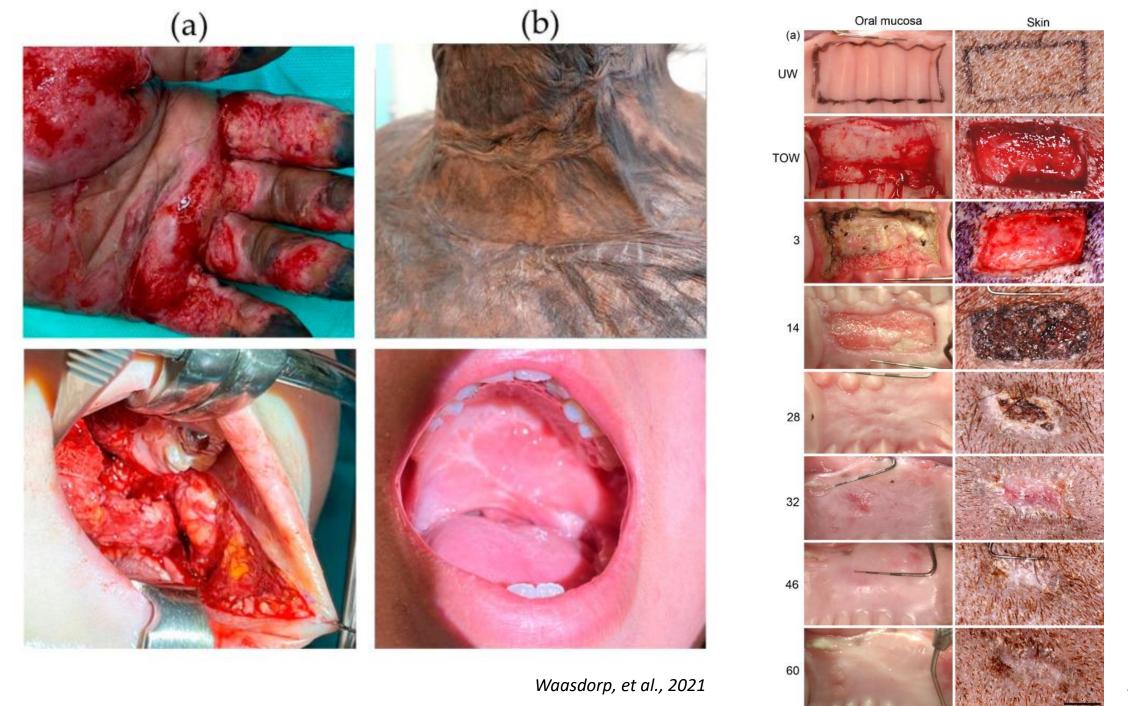
Lining epith. of the <u>floor of mouth</u> 20 days

Masticatory epithl. of <u>hard palate</u> 24 days

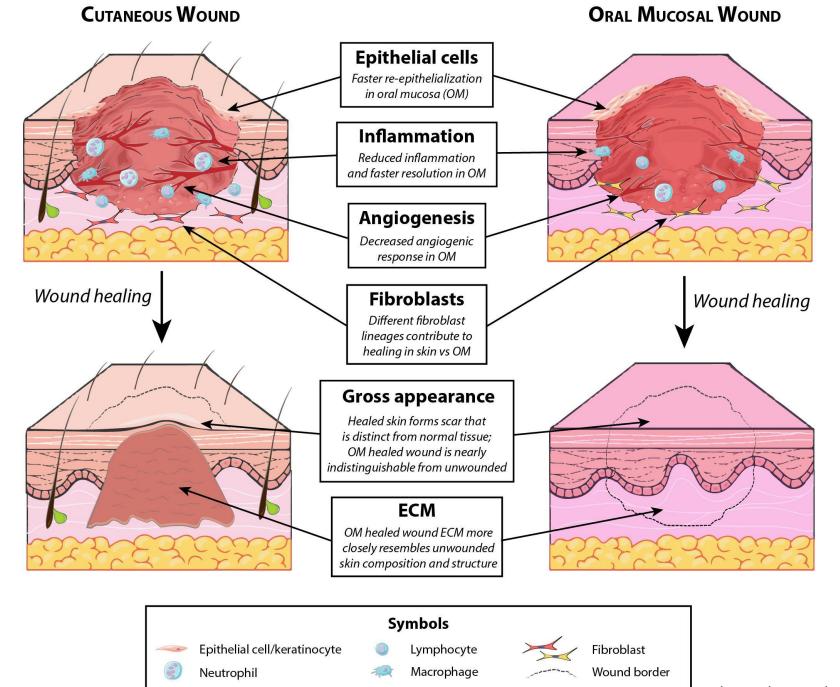
Epidermis of the face and neck frontal side 7 days Epidermis (rest) 30 days



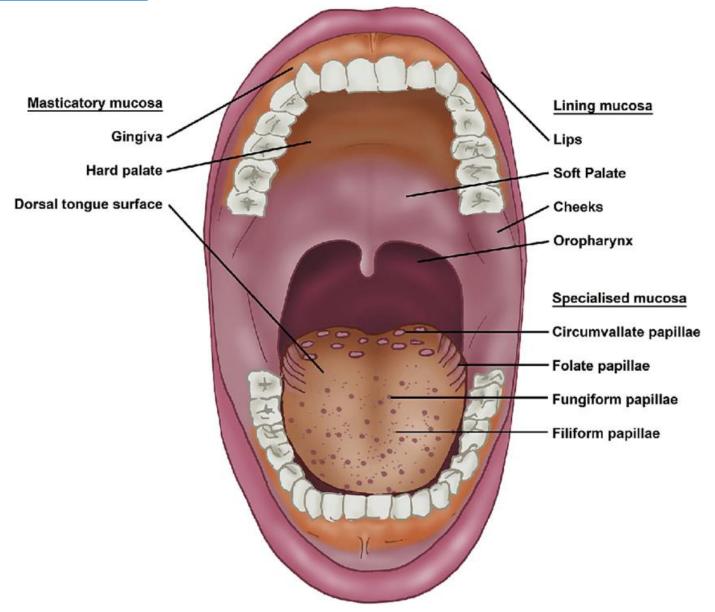
(faster turnover time in case of the face is probably caused by inductive effect of the ectomezenchyme)



Mak, et al., 2009



## Oral mucosa



Cook, Sarah et al. A food perspective. Food Hydrocolloids. 2017.

## Classification of oral mucosa

#### **Lining** (65 %)

Inner part of lips, cheeks soft palate, inferior aspect of the tongue, floor of the mouth and alveolar process (except of the gingiva)

Tela submucosa located under mucosa.

Soft and slightly movable (submucous coat)

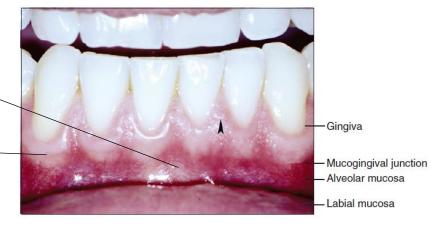
Lamina propria from loose connective tissue

#### Masticatory (25 %)

Hard palate and gingiva

Keratinized epithelium

Tela submucosa is usually missing



Lamina propria is composed from dense collagenous of irregular type and firmly connected with periosteum (mucoperiosteum)

#### Specialized (10 %)

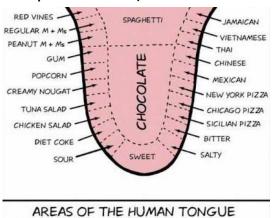
Dorsal surface of the tongue

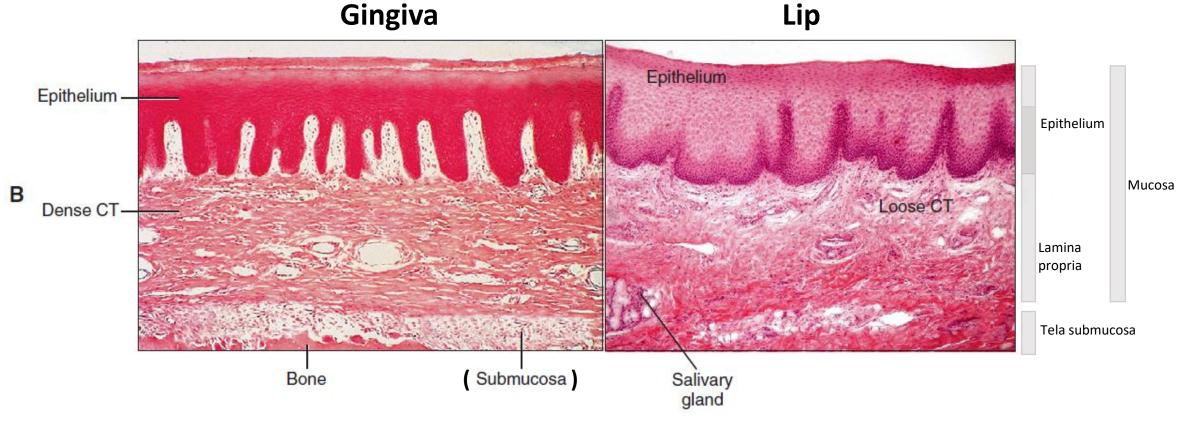
Mucosa protrudes into papillae

Tela submucosa is missing

Lamina propria connected with aponeurosis linguae







#### **Masticatory mucosa**

- Lamina propria from dense collagenous connective tissue of irregular type
- Firmly connected to periosteum (mucoperiosteum)

#### **Lining mucosa**

- Lamina propria from loose collagenous tissue
- Tela submucosa under mucosa
- Mucosa is slightly movable

**B,** In histologic sections, the **gingival** epithelium is seen to be tightly bound to bone by a dense fibrous connective tissue (CT), whereas the epithelium of the **lip (C)** is supported by a much looser connective tissue.

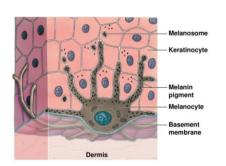
## Oral mucosa

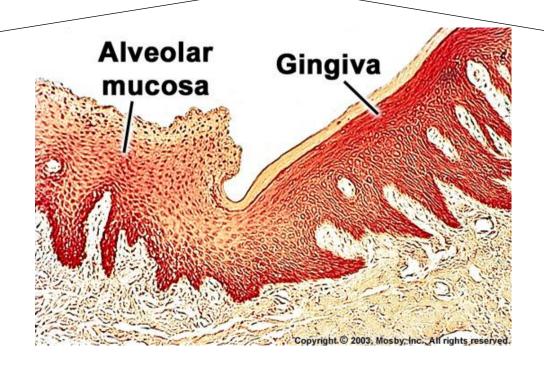
epithelium stratified squamous

Lamina epithelialis: tlustý vrstevnatý dlaždicový epitel

#### nonkeratinized

Lining mucosa





#### keratinized

- Masticatory mucusa
- Specialized mucosa\*



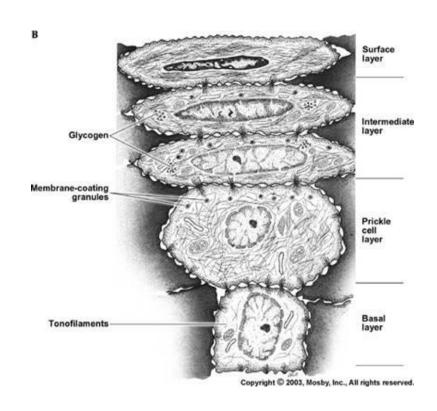
(Yadav et al., 2012)

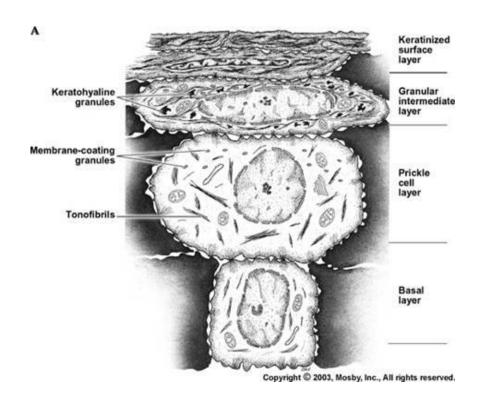
#### Lamina propria mucosae

- Contains numerous of melanocytes or melanophages; Merkel cells
- Multiple papilae projected against the epithelium. Their shape and density are spatially different
- (depends on different mechanical needs of oral mucosa)
- Differences between: Melanophages, melanocytes, (melanophores), melanosomes a melanin

Squid skin
https://youtu.be/OwtLrllKvJE?t=12

#### Classification of cell layers in the epithelium - similar as in the epidermis





#### **Nonkeratinized**

Stratum basale Stratum spinosum Stratum intermedium Stratum superficiale - melanin

#### Keratinized

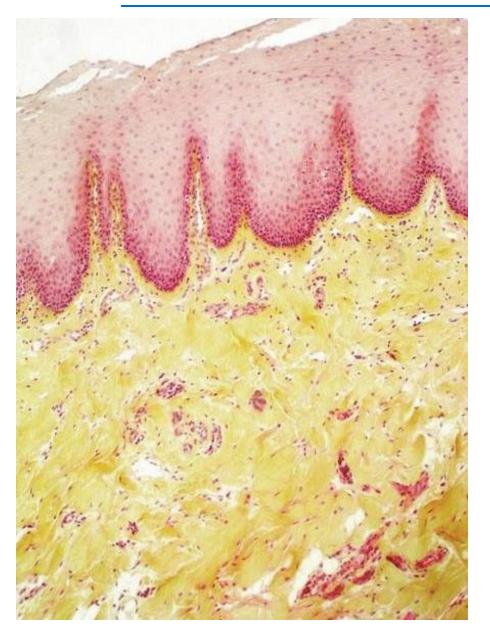
Stratum basale - melanin

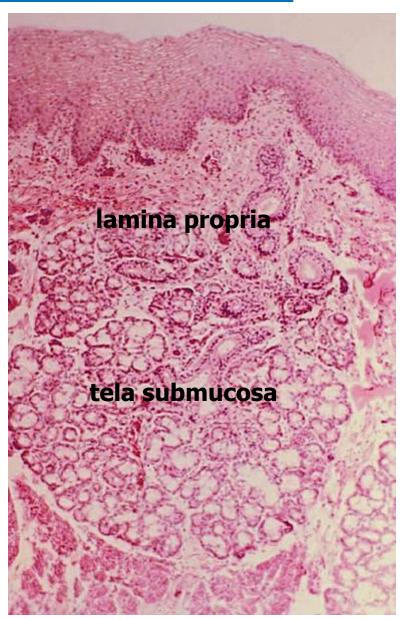
Stratum spinosum

Stratum granulosum - keratohyalin

Stratum corneum - keratin

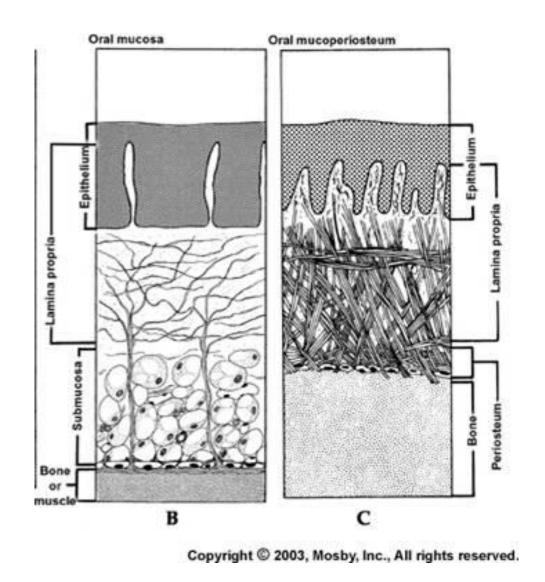
# Lining mucosa

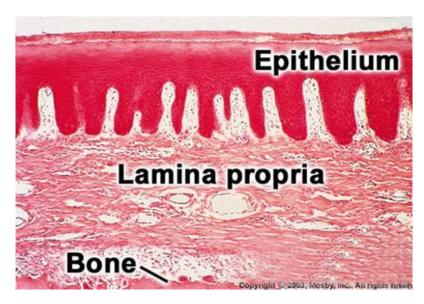


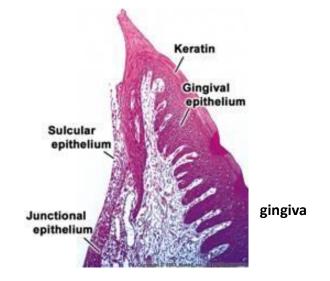


# Masticatory mucosa

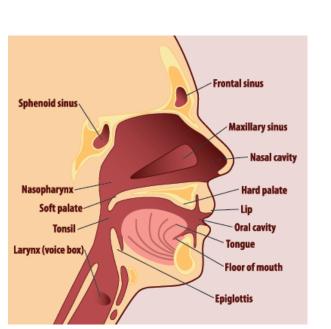
#### mucoperiosteum

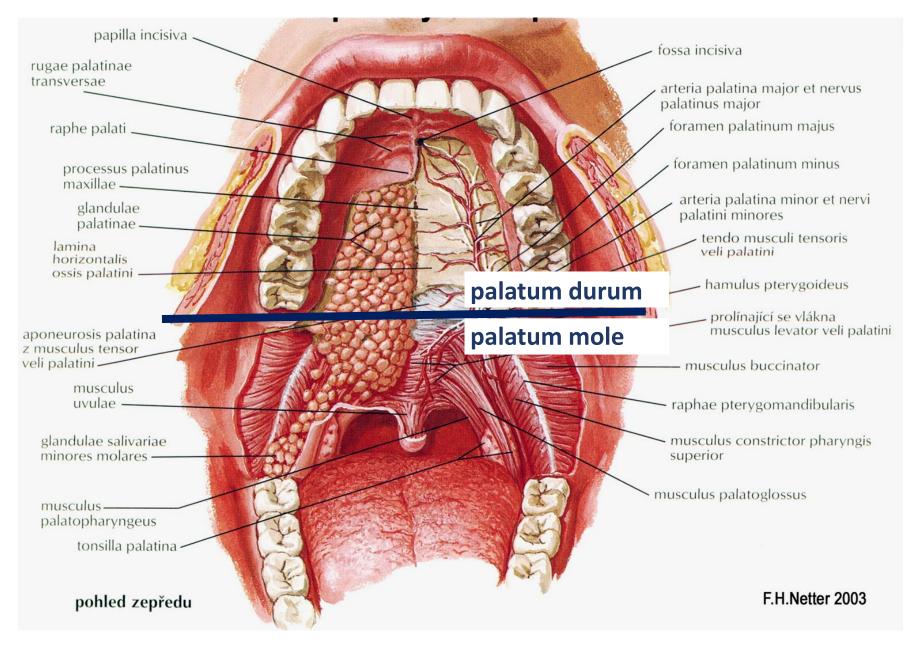






## Palate





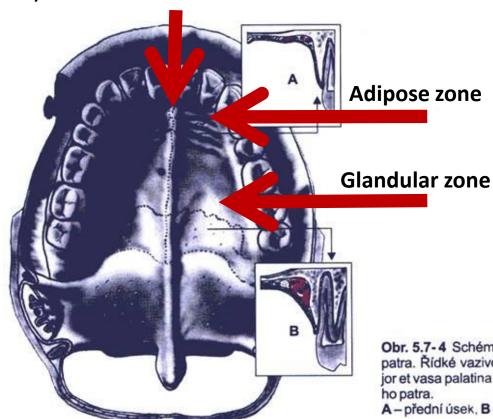
# Hard palate (palatum durum)

#### Masticatory mucosa:

- Epithelium stratified squamous keratinizing
- Tela submucosa is usually missing

High regional variability:

raphe palati (in the middle line)





Obr. 5.7-4 Schéma uspořádání měkkých tkání tvrdého patra. Řídké vazivo (růžově) obsahuje n. palatinus major et vasa palatina majora. Znázorněn průběh švů tvrdé-

A-přední úsek, B-zadní úsek

# Local differences in hard palate structure

#### Raphe palati

- Midline area from papilla incisiva to soft palate, mucosa of raphe palati is without glands and adipocytes
- Formed by fusion of the maxillary processes (origin of clefts)

#### Foramen incisivum

- Location on the papilla incisiva
- In the fetal period, forms opening between the nasal and oral cavities
- Before or shortly after birth, the connection is closed

# Anterior palatine branch of the nasopalatine nerve Greater palatine foramen Lesser palatine foramen

#### Adipose zone

- Paired structure
- Medially divided by papilla incisiva and raphe palati, Laterally bordered by gingiva and premolars
- Mucosa is thickened into 3-5 transversal plicae *plicae palatinae transversae*, core of plicae is formed by stripes of dense colagenous connective tissue interlaced with adipocytes

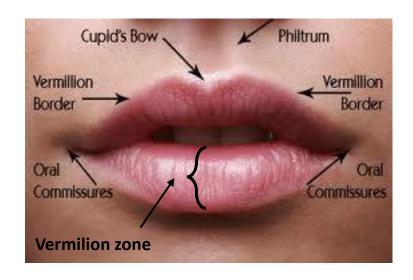
#### Glandular zone

- Paired structure
- Mucosa is smooth and contains <u>mucous</u> glands *gll. palatinae*



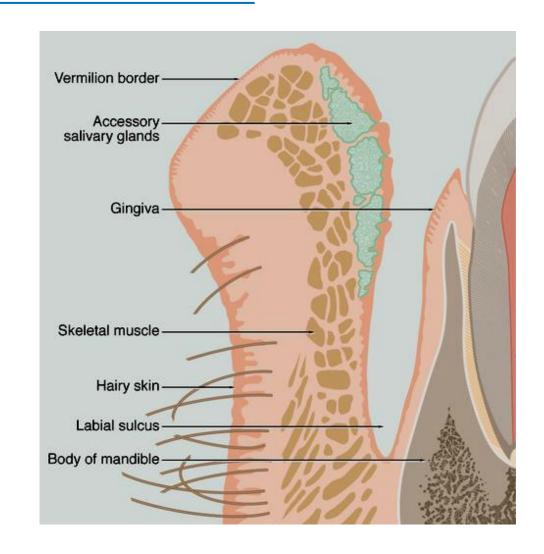
Hard palate – glandular zone

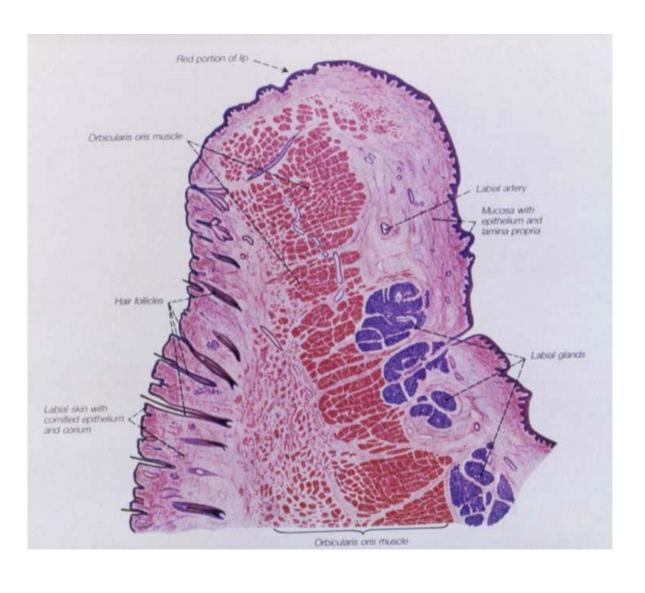
# Lips



#### Sagitally:

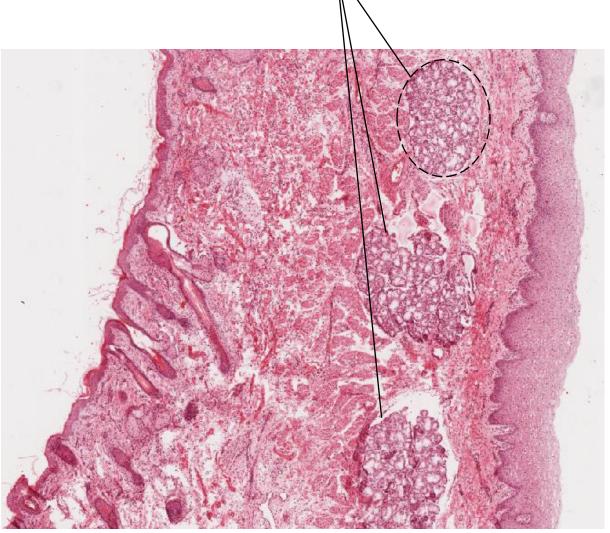
- ventral aspect of the lip (skin)
- dorsal aspect of the lip (mucosa)
- Structural support: *m. orbicularis oris*
- Vermilion zone





## glandulae labiales

(mixed glands)

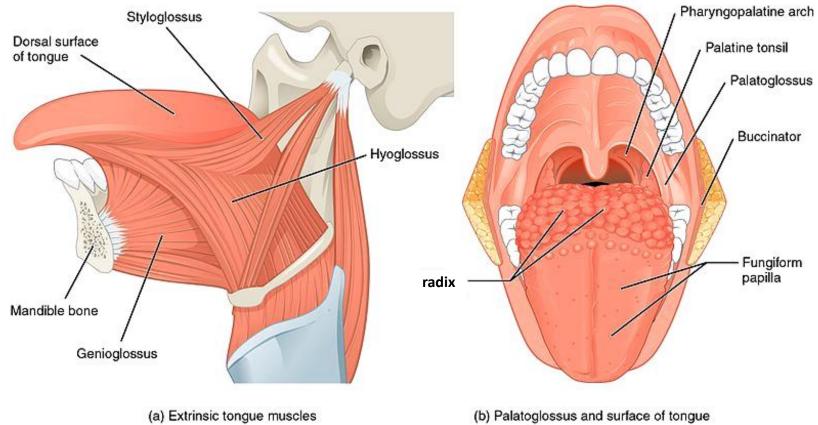




# Tongue

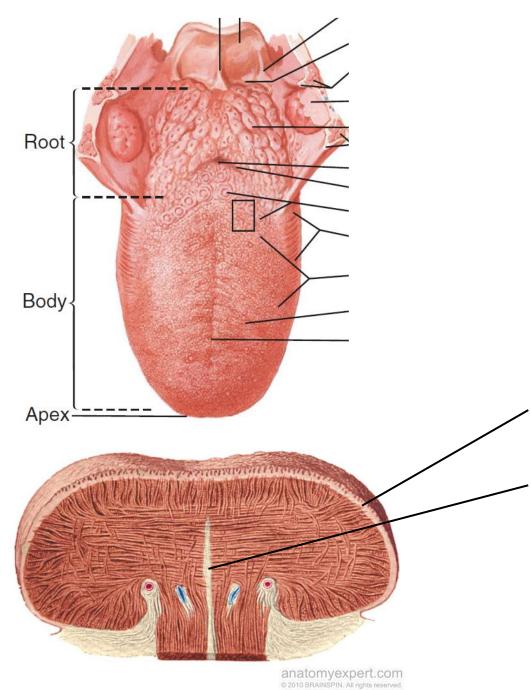
Lingua (lat.) Glossa (gr.)





**Base:** intra- and extraglossal striated muscles

**Evolutionary:** developed in terrestrial vertebrates and amphibians (tetrapods) from muscles of oral floor



#### **Surface**

**Dorsum linguae**Specialized oral mucosa

**Inferior aspect**Lining mucosa

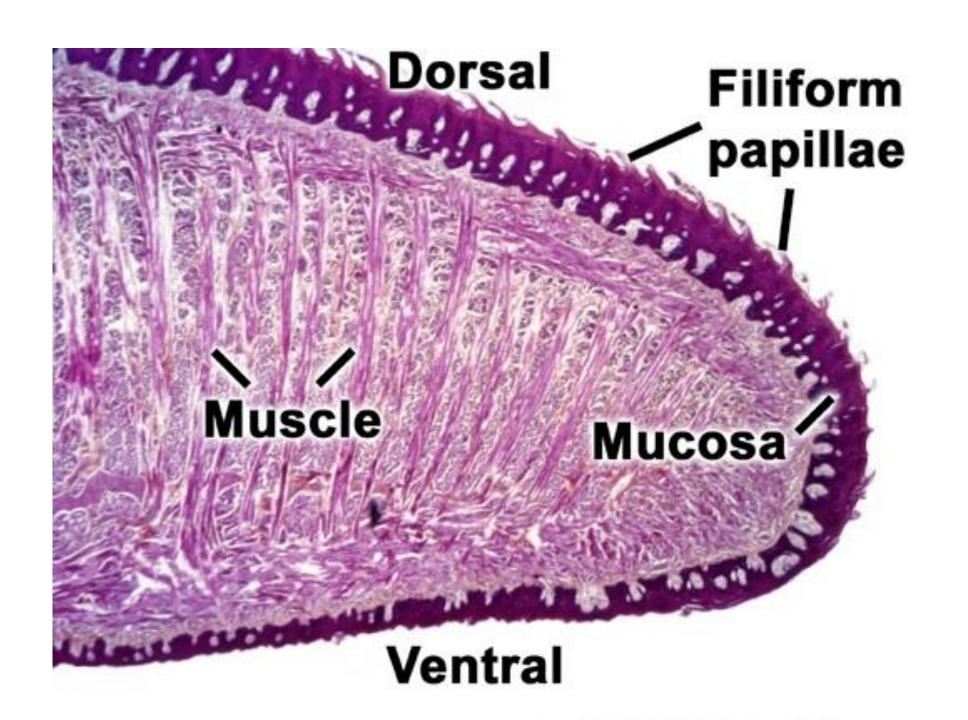
### **Fibrous parts**

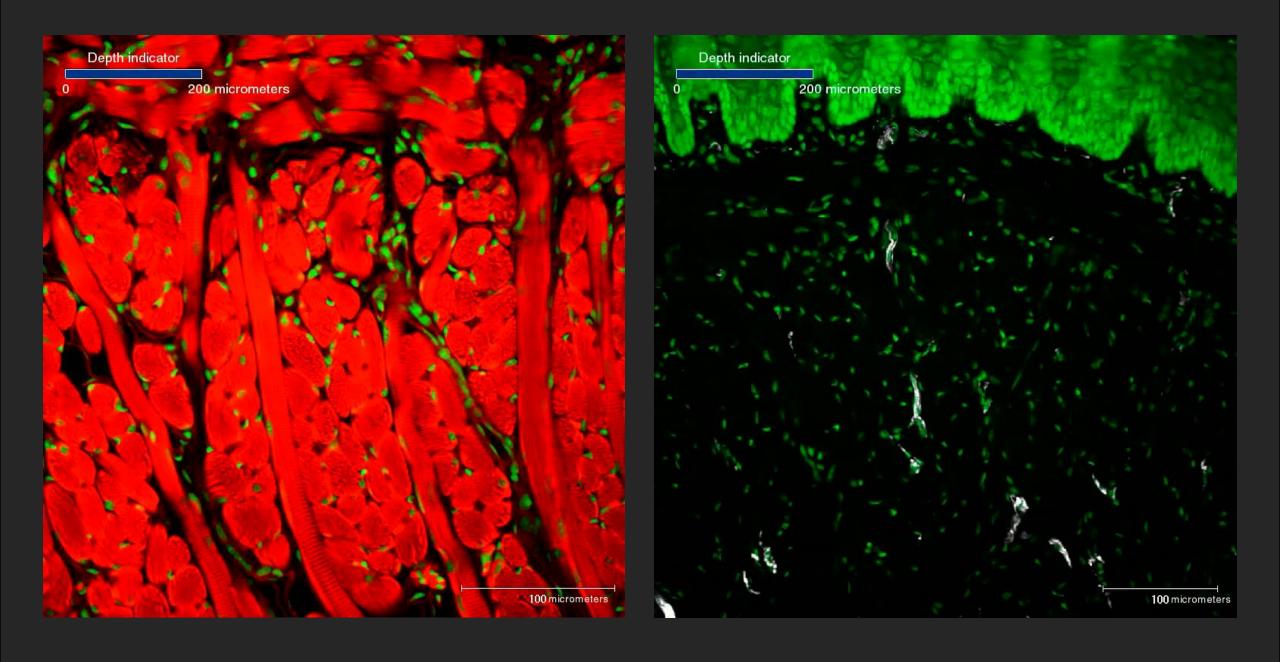
Aponeurosis linguae

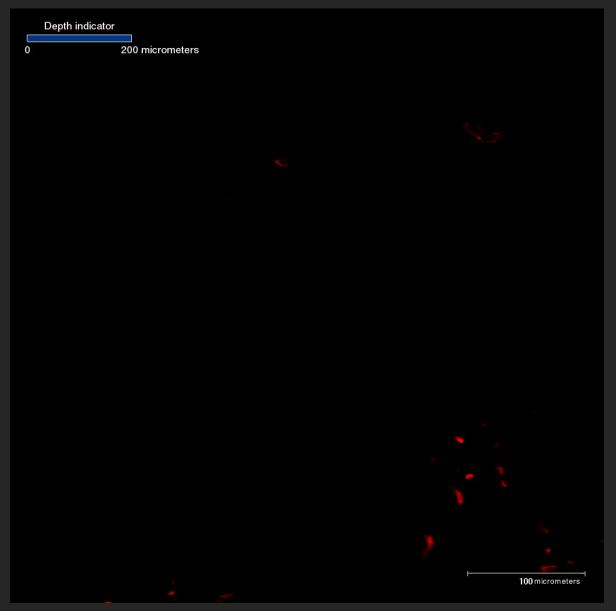
- very stiff fibrous membrane

Septum linguae

composed by dense collagenous tissue

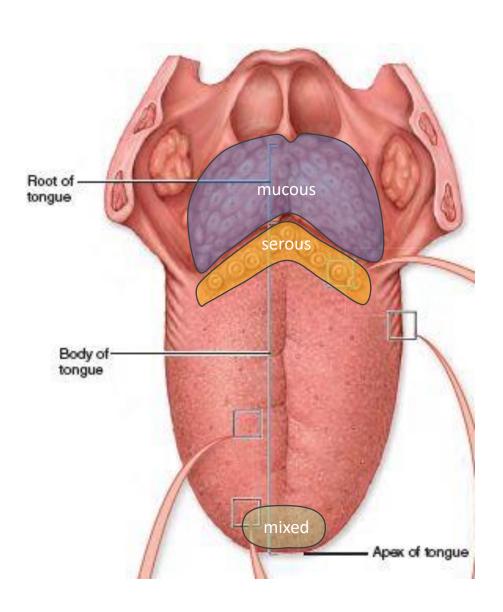








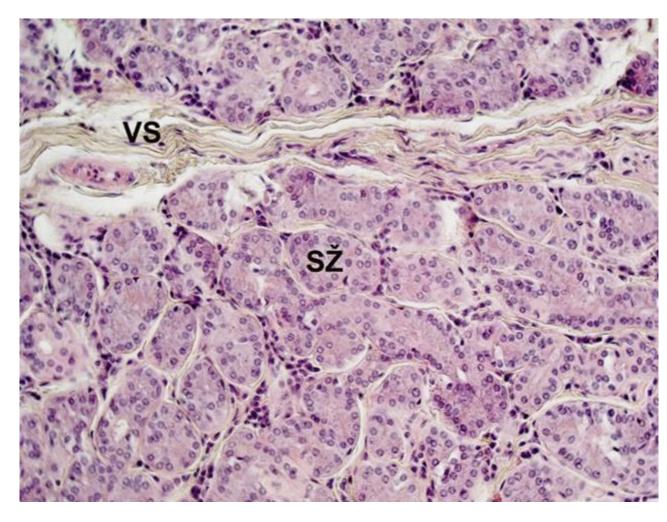
# Glands of tongue



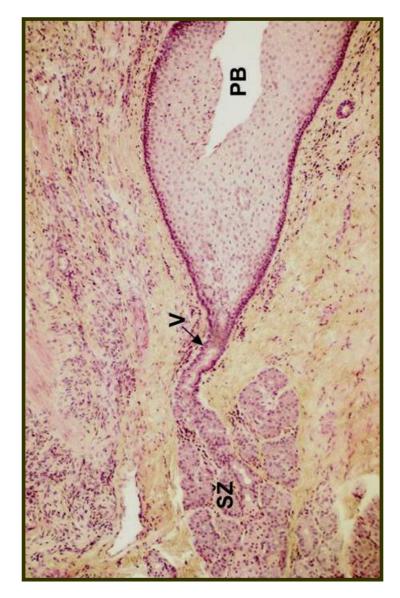
Glandula apicis linguae (gl. Blandini)

mixed gland

# **Ebner's glands** - *gll. gustatoriae* serous

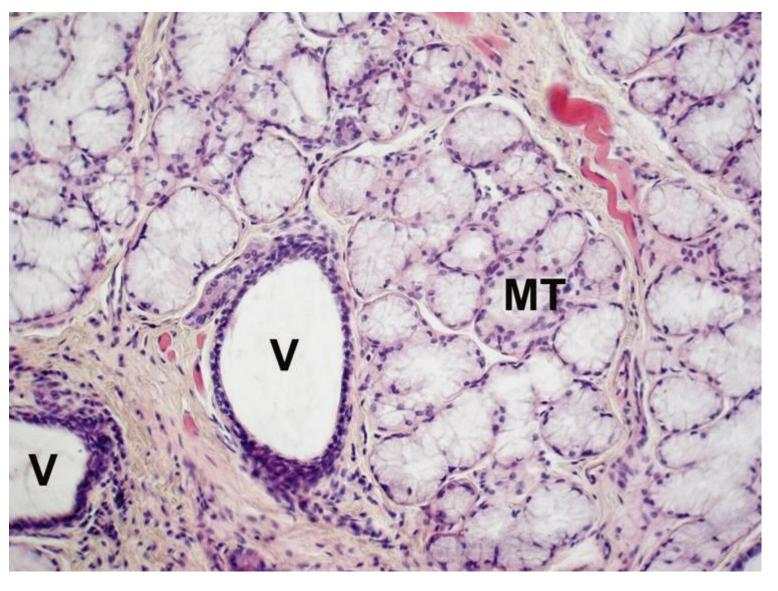


**Ebner's serous glands** (SŽ) with secretory parts of tubular character (VS – septum of connective tissue)



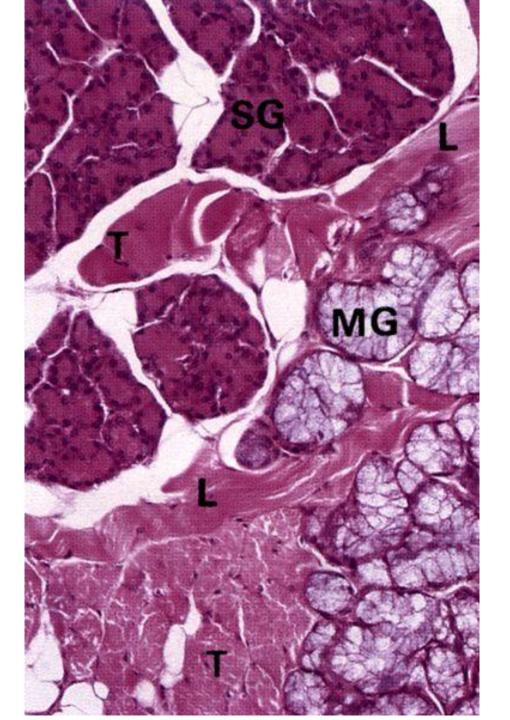
Duct (V) of Ebner's gland (SŽ)

**Weber's glands** - *gll. linguales post* mucinous



Weber's mucinous glands

MT – mucinous tubules, V – duct.



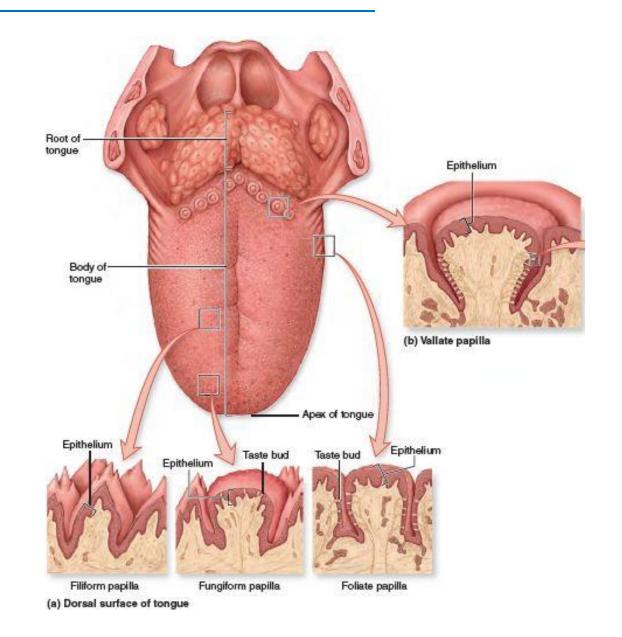
**Ebner's glands** - *gll. gustatoriae* serous

**Weber's glands** - *gll. linguales post* mucinous

# Dorsum linguae

## Specialized oral mucosa

- Firmly connected with *aponeurosis linguae*
- Rough surface
- Mucosal outgrowths lingual papillae
- Covered by nonkeratinized squamous stratified epithelium (except of papillae filiformes)

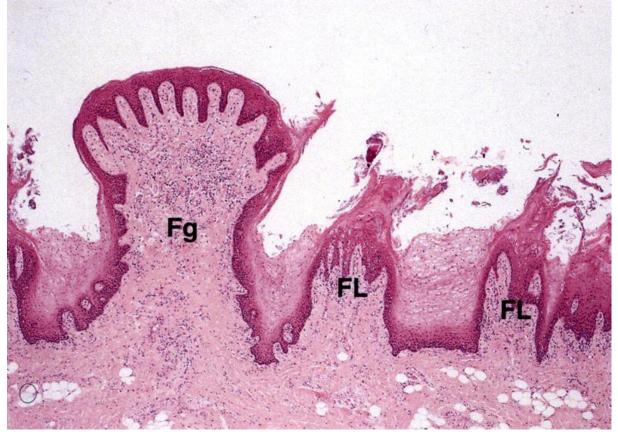


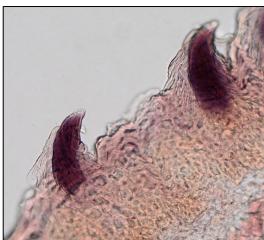
## Papillae filiformes

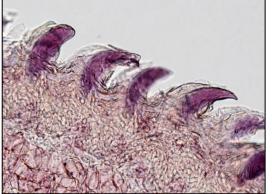
The most abundant and distributed over the entire dorsal surface of the tongue; Brush-like appearance (0.5 - 1 mm in height, 0.2 - 0.3 mm in width); The stratified squamous epithelium is often keratinized

## Papillae fungiformes

Apex; Mushroom-shape (0.5 - 1.5 in height, 0.5 - 1.0 mm in width) Taste buds in epithelium

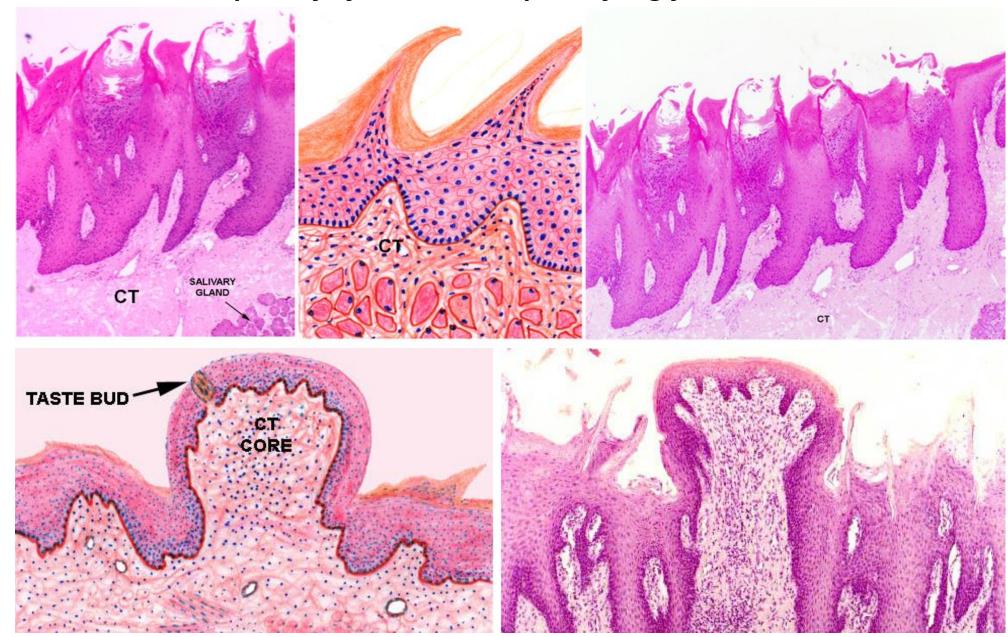


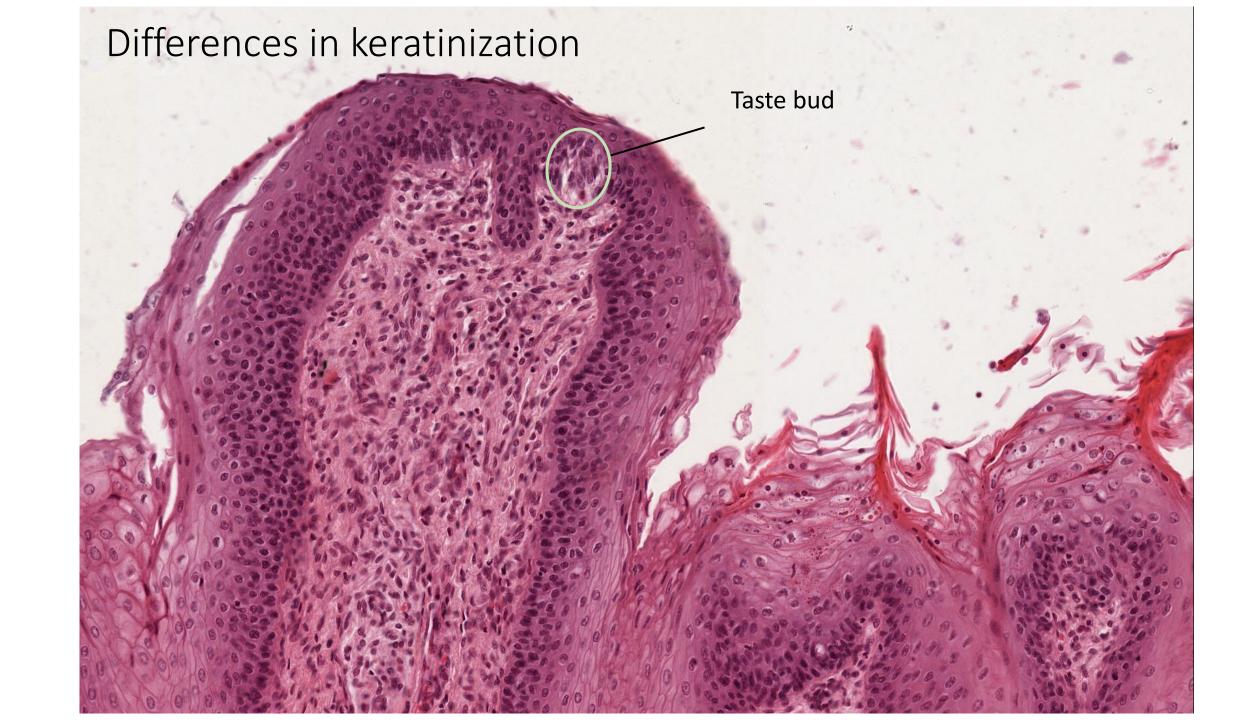






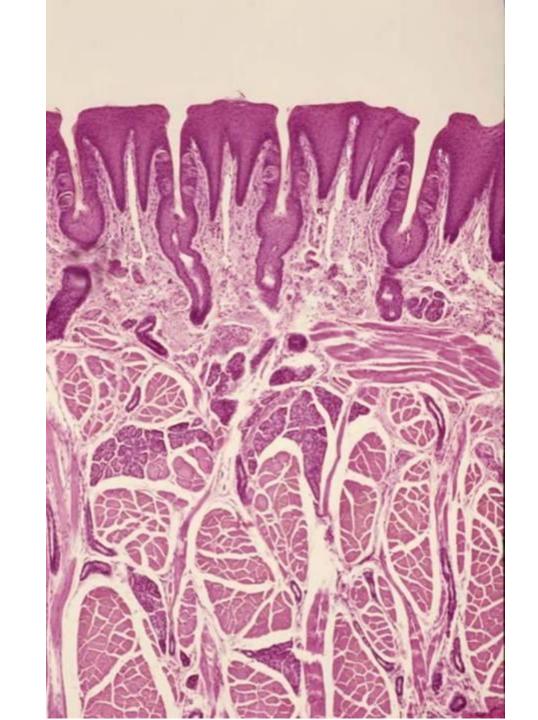
# Papillae filiformes vs. Papillae fungiformes





# Papillae foliatae

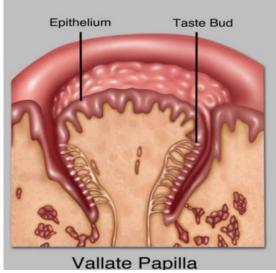
- Count: 3 8
- Vertically-oriented
- Rudimental
- Laterally on the edge of the main body and root of tongue
- Taste buds

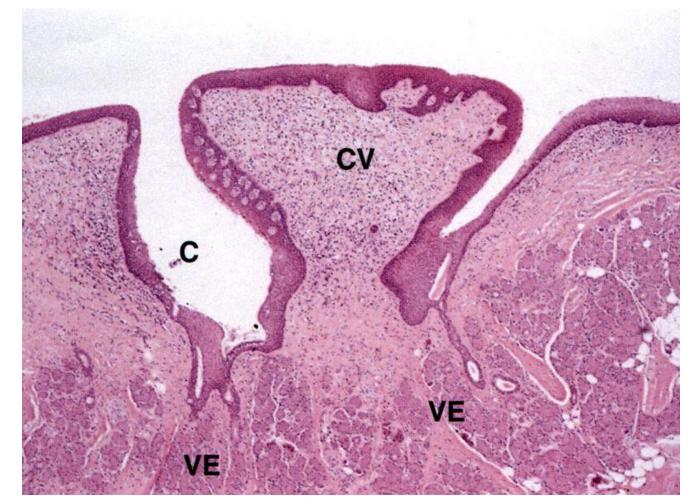


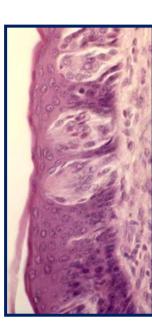
## Papillae vallatae

Largest (1-4 mm in height, 1-3 mm in width), 7–12 just in front of sulcus terminalis, submerged into mucosa. Deep circumpapillary furrow. Taste buds

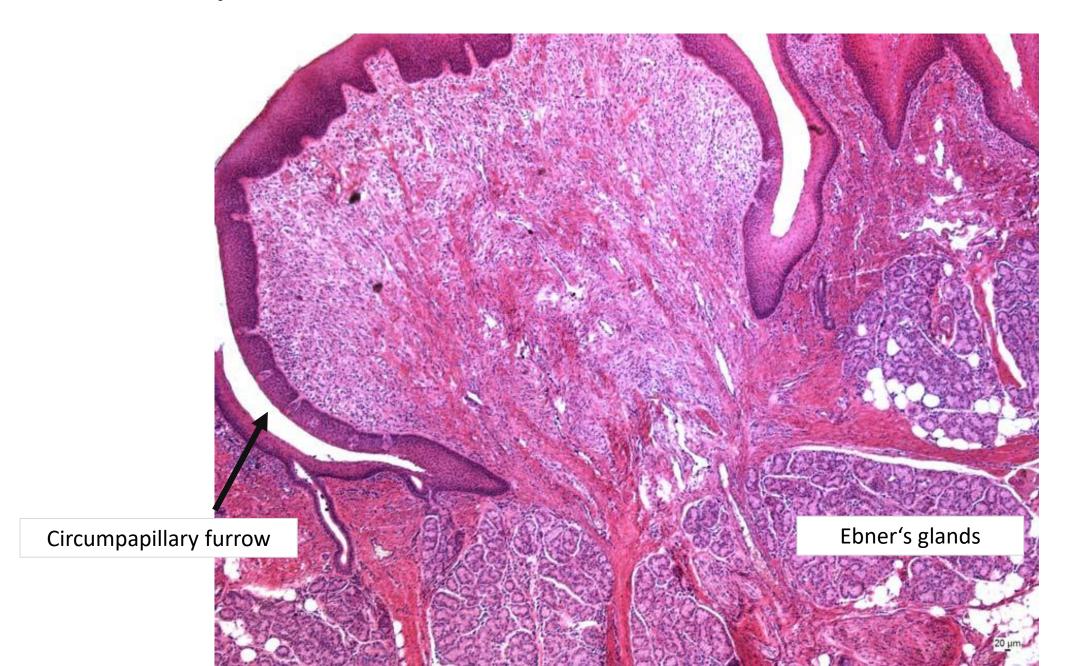








# Papilla vallata



# Taste

### **Basic tastes:**

Sweet

Salty

Sour

Bitter

Umami

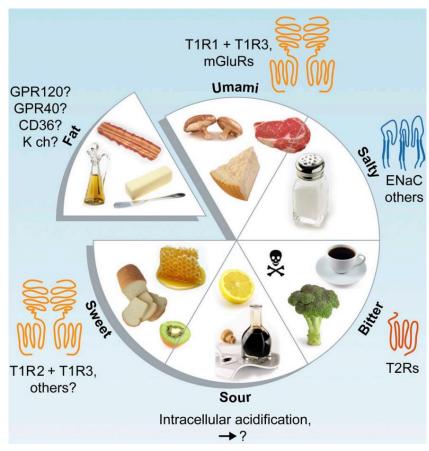
## **Suggested (still discussed):**

Fatty

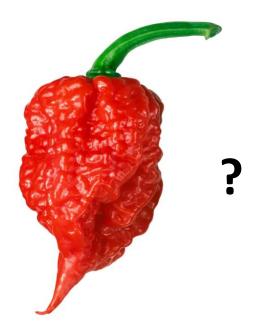
Metalic



## Why do we perceive tastes?







# Taste buds

(caliculi gustatorii)

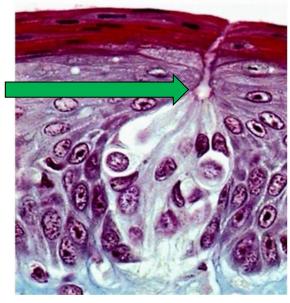
### **Intraepithelial structures**

#### Localization:

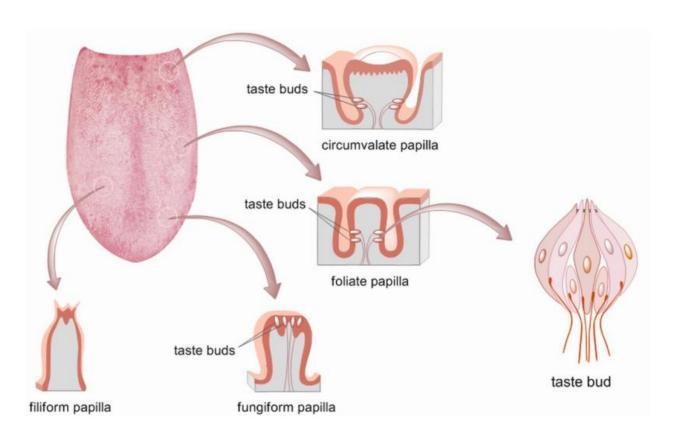
- In epithelium of vallate papillae + circumpapillar furrows
- In epithelium of fungiform papillae and foliate papillae
- Rarely in other places

The number is reported to be between 2,000 - 10,000 (their amount decreases with age)

# Porus gustatorius



(Kikut-Ligaj, et al., 2015)



# Taste buds

(caliculi gustatorii)

### **Cellular organization:**

- Each taste bud consits of about 30-80 spindle-shaped neuroepithelial cells
- There are 3 (+1) cell types in the taste buds distinguishable by electron microscopy and immunohistochemical staining:

### Type I

Most abundant (about half of the cells)
Support function, possible contribution to taste perception

### Type II

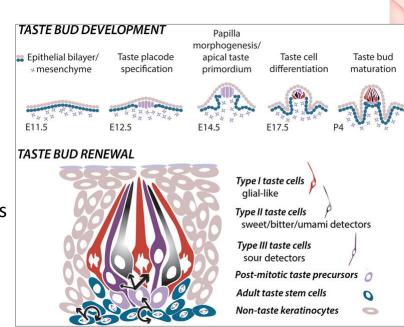
Adjacent to intraepithelial nerves
No synapses, signal transmission by
purinergic signaling

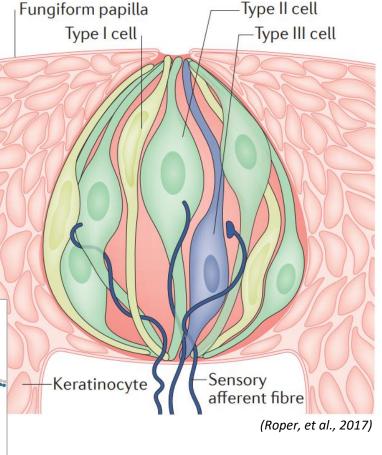
## Typ III

They are the only ones with identifiable synapses

Basal cells (Type IV)

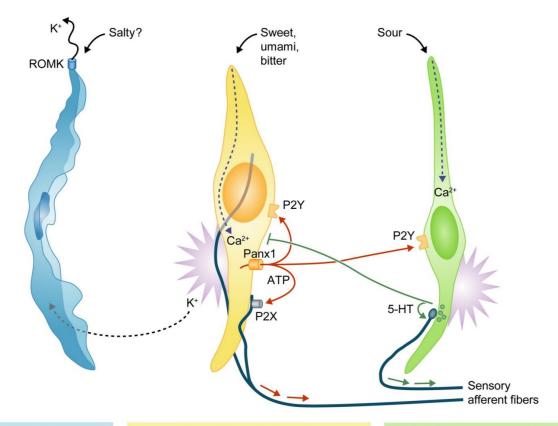
Non-differentiated cesll, stem function





(Barlow, 2021)

# Taste detection



#### Type I glial-like cell

#### Neurotransmitter clearance

GLAST Glutamate reuptake

NTPDase2 Ecto-ATPase

NET Norepinephrine uptake

#### Ion redistribution and transport

ROMK K+ homeostasis

#### Other

OXTR Oxytocin signaling?

### Type II receptor cell

#### Taste transduction

T1Rs, T2Rs Taste GPCRs mGluRs Taste GPCRs  $G\alpha$ -gus,  $G\gamma$ 13 G protein subunits

PLCβ2 Synthesis of IP3

TRPM5 Depolarizing cation current

#### Excitation and transmitter release

Na<sub>v</sub>1.7, Na<sub>v</sub>1.3 Action potential generation

Panx1 ATP release channel

### Type III presynaptic cell

#### Surface glycoproteins, ion channels

NCAM Neuronal adhesion

PKD channels Sour taste?

#### Neurotransmitter synthesis

AADC Biogenic amine synthesis

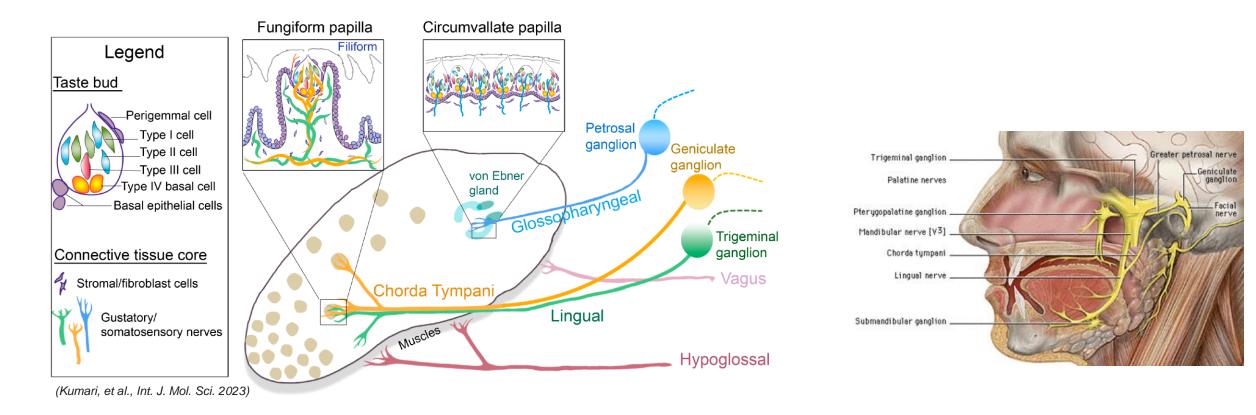
GAD67 GABA synthesis
5-HT Neurotransmitter

Chromogranin Vesicle packaging

#### Excitation, transmitter release

Na<sub>v</sub>1.2 Action potential generation
Ca<sub>v</sub>2.1, Ca<sub>v</sub>1.2 Voltage-gated Ca<sup>2+</sup> current
SNAP25 SNARE protein, exocytosis

# Transmission of taste signals



#### Inervation of taste buds

- Taste buds on fungigorm papillae n. facialis chorda tympani (through lingual nerve)
- Taste buds on *foliate papillae* and vallate papillae *n. glossopharyngeus*
- Taste buds in other locations (radix of the tongue, the isthmus faucium n. vagus