

Development of the skin and its derivatives

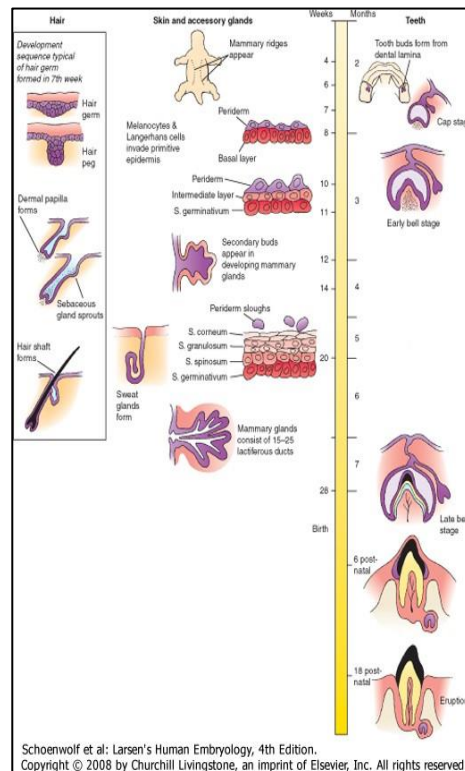
รองศาสตราจารย์ ดร. วิภาวี หีบแก้ว
สาขาวิชากายวิภาคศาสตร์





Learning objectives

To understand



Skin function and anatomy

Skin origins

Development of the overlying epidermis

Development of epidermal appendages:

Hair follicles

Glands

Nails

Teeth

Development of melanocytes

Development of Langerhans cells

Development of the Dermis

Skin Function and Anatomy

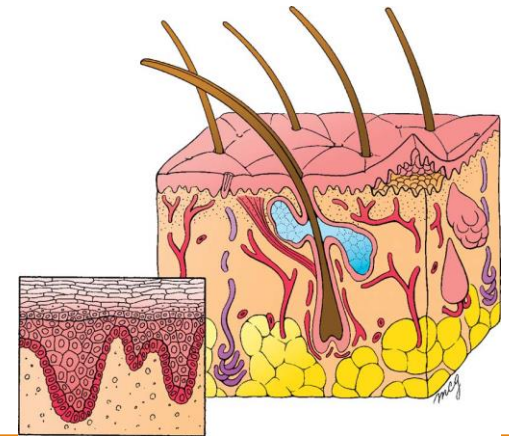
Largest organ of our body (20%)

- Protects inner body from outside world (pathogens, water, sun)
- Thermoregulation

Diverse: thick **vs** thin skin, scalp skin **vs** face skin, etc

Consists of:

- Overlying epidermis
- Epidermal appendages:-
 - Hair follicles,
 - Glands: sebaceous, sweat, apocrine, mammary
 - Nails
 - Teeth
- Melanocytes
- Langerhans cells
- (Merkel Cells)
- Dermis
- Hypodermis



Derivatives of the three Germ Layers

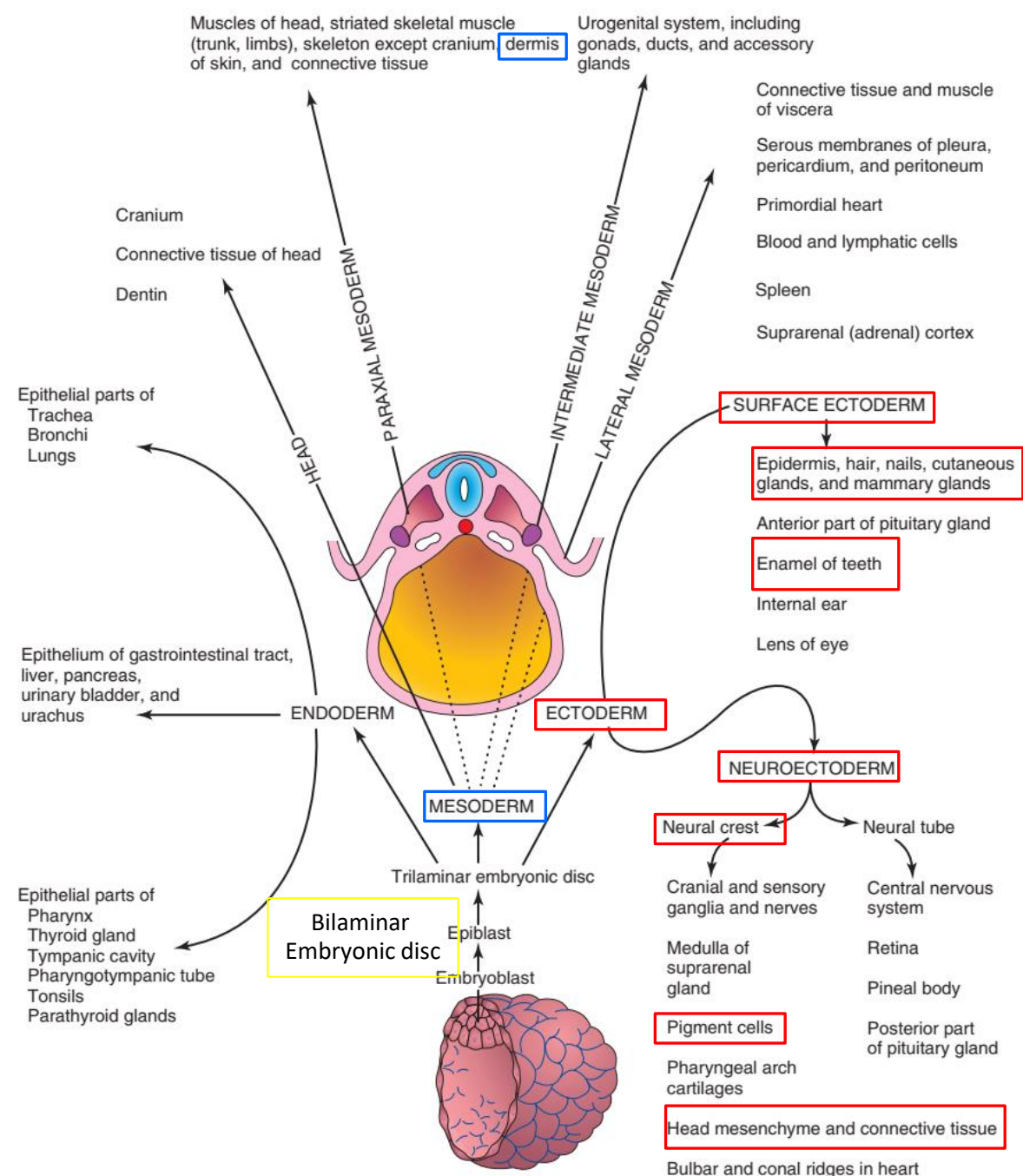
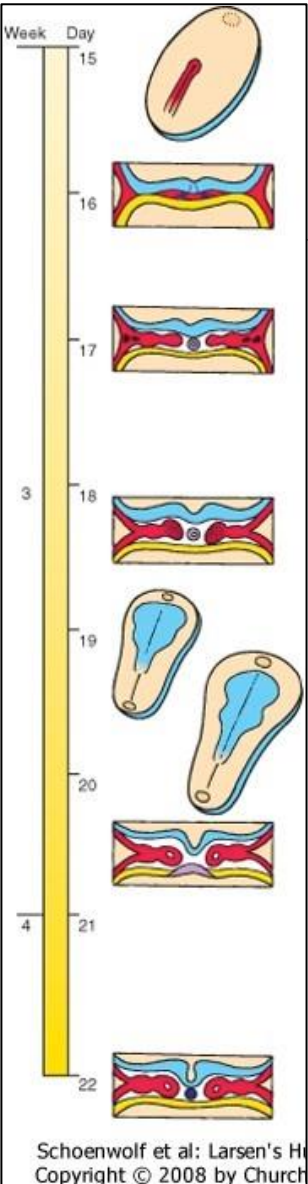


FIGURE 5-5 Schematic drawing of derivatives of the three germ layers, ectoderm, endoderm, and mesoderm. Cells from these layers contribute to the formation of different tissues and organs.

Ectoderm

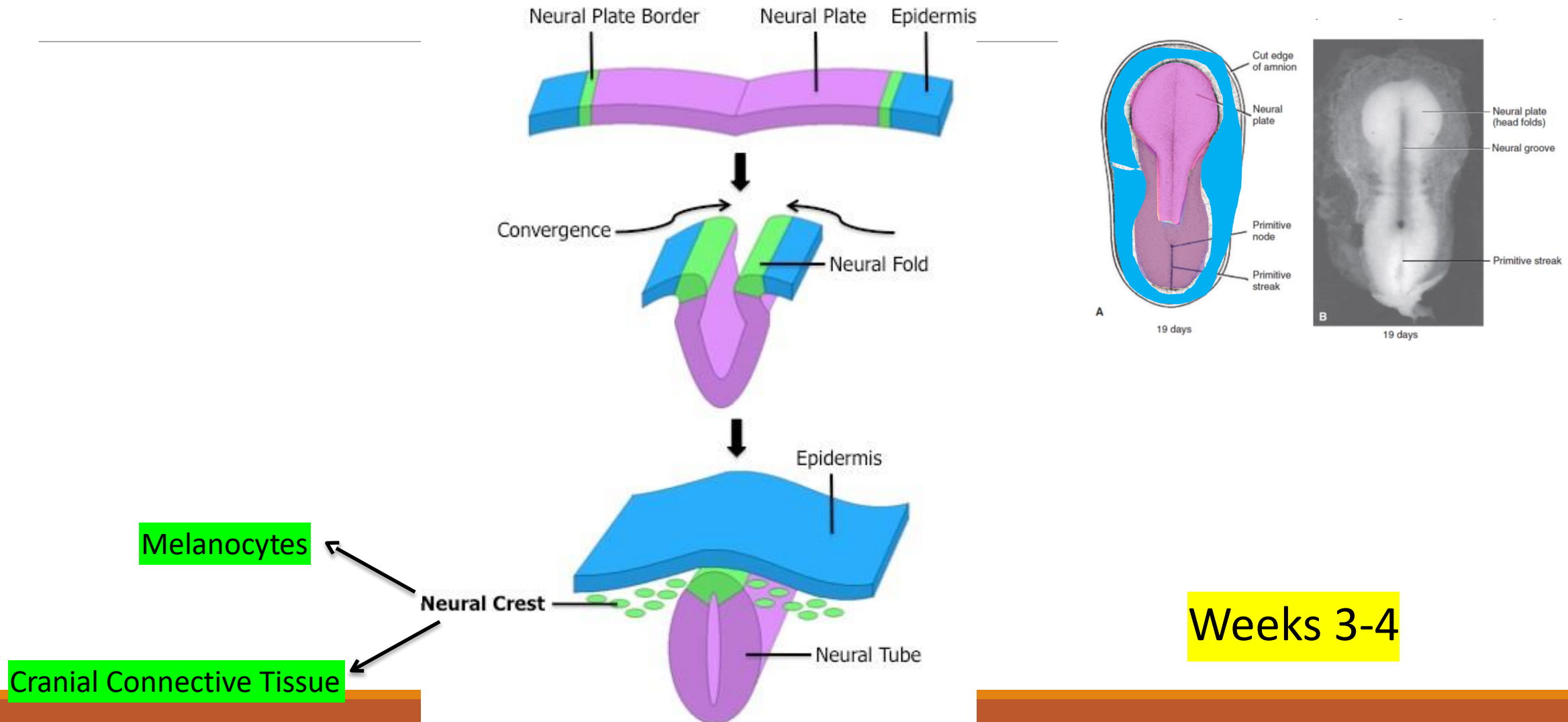
Overlying epidermis, epidermal appendages, nerve endings



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Neural Crest

Melanocytes + Cranial Connective Tissue



Dermis

Connective tissue with:

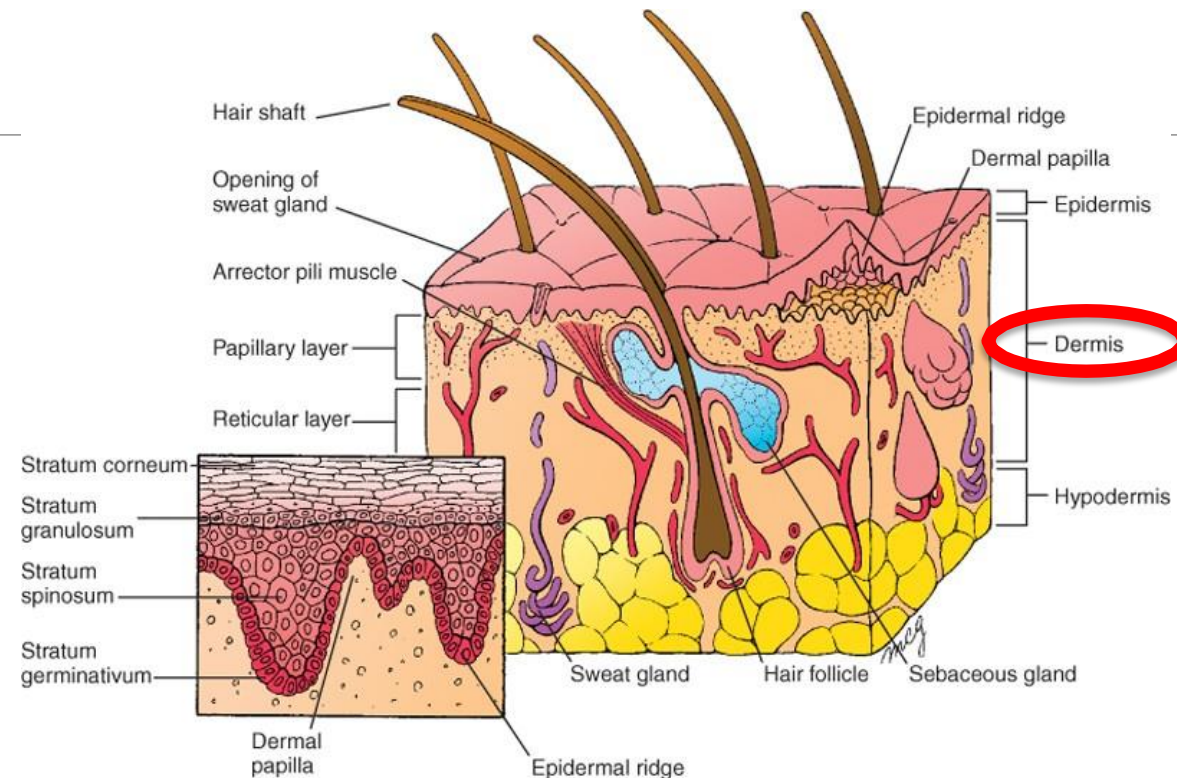
Fibroblasts

Blood vessels

Nerve endings

Sensory receptors

Muscle bundles



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Dermis development



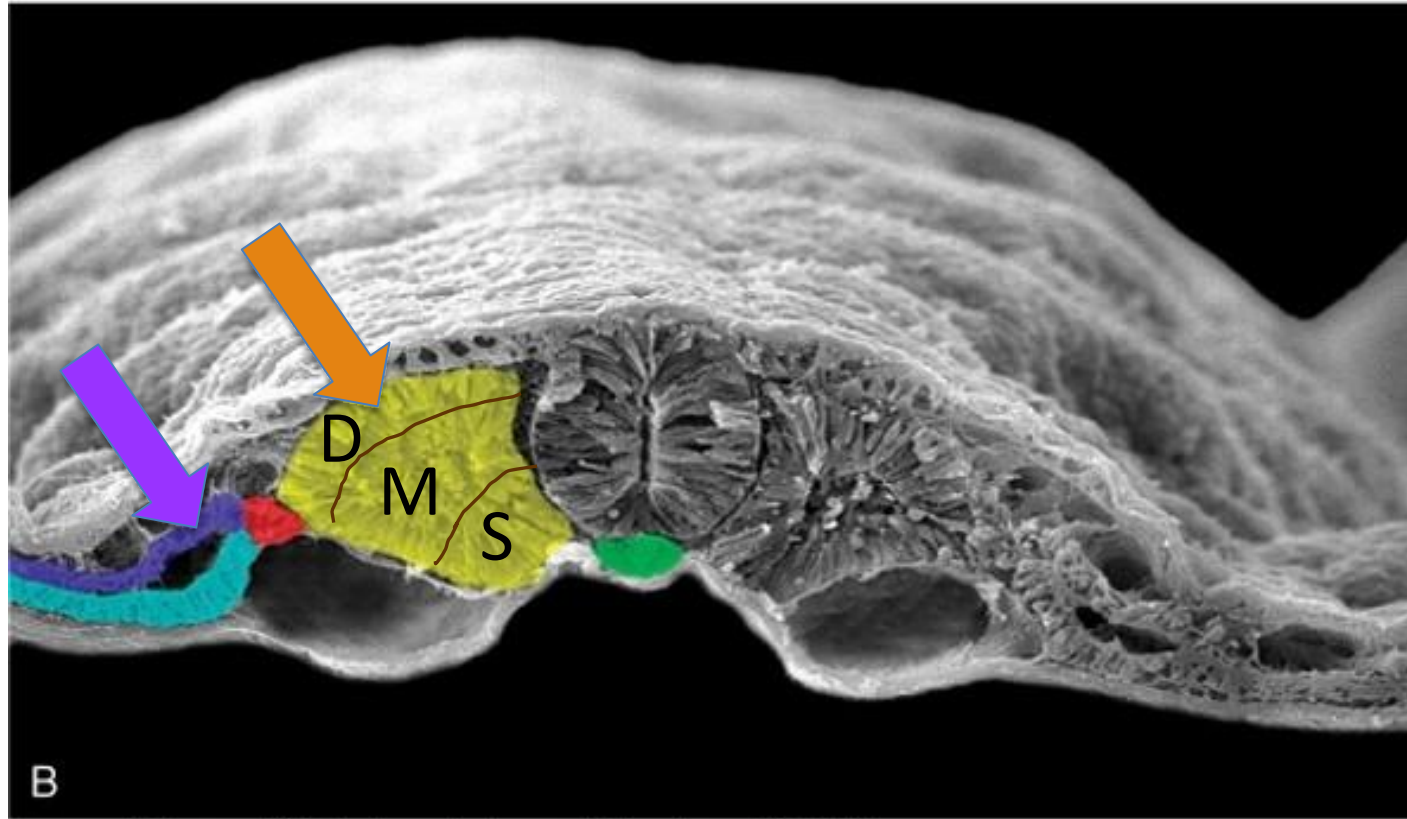
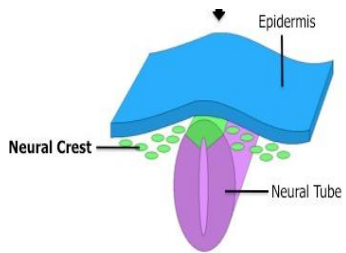
Somitic mesoderm -> trunk (back)



Somatic lateral plate mesoderm -> trunk (anterior), limbs



Cranial neural crest derived mesenchyme -> cranial region



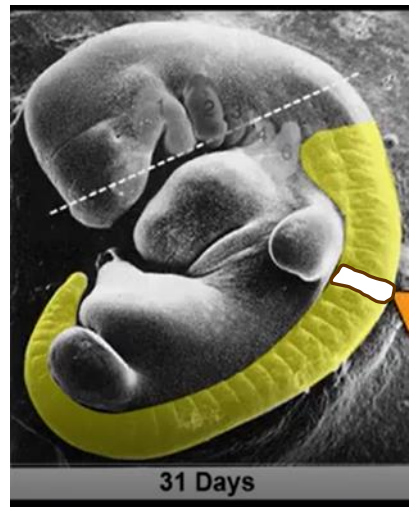
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Dermatome : area of skin supplied by a single spinal nerve level

Dorsal ramus

(Somitic mesoderm -> trunk (back))

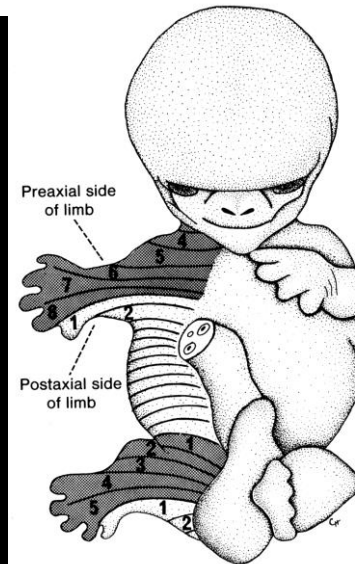
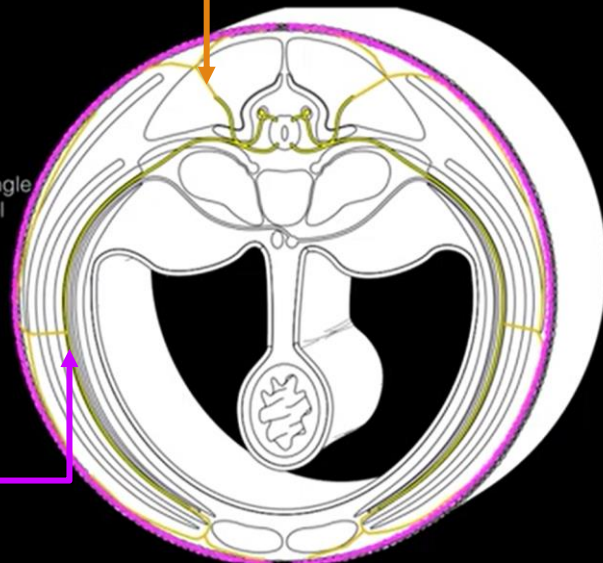


Somite formation

- Somites

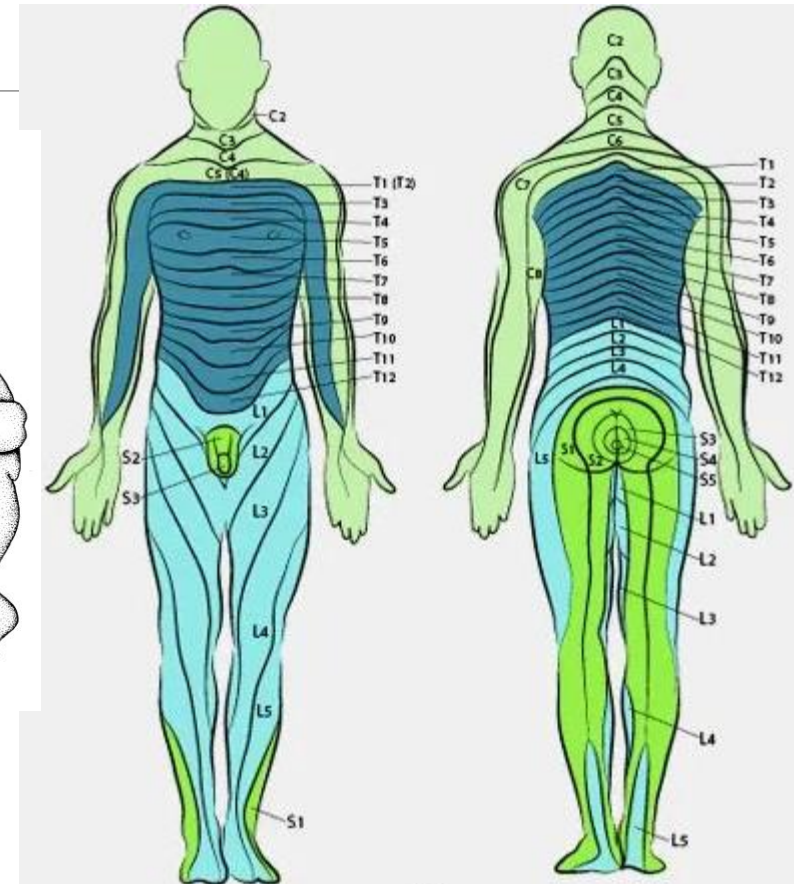
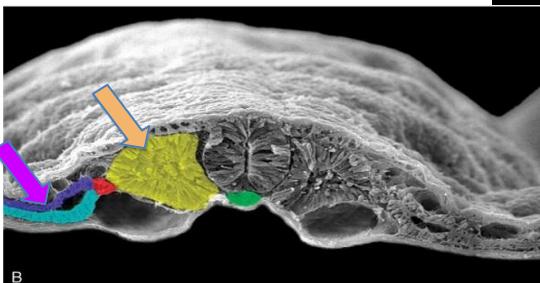
- Dermatome

An area of skin supplied by a single spinal nerve level



Ventral ramus

Somatic lateral plate mesoderm -> trunk (anterior), limbs



Epidermal development

Surface ectoderm

Epidermis: stratified squamous epithelium

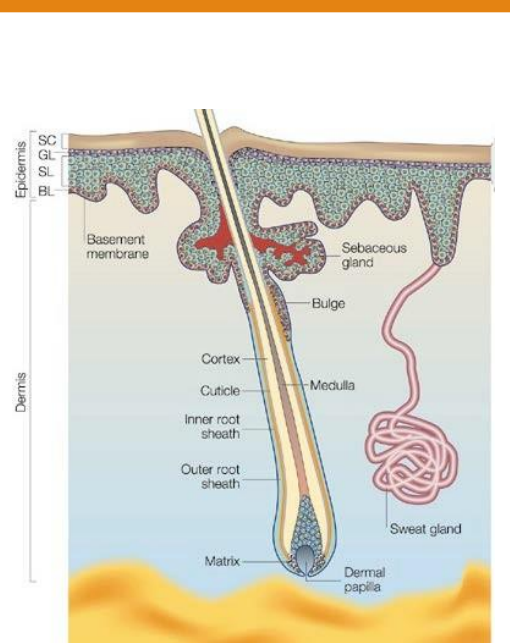
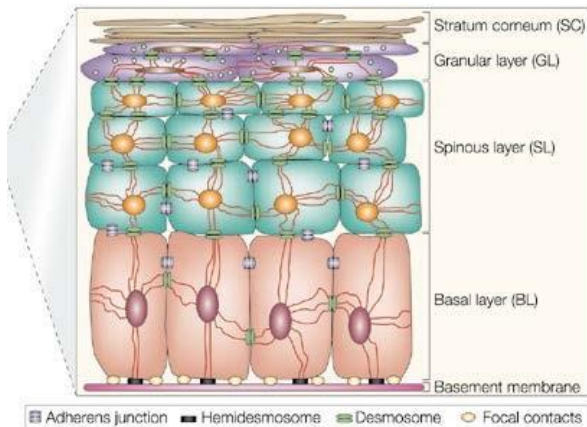
Appendages:

Hair follicles

Glands (sebaceous, sweat, apocrine, mammary, lacrimal, salivary)

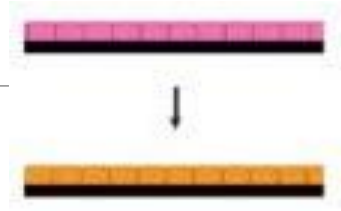
Nails

Teeth



Development of Skin/interfollicular epidermis (IFE)

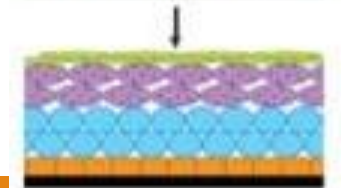
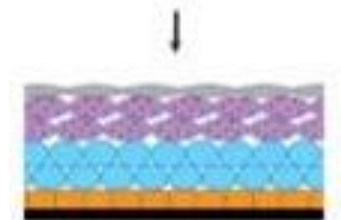
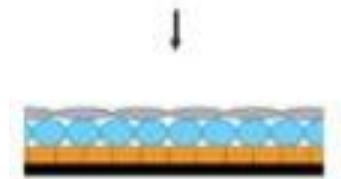
Week 4



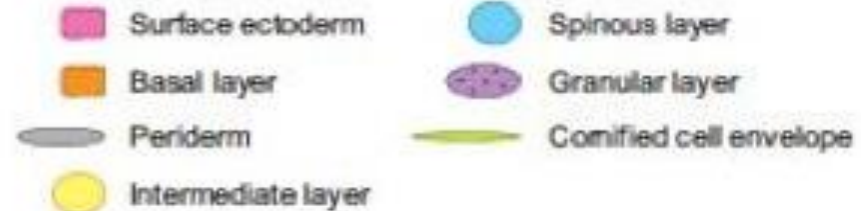
Week 11



Onset of stratification



Week 24

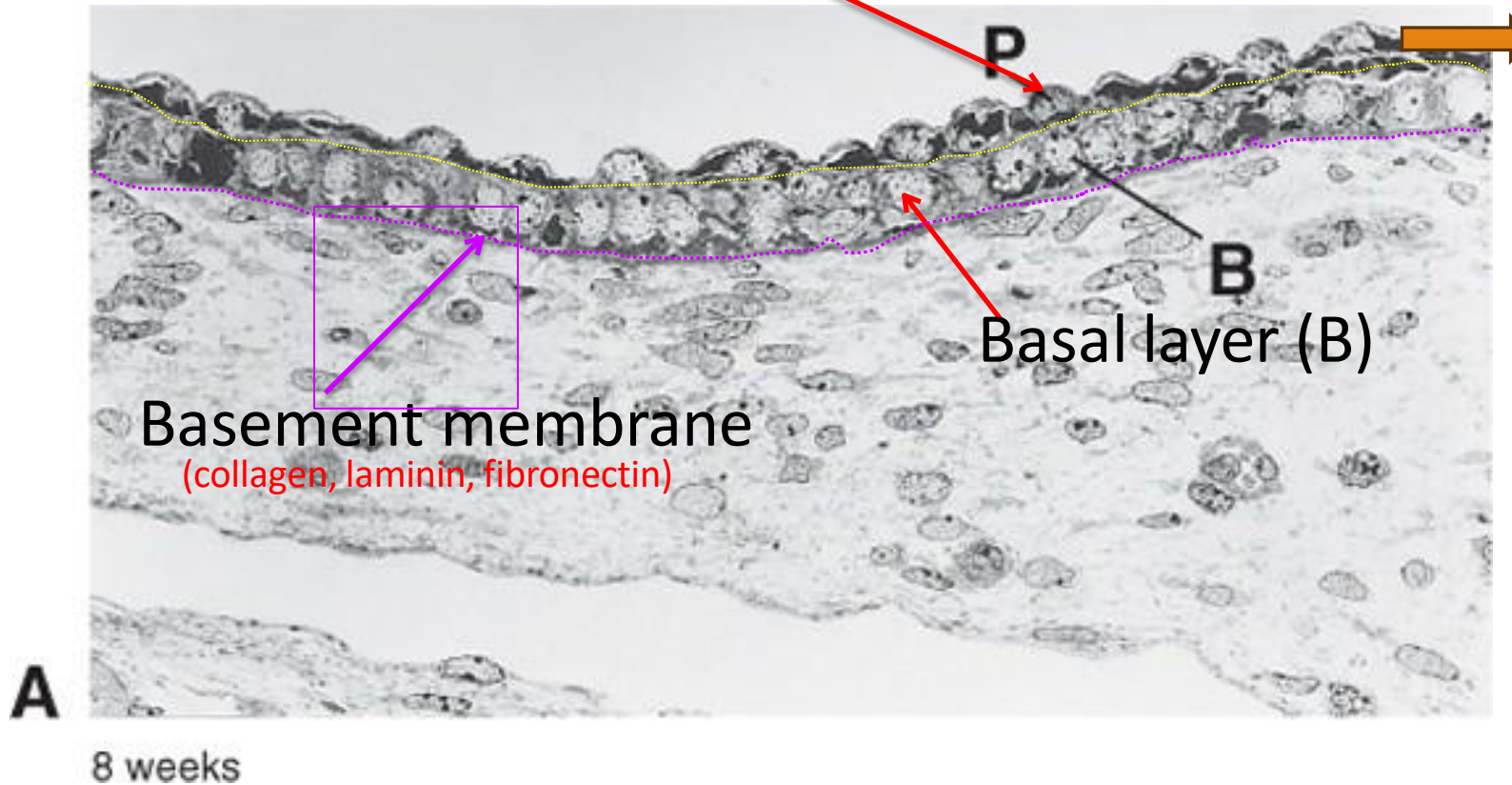


Development of Skin



Periderm formation: **4 weeks**

Periderm (P)



1. แลกเปลี่ยนน้ำและสารละลายระหว่างตัวอ่อนกับน้ำคร่ำ
2. ป้องกันเซลล์ชั้นล่าง

Development of Skin/Stratification

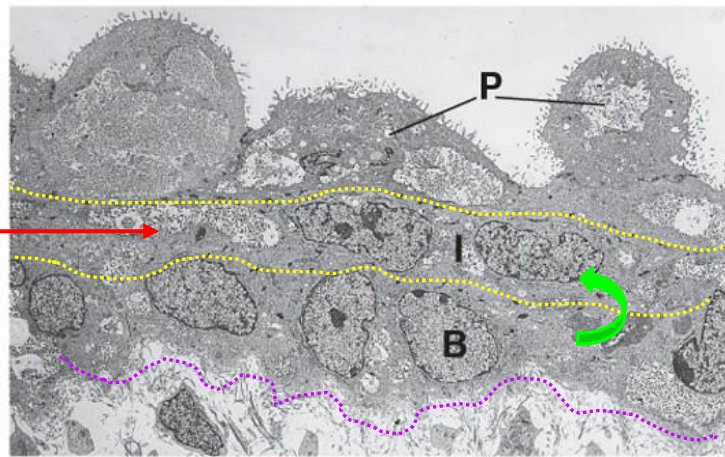
Onset of stratification

11 weeks (3 Mo): intermediate layer

Periderm (P)

Intermediate layer (IL)

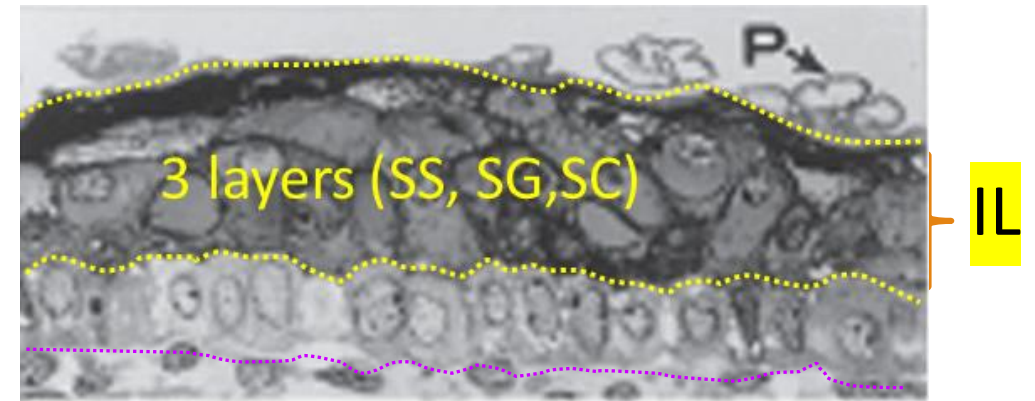
Germinative layer (B)
(Stratum germinativum: SG)
Layer of **stem cells**



11 weeks
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5 months

Periderm (P) is shed

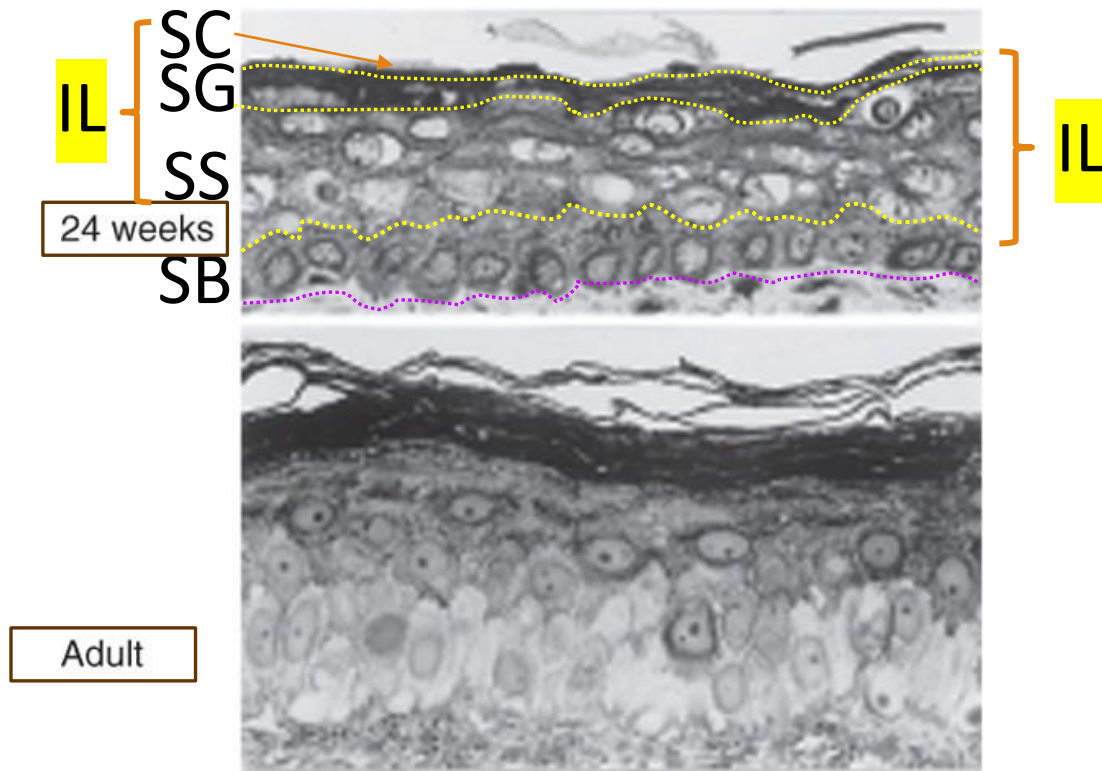


17 weeks

The basal layer later forms ridges and hollows, which are reflected on the surface of the skin in the fingerprint.

Development of Skin/Stratification

6 Months: Full thickness



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Development of Skin/Stratification

Onset of stratification

11 weeks: intermediate layer

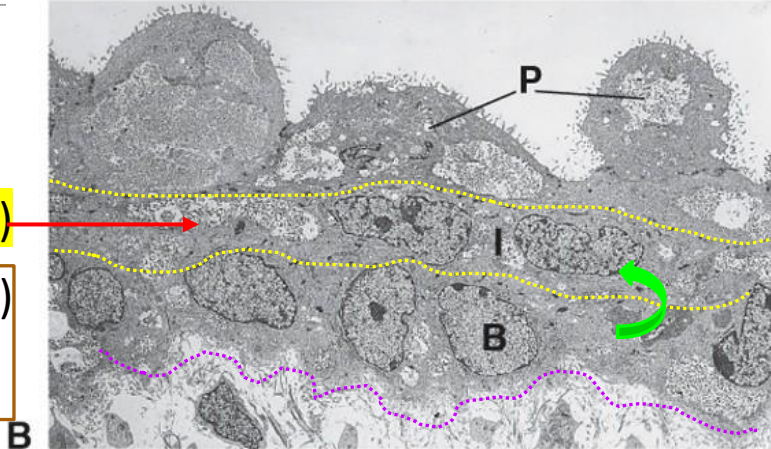
Periderm (P)

Intermediate layer (IL)

Germinative layer (B)

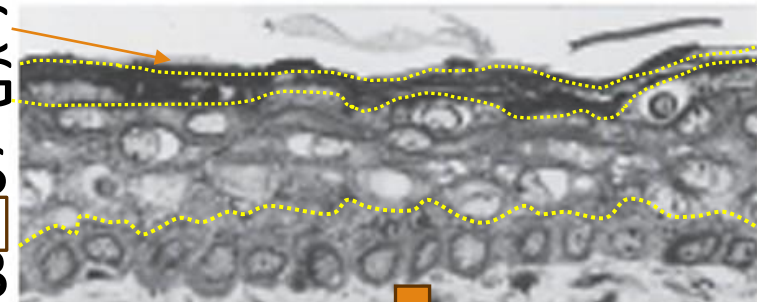
(Stratum germinativum: SG)

Layer of **stem cells**



11 weeks
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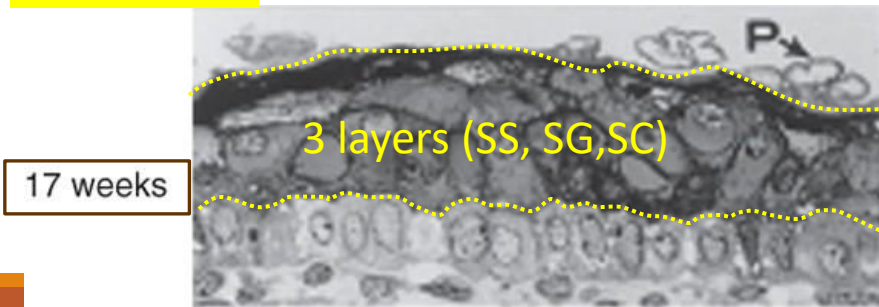
IL
SC
SG
SS
SB
24 weeks



IL

5 months

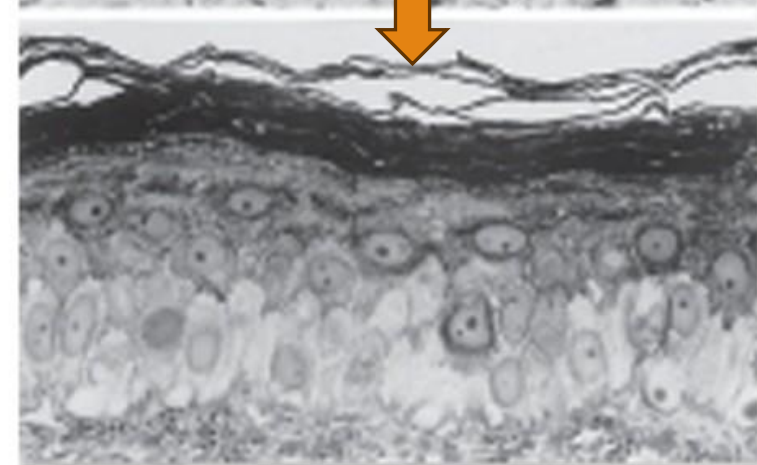
Periderm (P) is shed



17 weeks

3 layers (SS, SG, SC)

Adult



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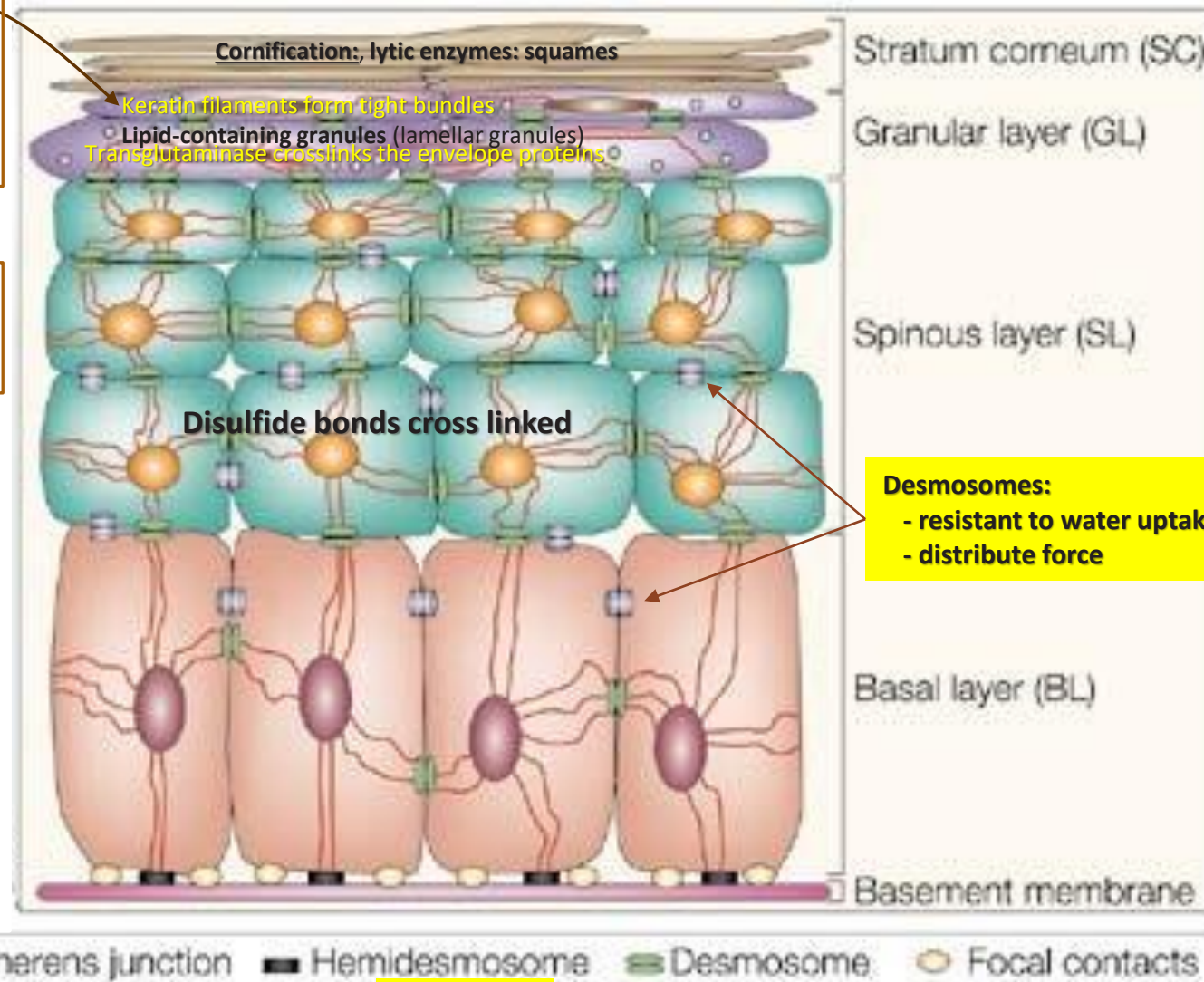
Stratified epidermis

Filaggrin: flatten cell

Envelope proteins { Loricrin
Involucrin
Envoplakin

Secondary Keratin proteins
Keratin 1/10

Primary Keratin filaments
Keratin 5/14



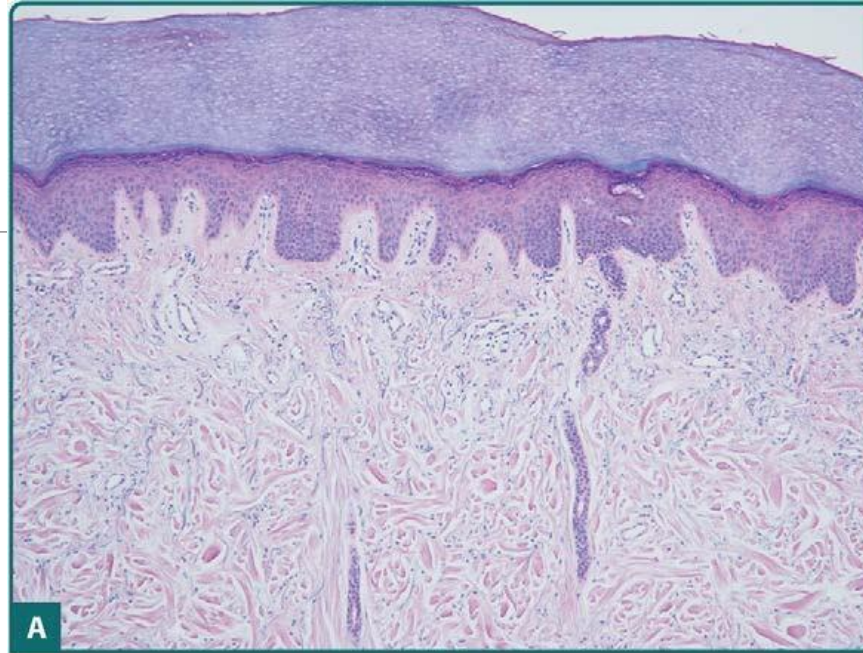
Desmosomes:

- resistant to water uptake or loss and infection
- distribute force

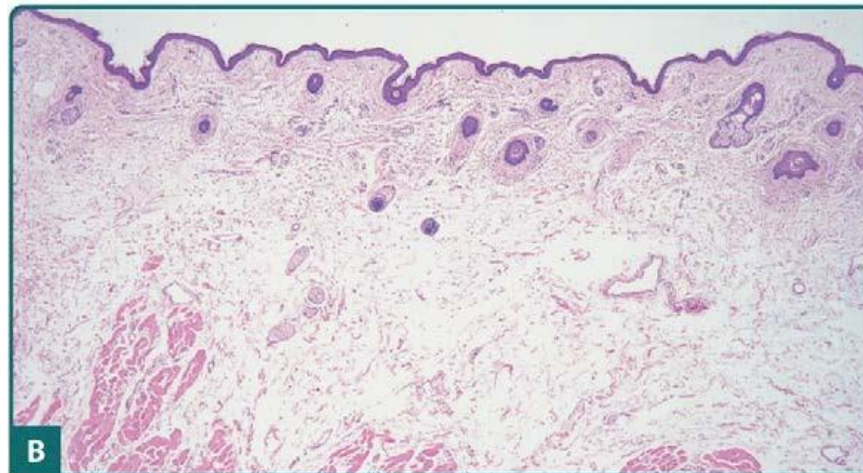
essential for cell survival and determines the orientation of cell divisions

Stratified skin

Thick skin



Thin skin

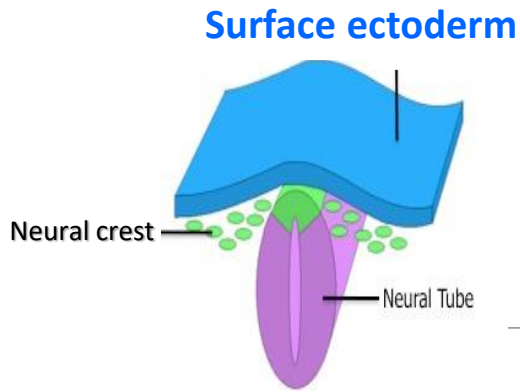


Keratinization of the Skin

Ichthyosis, excessive keratinization of the skin, is characteristic of a group of hereditary disorders that are usually inherited as an autosomal recessive trait but may also be X-linked. In severe cases, ichthyosis may result in a grotesque appearance, as in the case of a **harlequin fetus** (Fig. 21.2).



Figure 21.2 Ichthyosis in a harlequin fetus with massive thickening of the keratin layer, which cracks to form fissures between thickened plaques.

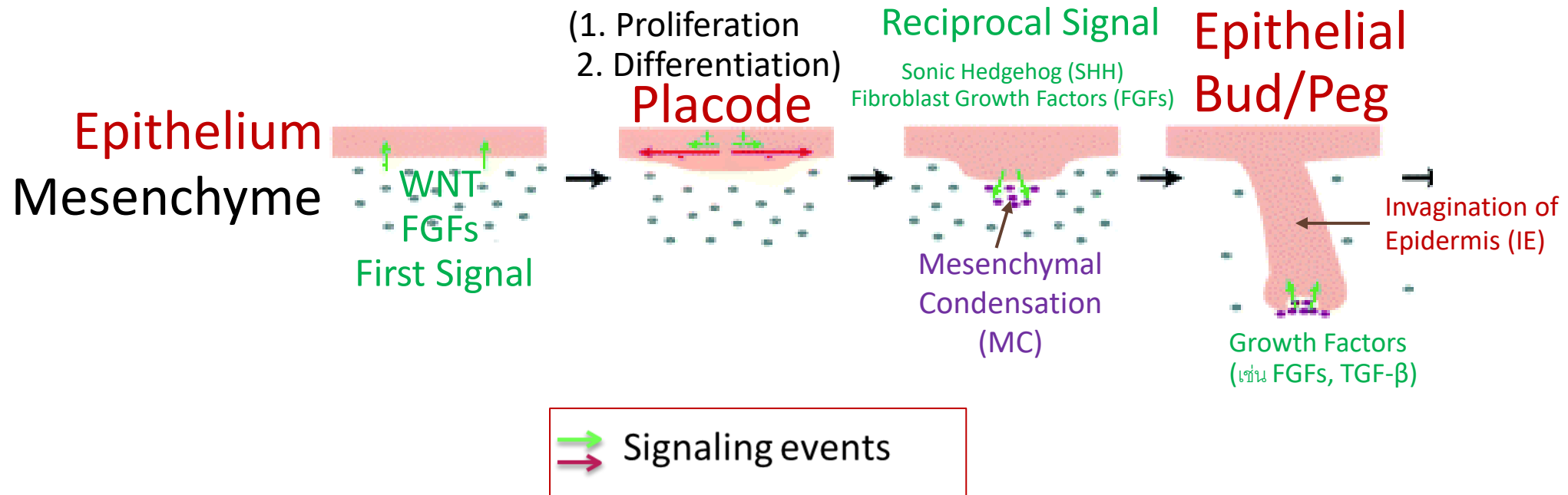


Epidermal Appendages

Surface ectoderm

- Hair follicles,
- Glands: **sebaceous, sweat, apocrine,** mammary, salivary, lacrimal
- Nails
- Teeth

Share common developmental mechanisms:



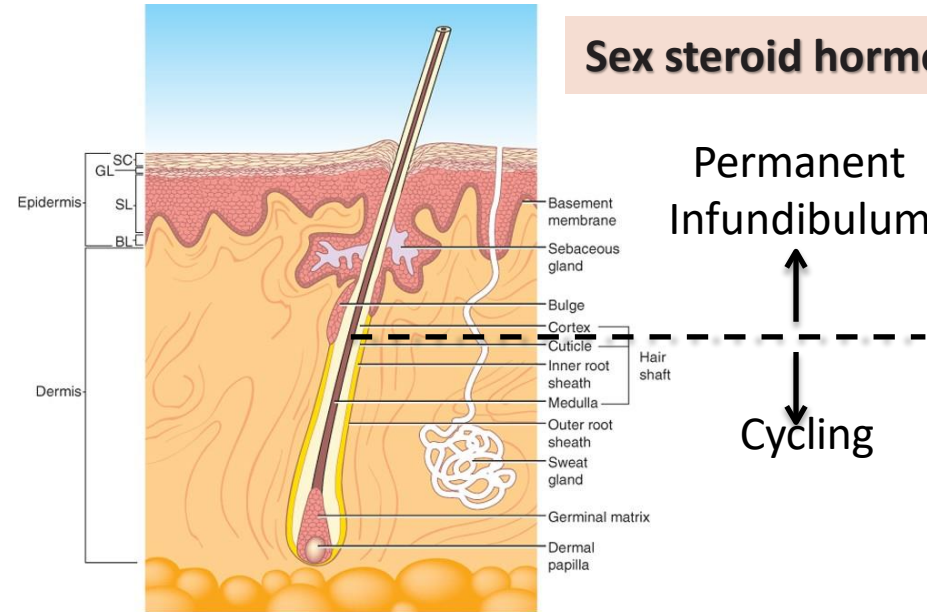
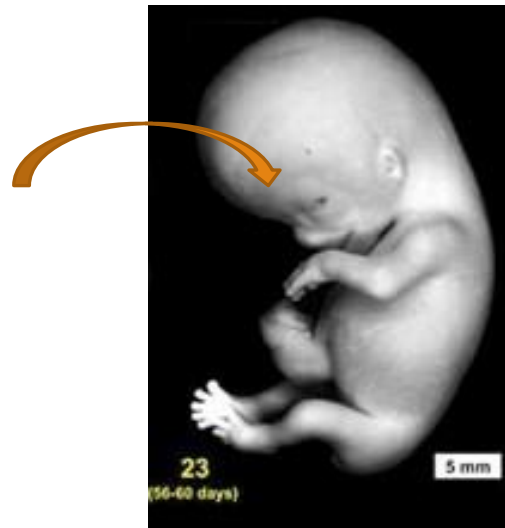
Epithelial-mesenchymal interaction induces the formation of the **placode (P)**, followed by **mesenchymal condensation (MC)** and the **invagination of the epidermis (IE)** into the dermis.

Hair follicle development and cycling

Epidermal appendage

Involved in thermoregulation/sun protection

eyebrows,
eyelids,
upper
lip,
chin.



Sex steroid hormones dependent

Months 2-4: Onset hair follicle development

Hair follicles develop **first in cranial region**

Month 5: most hair follicles present ≈ **5,000,000** hair follicles.

it is believed that novel hair follicles do not form after birth

Hair follicle development

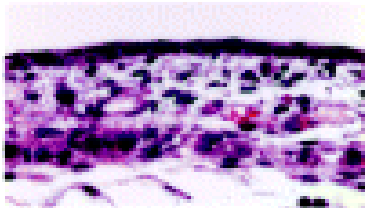
(= 3 months = 12 weeks)

Month 2

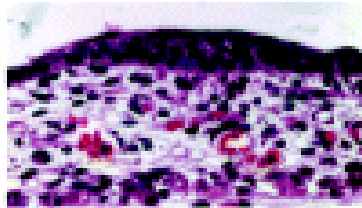
+ 1 Month

Birth: 5,000,000

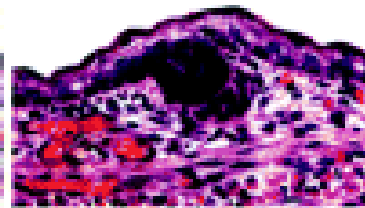
(i) UNDIFFERENTIATED
EPITHELIUM



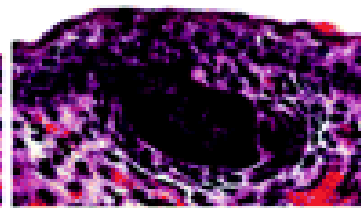
(ii) PLACODE



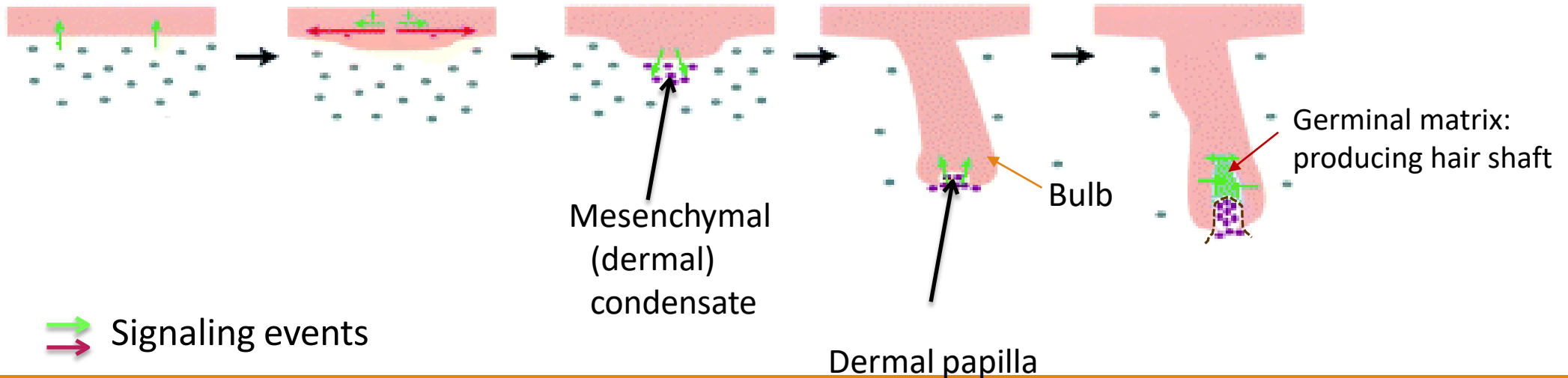
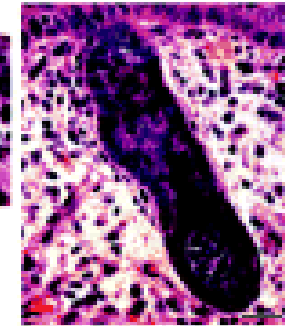
(iii) GERM



(iv) PEG



(v) BULBOUS PEG



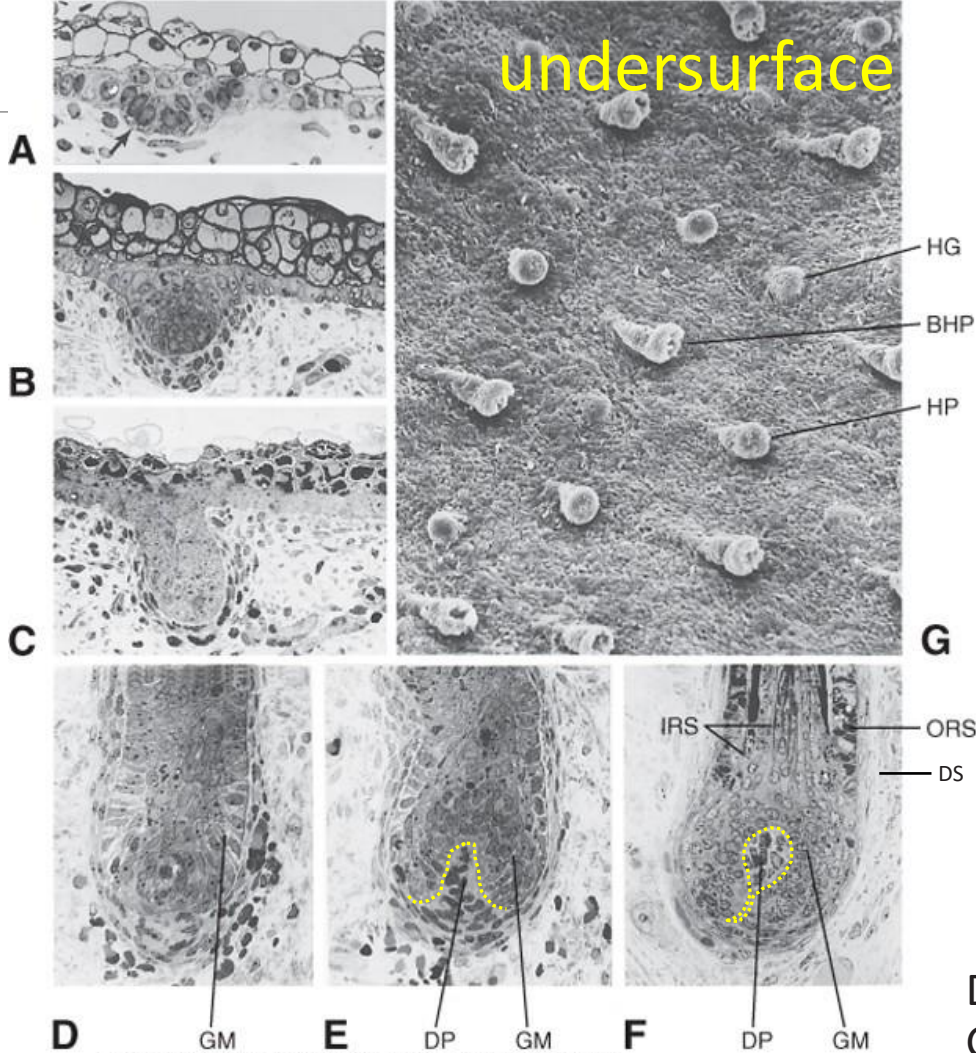
Hair follicle development

Hair germ (HG)

Hair peg
(HP)

Bulbous hair peg (BHP)

Invasion of dermal cells



Epidermal cells lining the follicular canal

 Inner root sheath (IRS) Outer root sheath (ORS)

Dermal root sheath (DS)

↓ associated with the hair follicle

Arrector pili muscle

 eyebrows

 eyelashes

DP: Dermal papilla

GM: Germinal matrix: producing the hair shaft

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Hair follicle development

Formation of:

Arrector pili muscle (APM)

Stem cells=

hair follicle bulge ★★

Germinal matrix

Inner and outer root sheaths

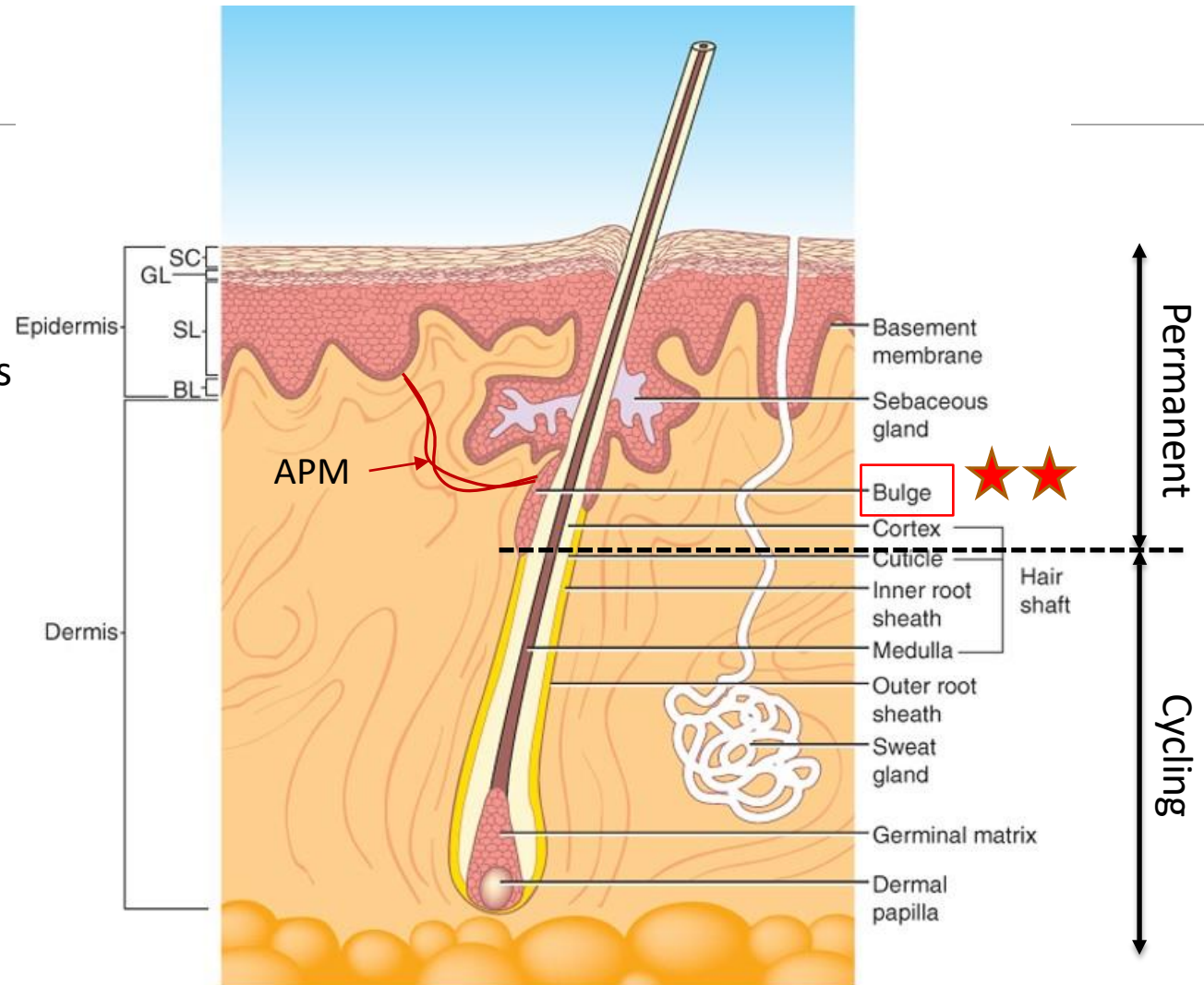
12th week

Babies are born with first generation of fine unpigmented hairs:

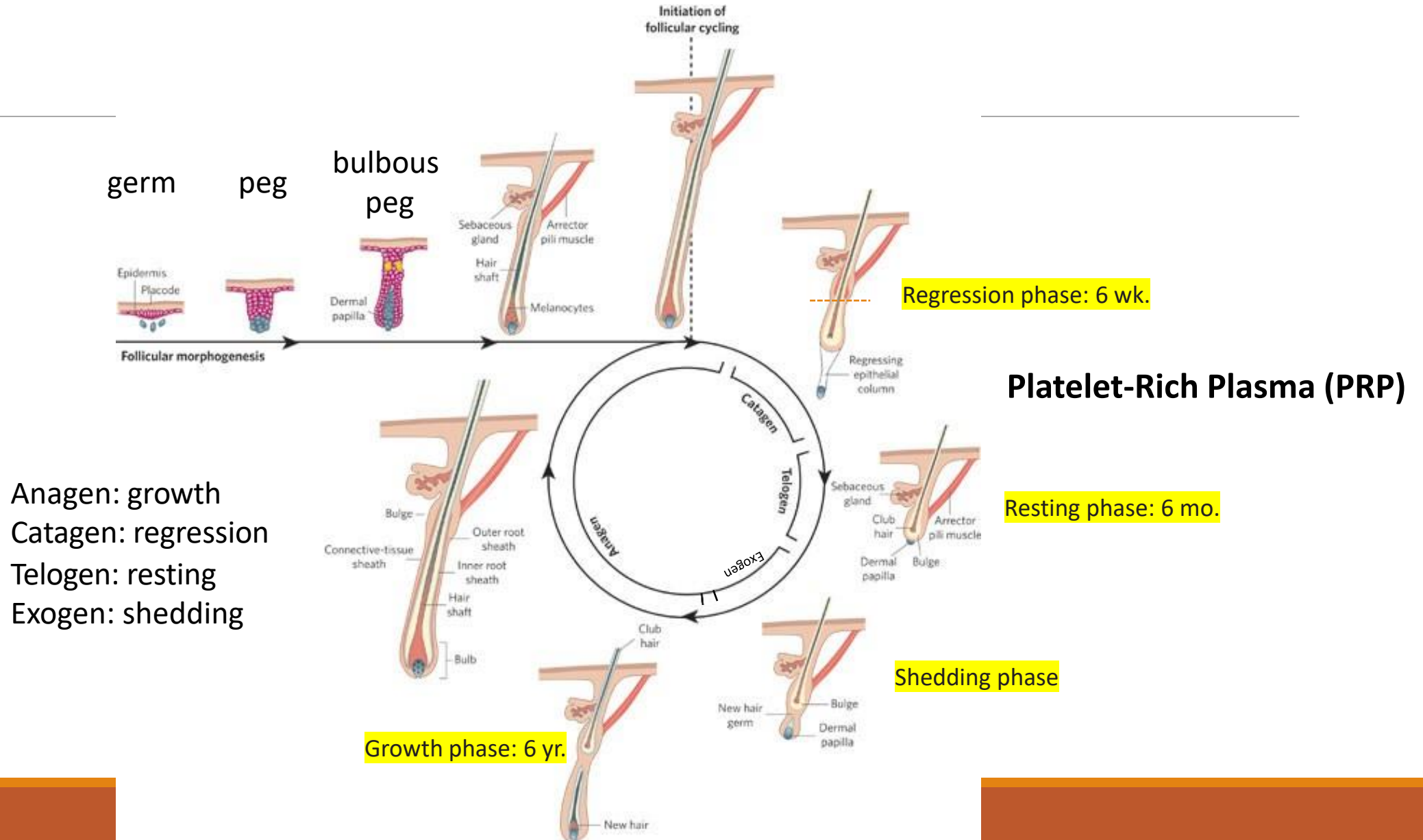
lanugo ★

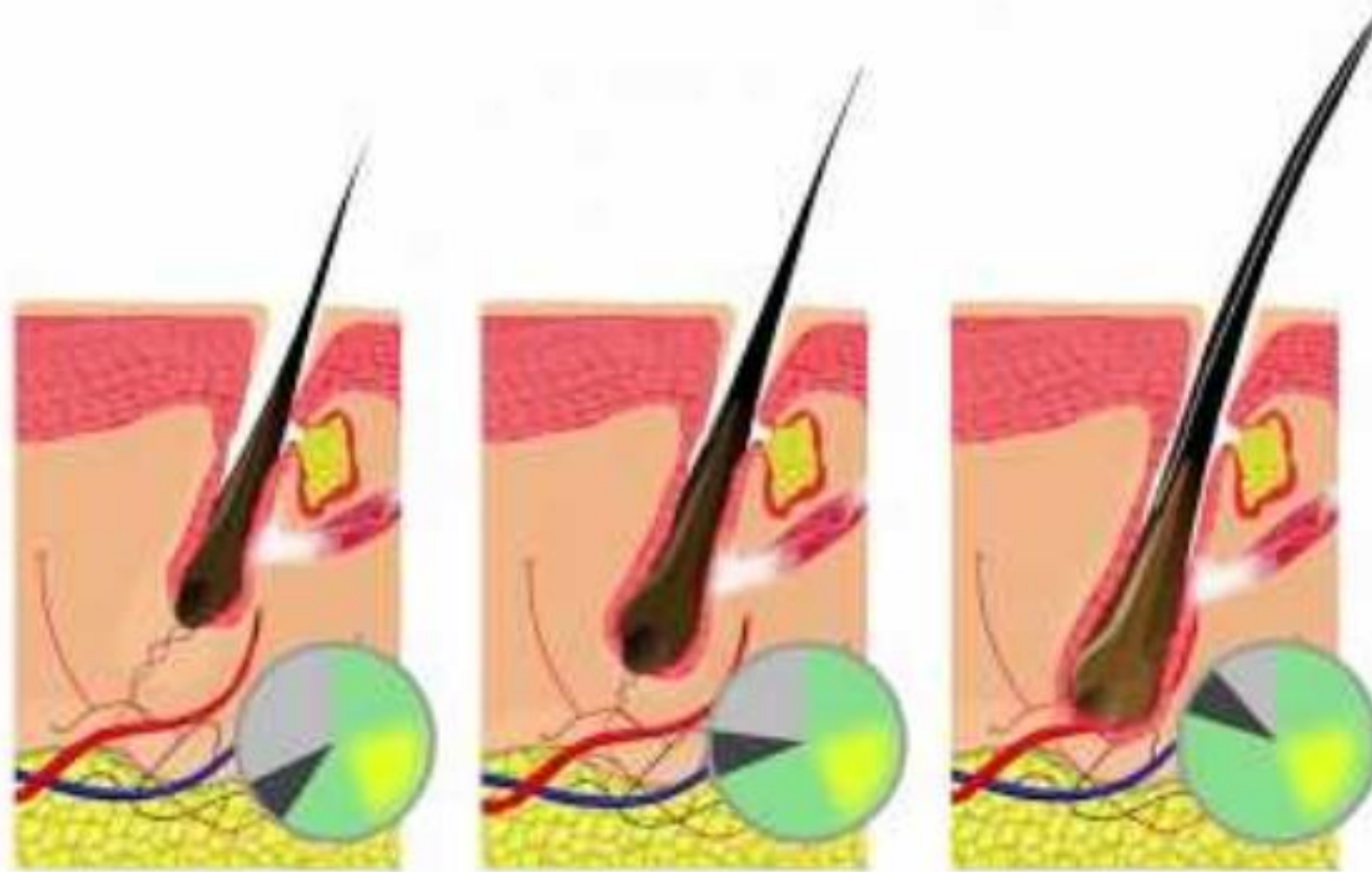
Postnatal regeneration:
Hair follicle cycling

Vellus & terminal hairs



Postnatal hair follicle cycling

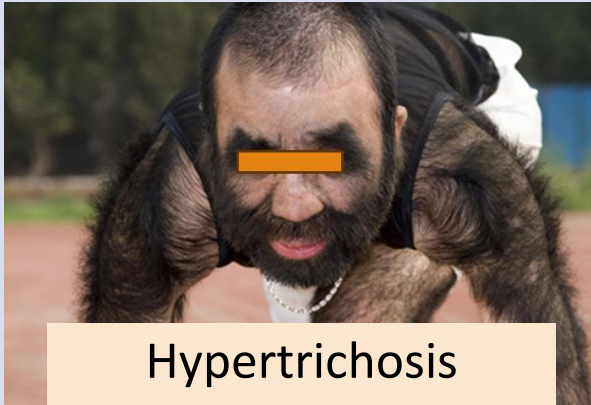




Abnormalities of Hair Distribution

Hypertrichosis (excessive hairiness) is caused by an unusual abundance of hair follicles. It may be localized to certain areas of the body, especially the lower lumbar region covering a spina bifida occulta defect or may cover the entire body (Fig. 21.4).

Atrichia, the congenital absence of hair, is usually associated with abnormalities of other ectodermal derivatives, such as teeth and nails.



Hypertrichosis



Hypotrichosis or Alopecia or Atrichia



Figure 21.4 Child with hypertrichosis.

Epidermal gland development

Epidermal appendages: outgrowths from E or HF



Sebaceous glands: sebum/vernix



Sweat glands



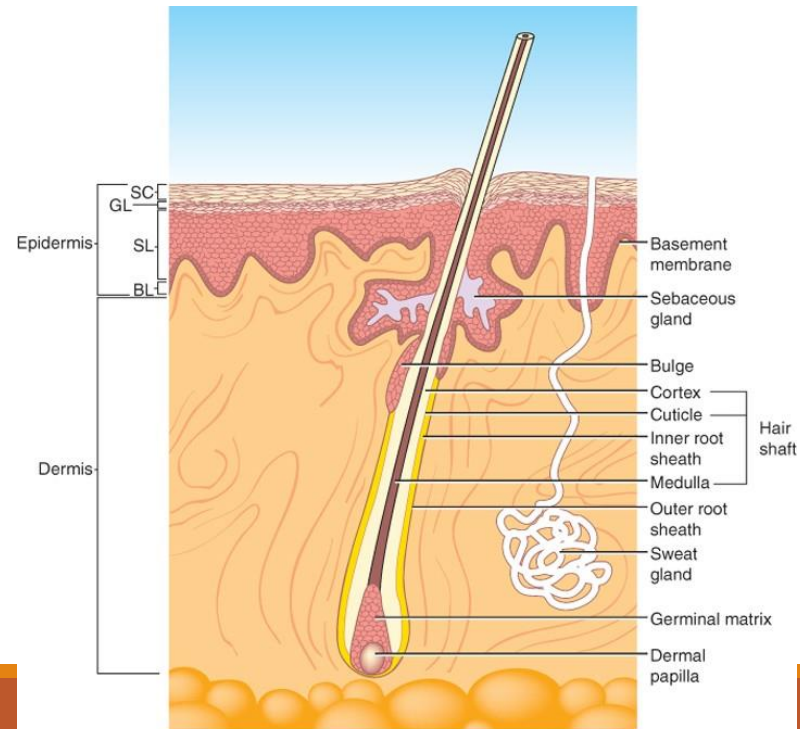
Apocrine glands (pheromones)



Mammary glands

Salivary glands

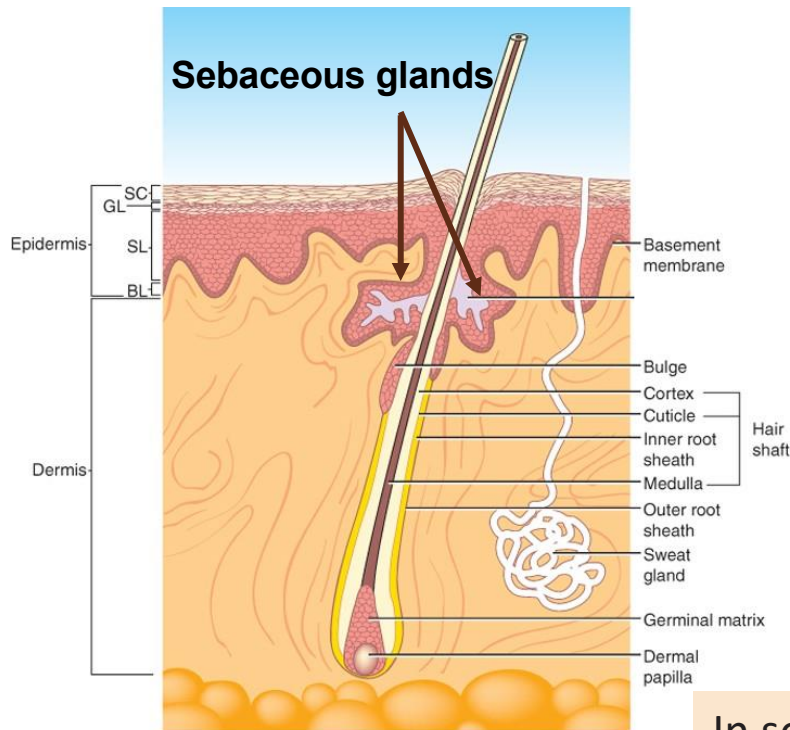
Lacrimal glands



Epidermal gland development

Epidermal appendages

Sebaceous glands

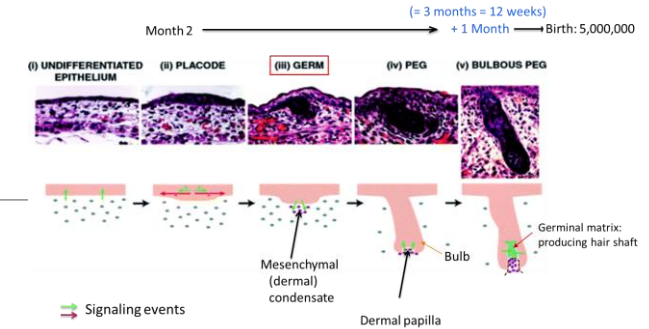


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- Develop from hair follicle (+4 weeks):
- Mature by 6 months: face
- Highly active in the fetus

(waterproof protective coating)

★ **vernix caseosa** = **sebum** + desquamating epidermal cells + remnants of the periderm



In some areas of hairless skin—such as the **glans penis** of males and the **labia minora** of females—sebaceous glands **develop as independent** down growths of epidermis.



★ **vernix caseosa** = **sebum** + desquamating epidermal cells + remnants of the periderm

After **birth**, the sebaceous glands become relatively **inactive**, but at **puberty** they again begin to secrete **large quantities** of sebum in response to the **surge** in circulating **sex steroids**.



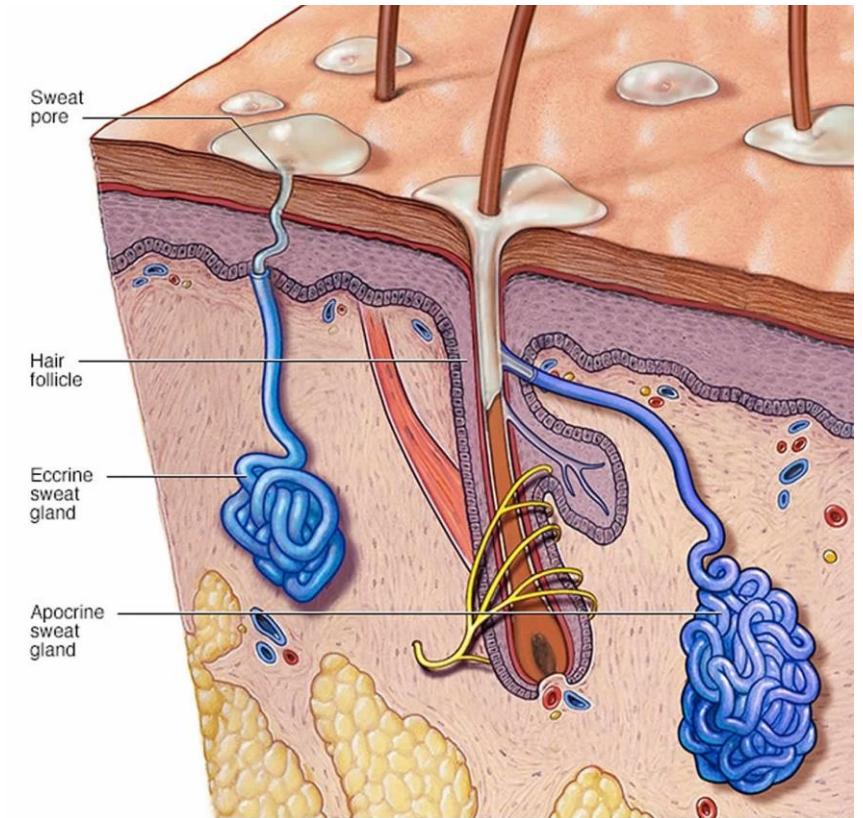
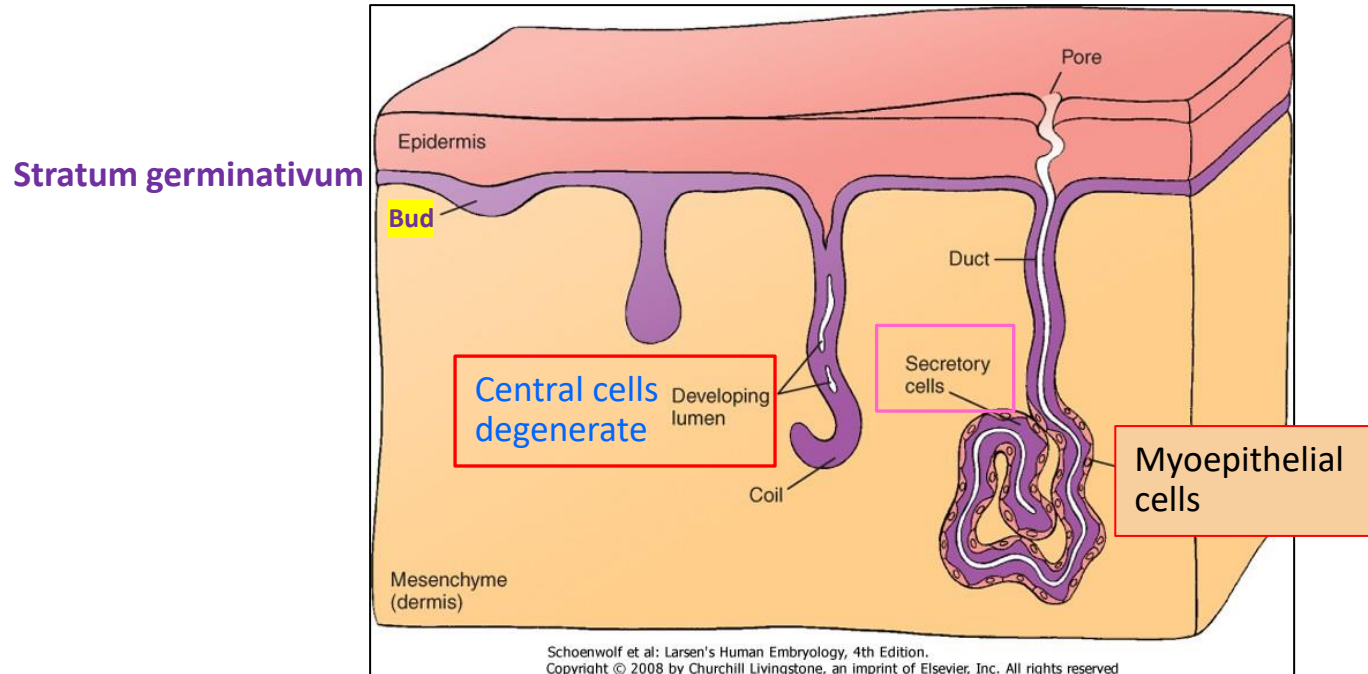
Milia (baby acne) are common in all ages and both sexes. They most often arise on the face, and are particularly prominent on the eyelids and cheeks, but they may occur elsewhere.

Epidermal gland development

Epidermal appendages

Sweat glands (eccrine): develop from **stratum germinativum (20 weeks+)**

Apocrine sweat glands: develop in association with HFs, **most lost** late in fetal development
except the axillae, mons pubis, prepuce, scrotum, and labia minora



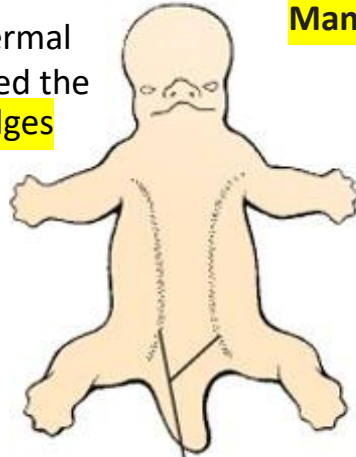
Epidermal gland development

Epidermal appendages

Mammary gland

A pair of epidermal thickenings called the mammary ridges

Mammary ridges

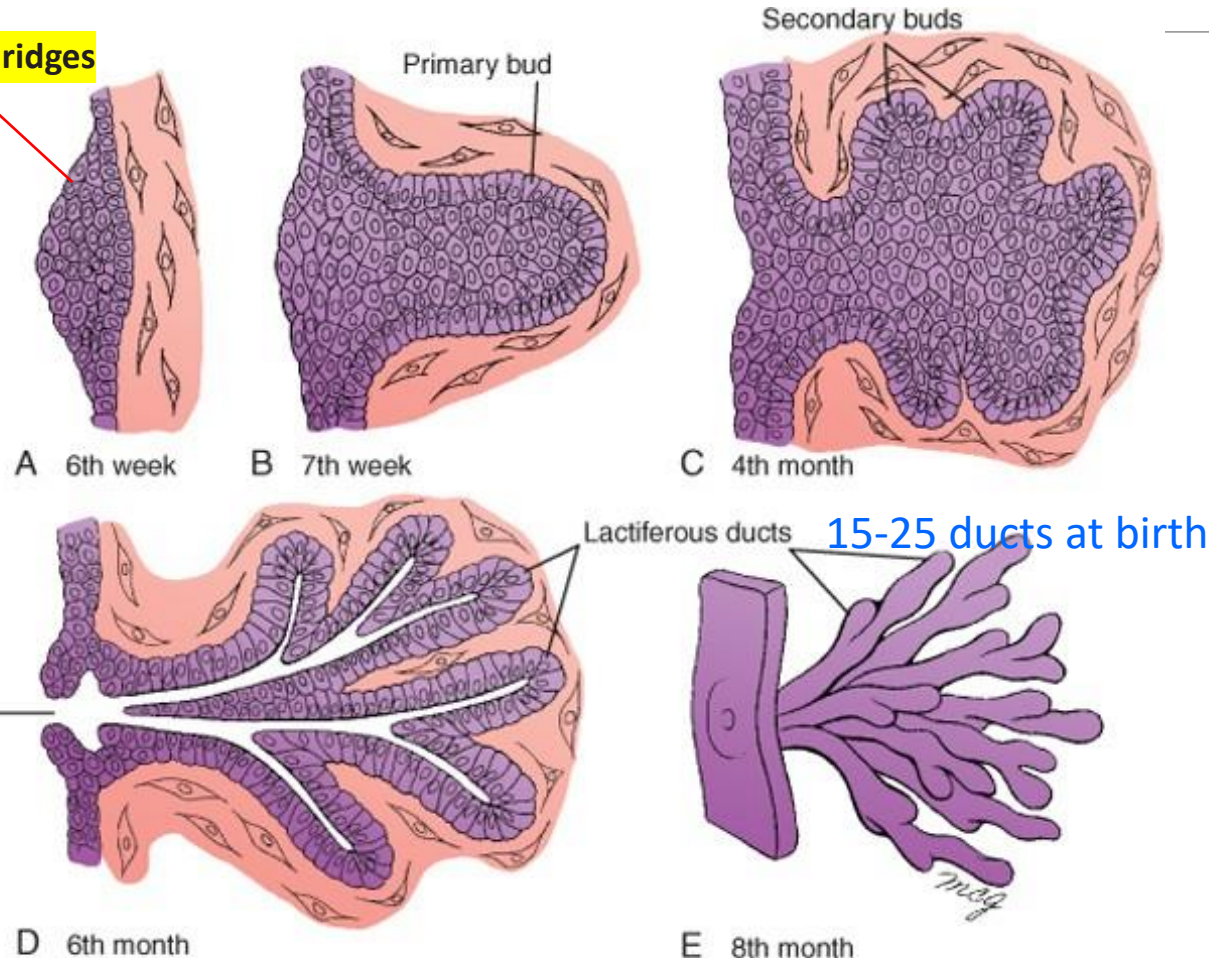


Mammary ridges

4th week

inverted nipple

หัวนมบอด



15-25 ducts at birth

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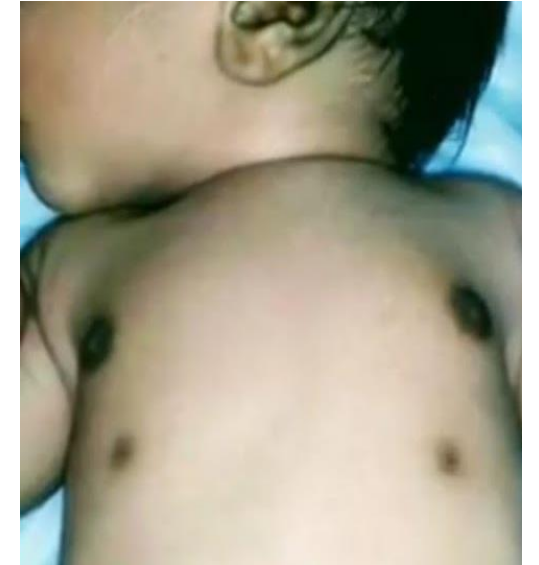
Postnatal regenerative growth



Polymastia:
supernumerary breasts



Polythelia:
supernumerary nipples

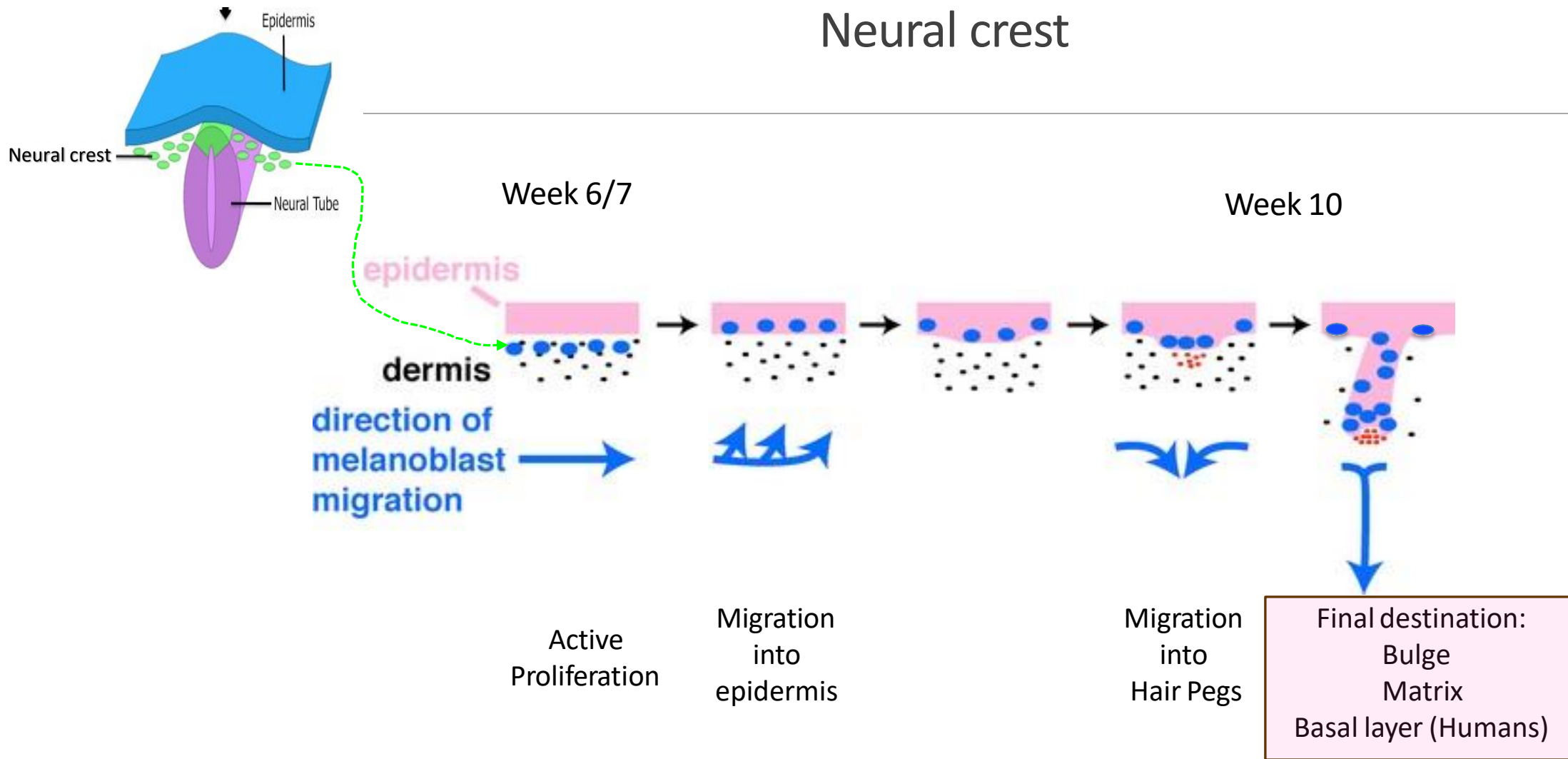


Melanocytes



Melanocyte development

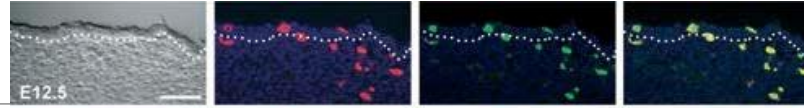
Neural crest



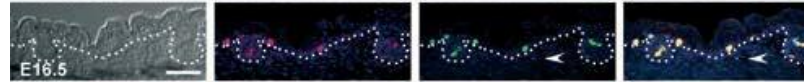
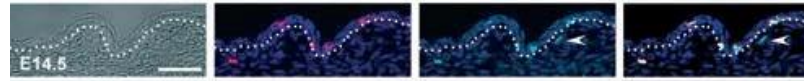
Melanocyte development

Neural crest

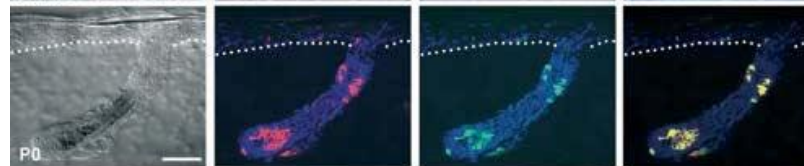
Before stratification



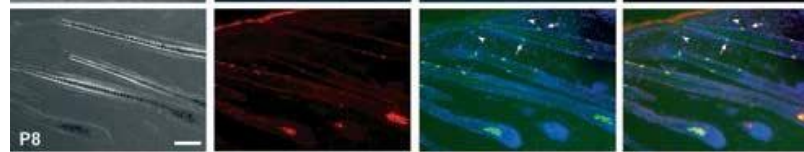
Onset stratification



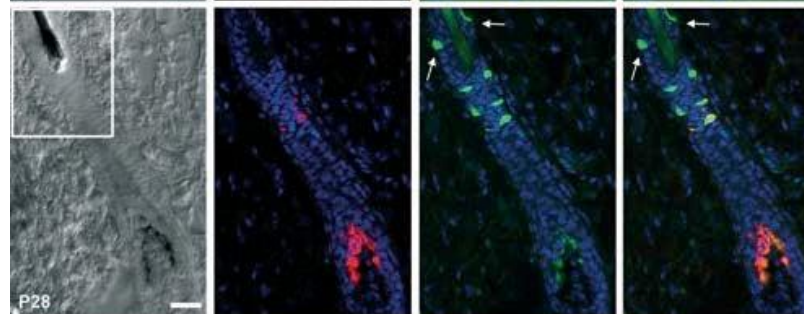
Bulbous hair pegs



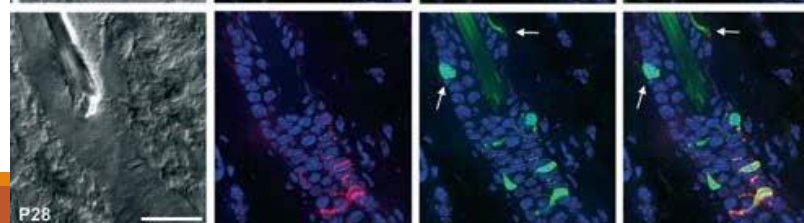
Anagen



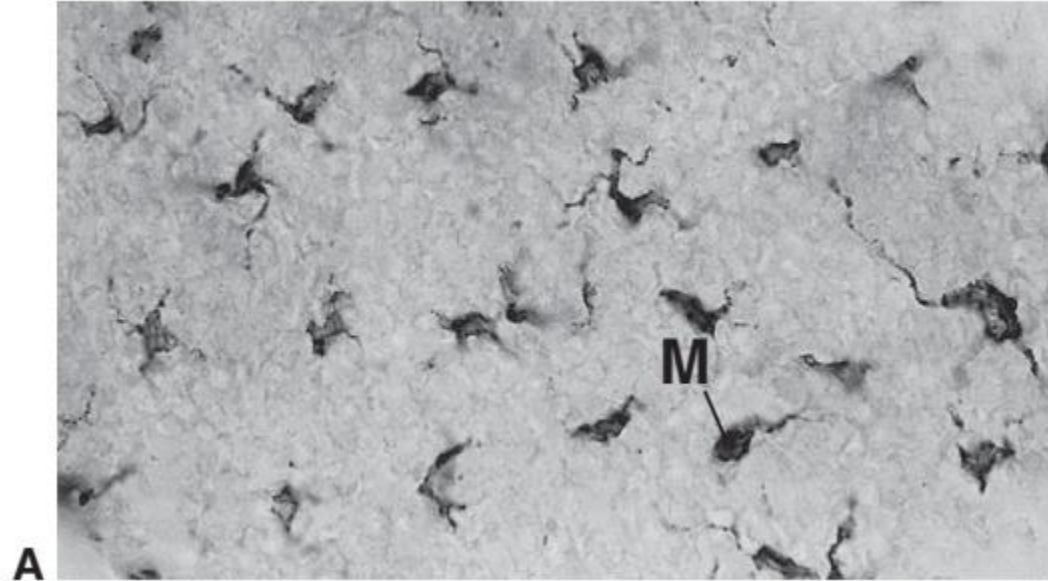
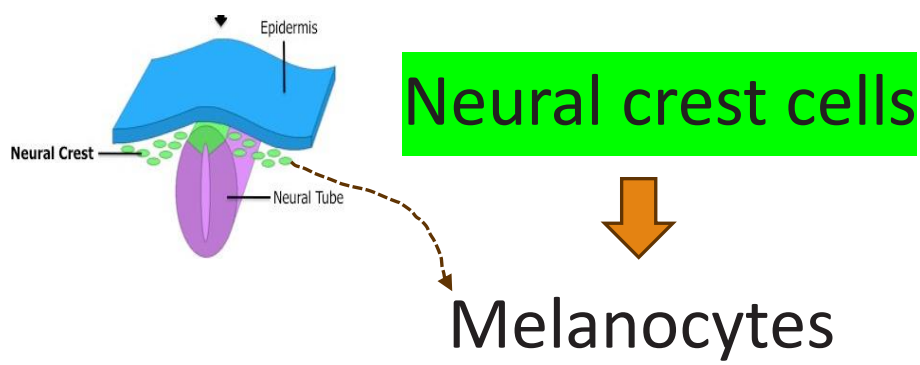
Matrix



Hair follicle Bulge



Djian-Zaouche,
Pigment Cell Melanoma Res 2012



6th to 7th weeks

(studies using monoclonal antibodies directed against antigens characteristic of **melanocyte precursors**)

Melanocytes represent between 5% and 10% of the cells of the epidermis in the adult.

Melanocytes function as a sunscreen

- ⊘ Sunburn
- ⊘ Cancer

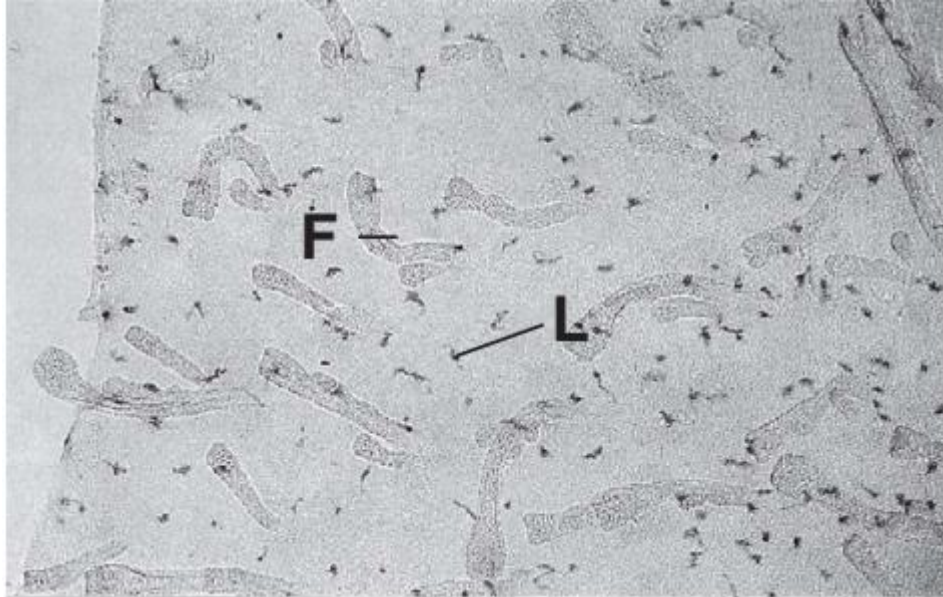
Melanoma



Bone marrow



Langerhans cells (macrophage)



7th week



throughout life

Langerhans cells function as the **immune cells** of the skin

⊘ contact sensitivity (allergic skin reactions)

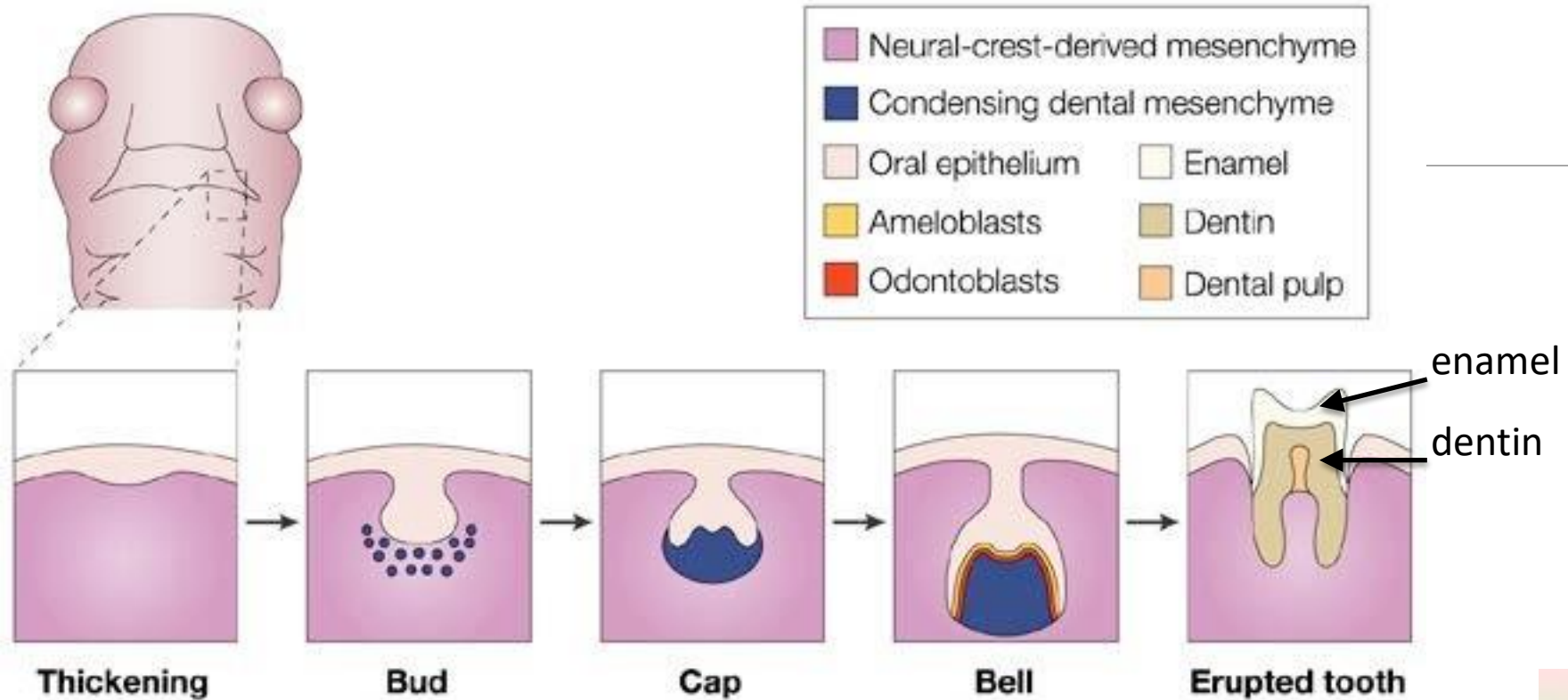
in immune surveillance

⊘ against invading microorganisms

Teeth



Tooth development

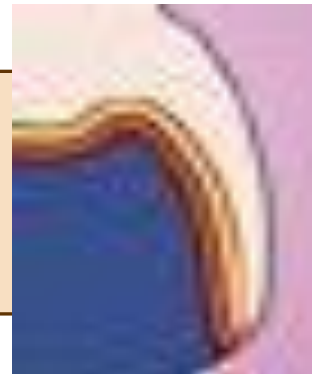


Week 6:
dental lamina
U-shaped ridge
of epidermis

Week 7:
Mesenchyme
condensation
(20)

Week 8:
Mesenchyme
invades base
tooth bud:
Dental papilla

Week 14:
❖ Epithelial cells: **Ameloblasts**: enamel
❖ **Neural crest cells**=Ectomesenchymal cells
= **Odontoblasts**: dentin



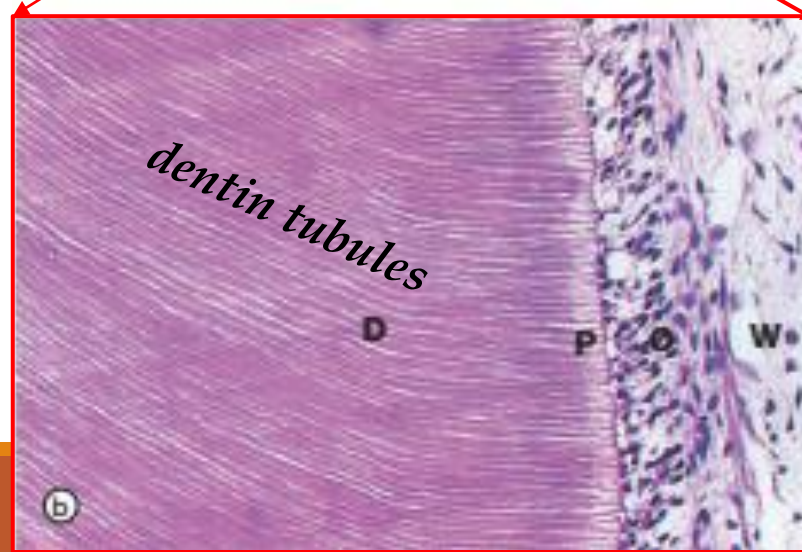
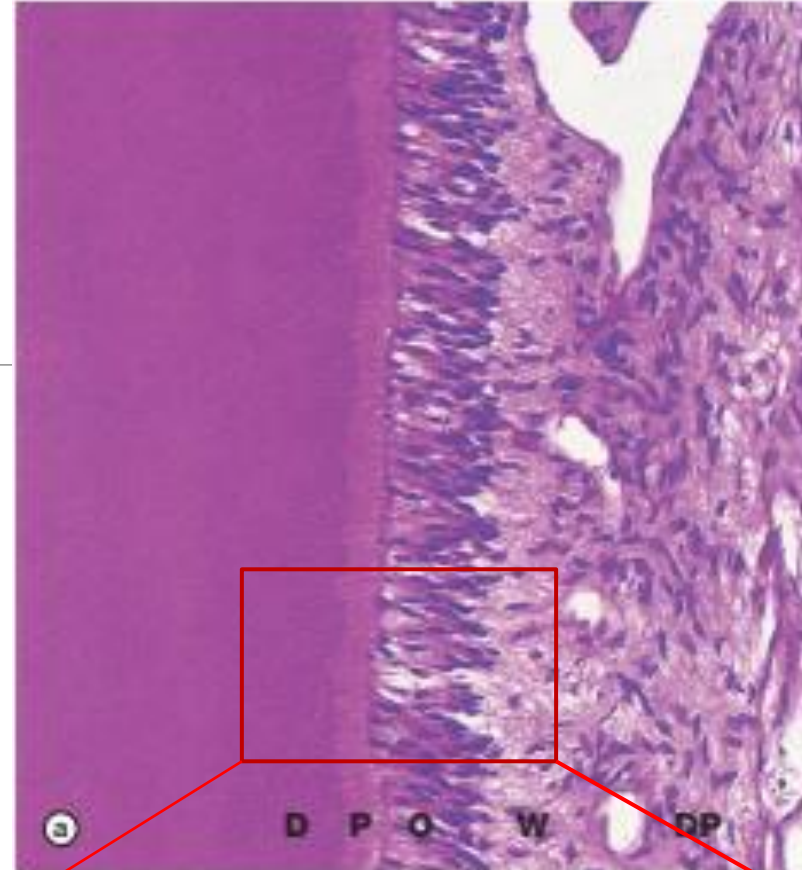
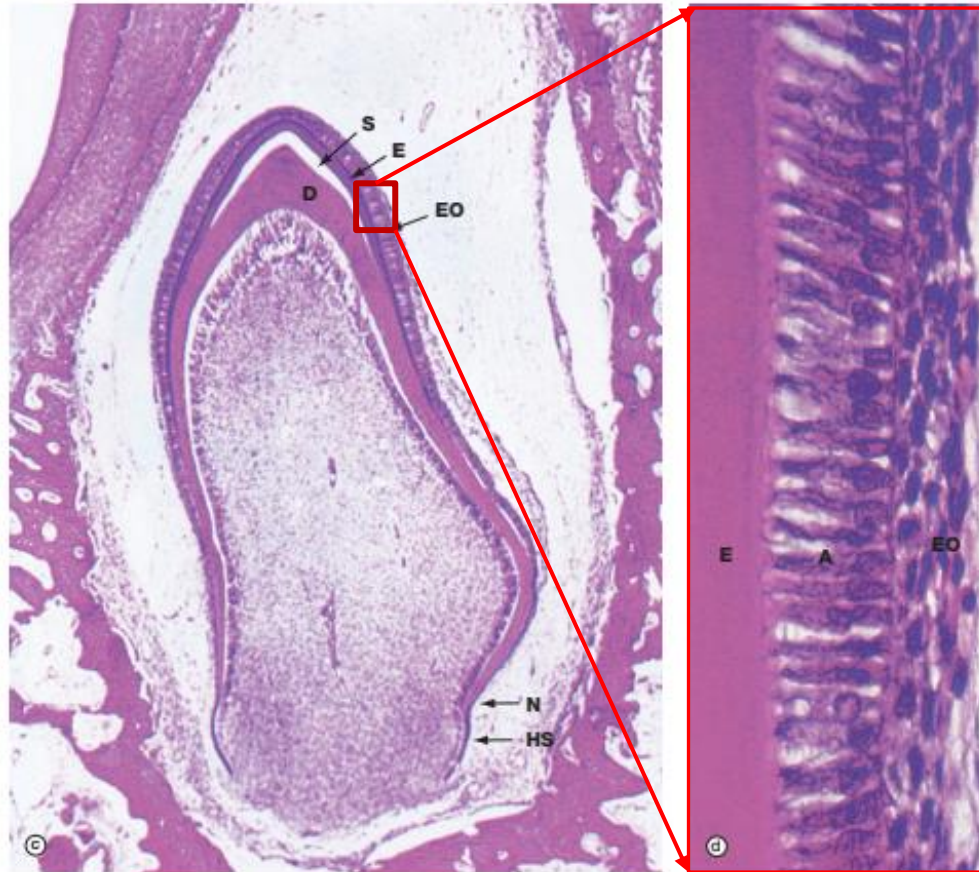
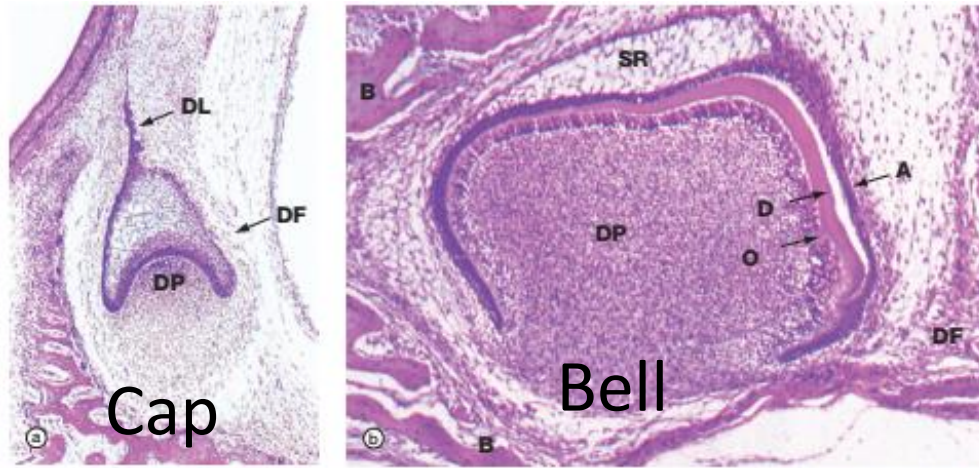
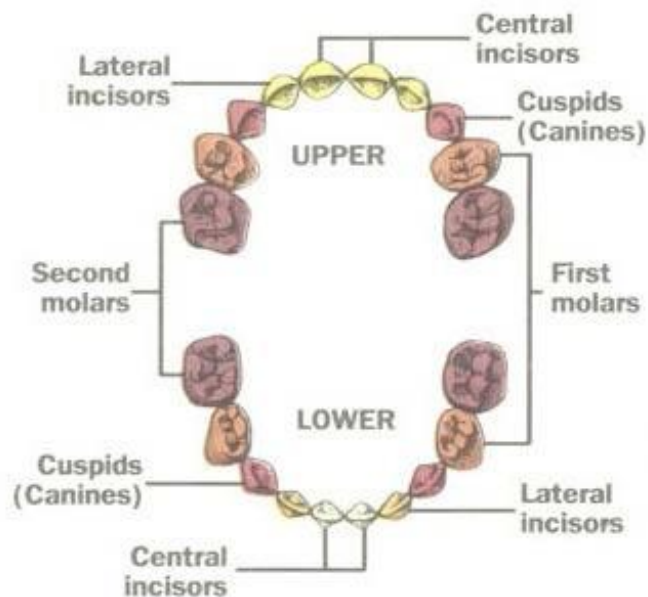


FIG. 13.5 Tooth development (caption opposite)
 (a) H&E, cap stage (LP) (b) H&E, bell stage (MP) (c) H&E, onset of root development (LP) (d) H&E, ameloblasts (HP)

Tooth development

Primary teeth

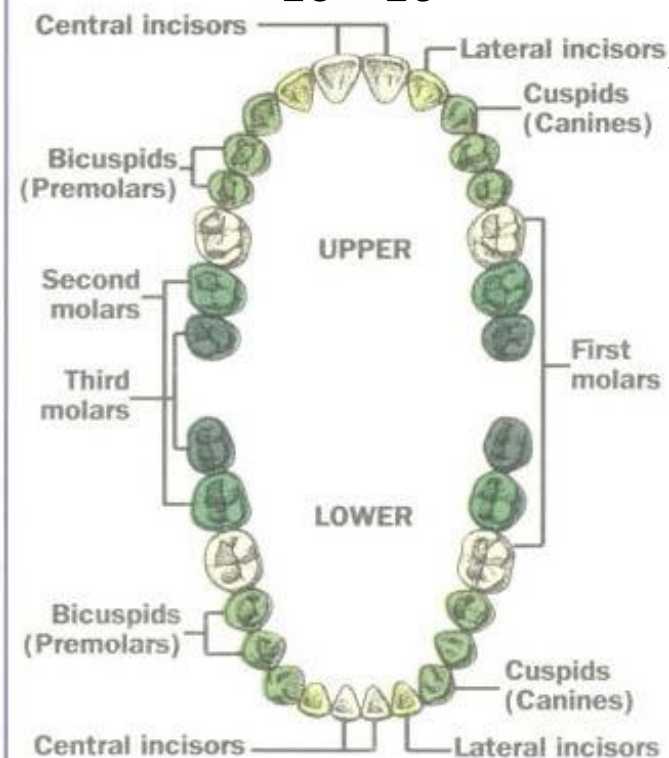
10 + 10



6-10 months	13-19 months
8-13 months	16-23 months
10-16 months	23-33 months

Adult teeth

16 + 16



6-8 years	11-13 years
7-9 years	17-21 years
9-12 years	

Nails



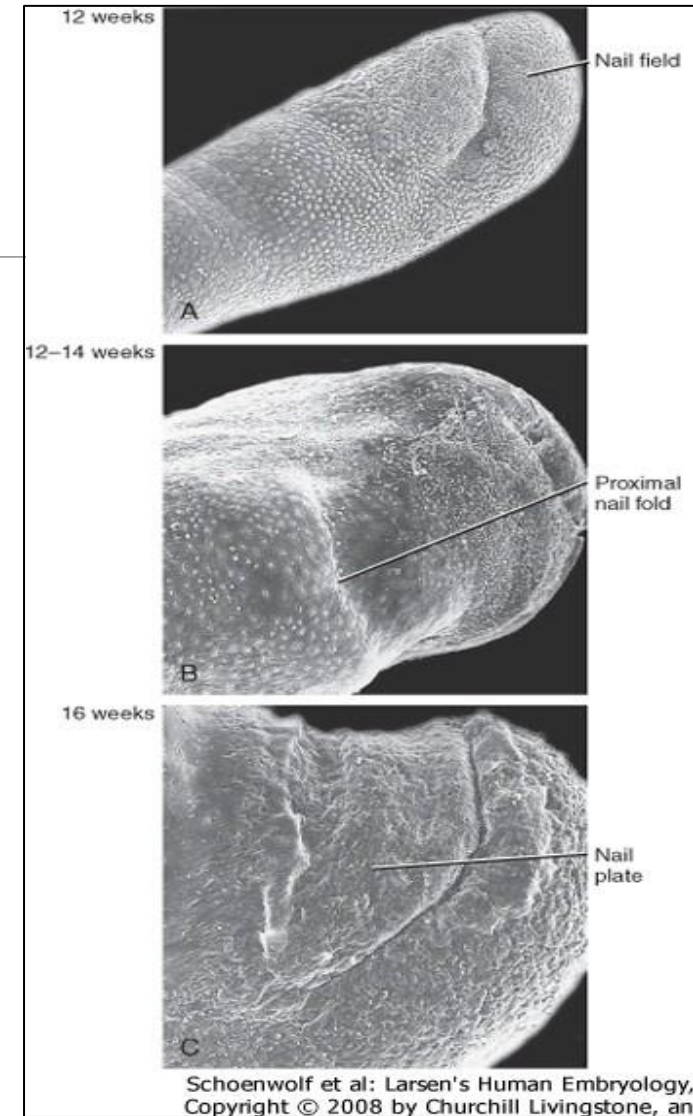
Nail development

Ectoderm

Week 10: epidermal thickening, **nail field**

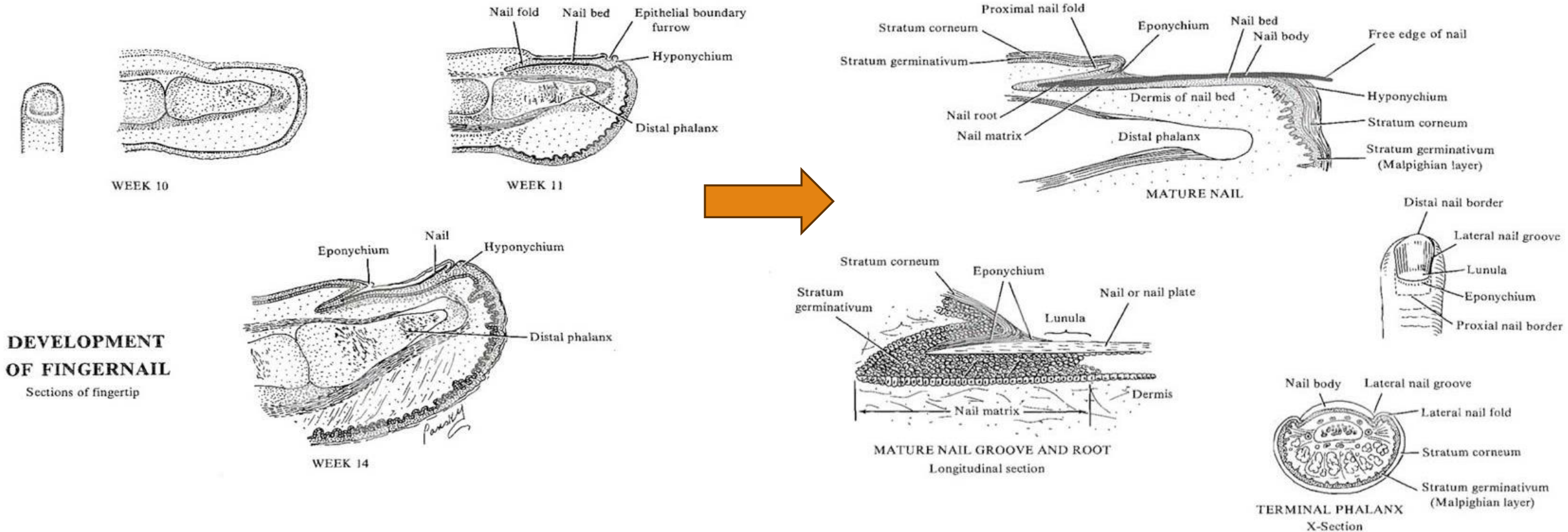
Week 11: nail field **moves proximally**, nail folds, proximal nail fold: formation of formative zone

3rd – 9th months



Nail development

Ectoderm



X-linked genetic disorder



Ectodermal Dysplasia

- **Skin**: dry, thin, or prone to infections due to abnormalities in sweat glands.
- **Hair**: sparse, brittle, or absent (hypotrichosis or alopecia).
- **Nails**: thin, brittle, or misshapen.
- **Teeth**: hypodontia (fewer teeth) or anodontia (absence of teeth). Teeth may also be small, pointed, or misshapen.
- **Sweat Glands**: Reduction or absence of sweat glands can lead to heat intolerance and difficulty regulating body temperature.

Lecture conclusion

Skin

Skin origins

Development of the overlying epidermis

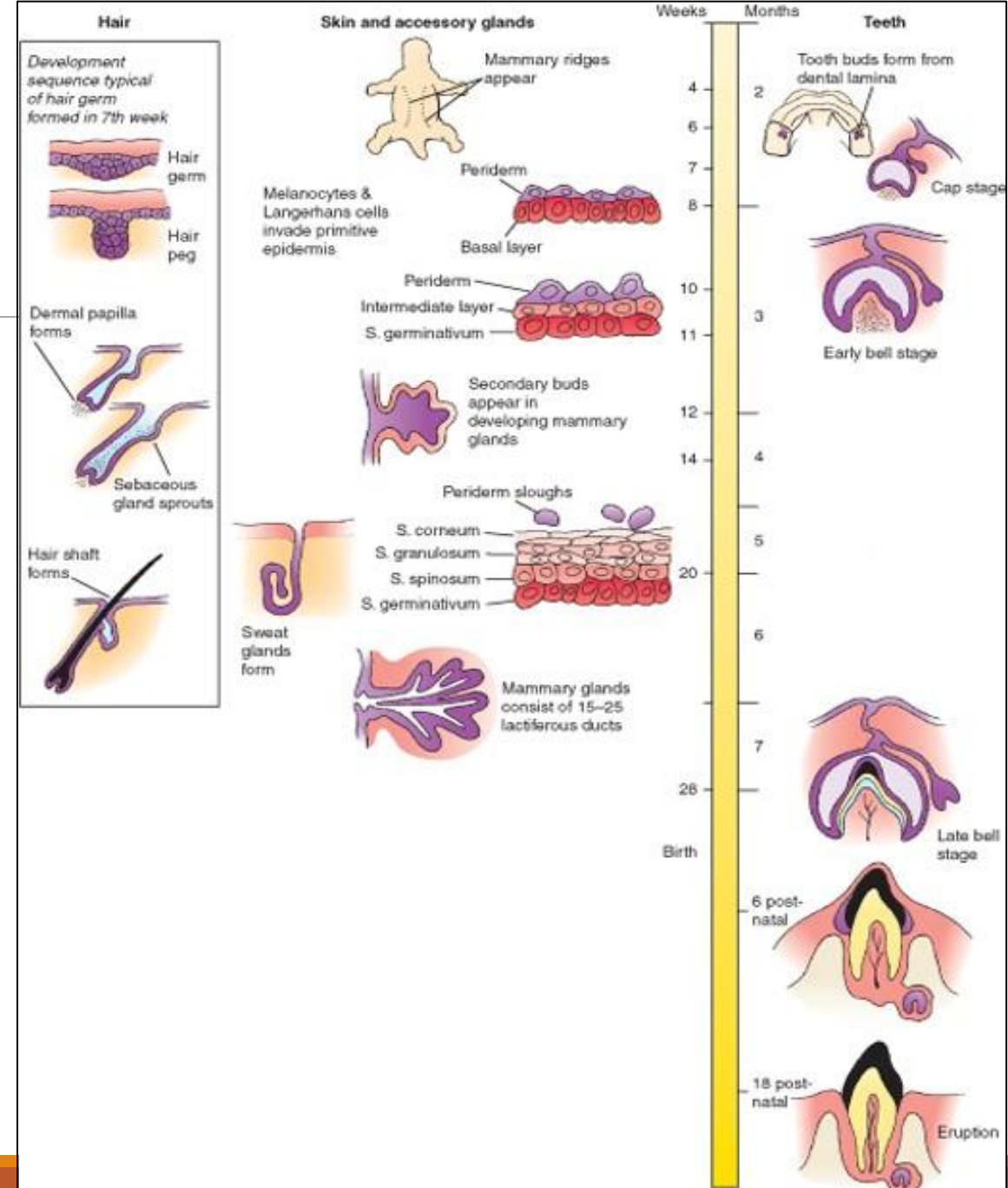
Development of epidermal appendages:

Hair follicles Glands
Mammary glands Nails
Teeth

Development of Melanocytes

Development of Langerhans cells

Development of the Dermis



Larsen's Human Embryology

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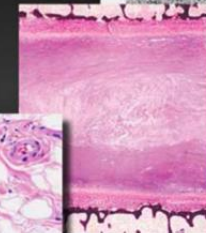
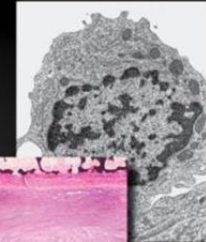
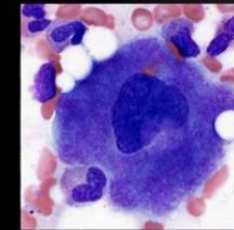
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