The normal cervix: cytology, colposcopy, and histology

Alan G. Waxman, MD, MPH
University of New Mexico School of Medicine
Department of Obstetrics and Gynecology
Albuquerque, New Mexico



Disclosures

- Alan G. Waxman, MD, MPH Faculty No Disclosures
- Please see www.asccp.org/CompOnlineCME for full program disclosures

Images used with permission:

- Apgar B, Brotzman G, Spitzer M. Integrated Colposcopy: A Text and Atlas. Elsevier; 2002, 2008. (ABS)
- Ferris D, Cox T, O'Connor D, Wright C. Modern Colposcopy. Wolters Kluwer, ASCCP; 2002
- Personal collections as noted on slides





Objectives

- Review the epithelial features of the normal transformation zone
- Discuss the process of squamous metaplasia
- Explain the normal features of the transformation zone according to age
- Describe features that define a satisfactory colposcopy



Origin of cervical epithelium

- Vagina is originally lined by <u>columnar epithelium</u> derived from fusion of the *Mullerian ducts*
 - Lines the endocervix and is continuous with the endometrium
- Gradually replaced by a core of <u>stratified squamous</u> epithelium originating in the *urogenital sinus*
 - Lines the vagina, portio vaginalis of the cervix
- Rudimentary cervix by 16 weeks gestation



Types of cervical epithelia

Stratified squamous

- Covers the ectocervix and vagina
- Multilayered epithelium, rests on basement membrane
- ***Smooth pink color

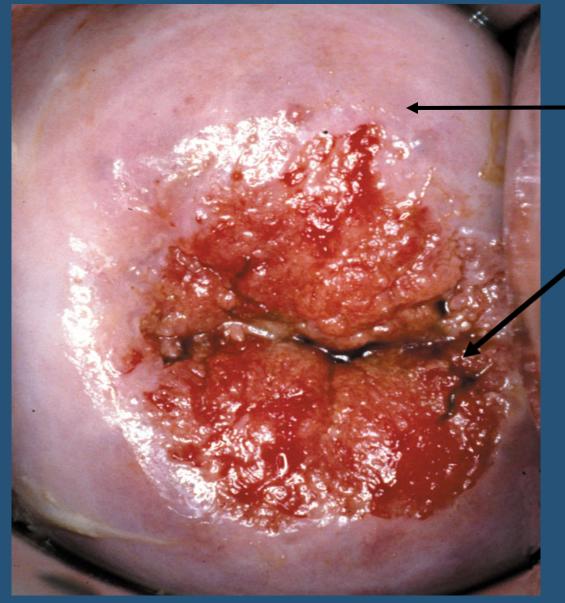
Tall columnar

- Lines the endocervix
- One cell layer thick, mucin-secreting, numerous folds
- ***Bright red irregular color

Metaplasia: immature and mature

- Lies between columnar and squamous epithelium
- ***Faint acetowhite color after vinegar application

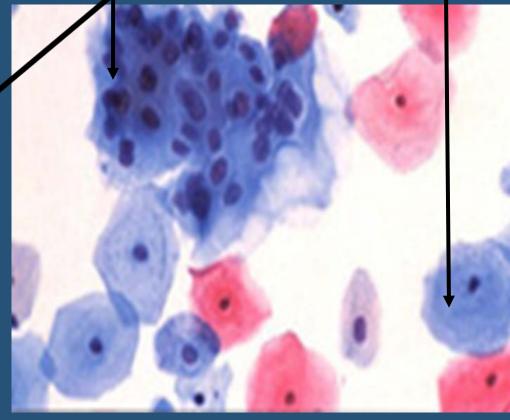




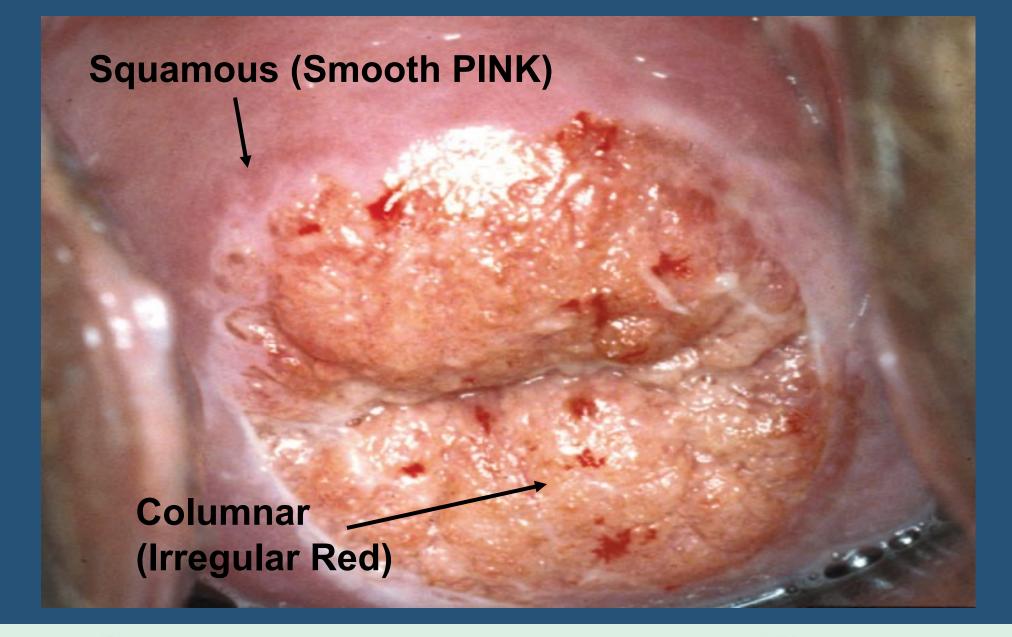
2 types of epithelium

Squamous

Columnar

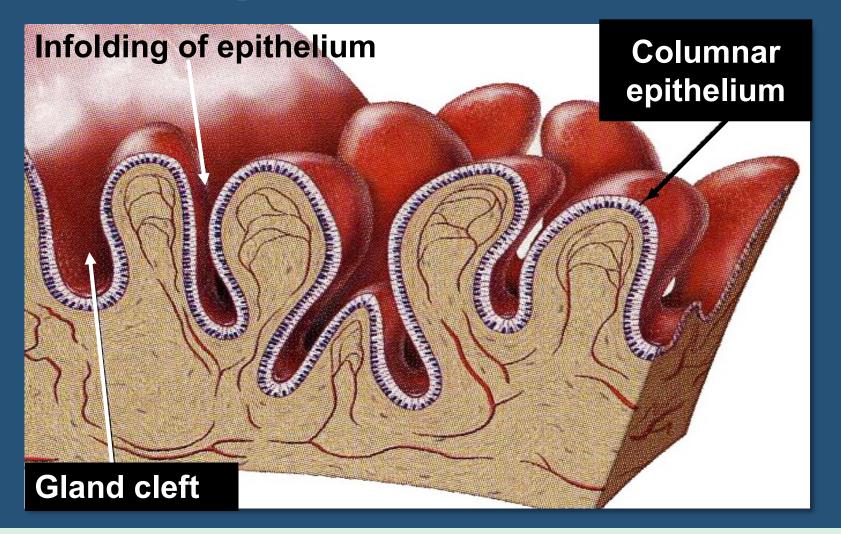








Columnar epithelium



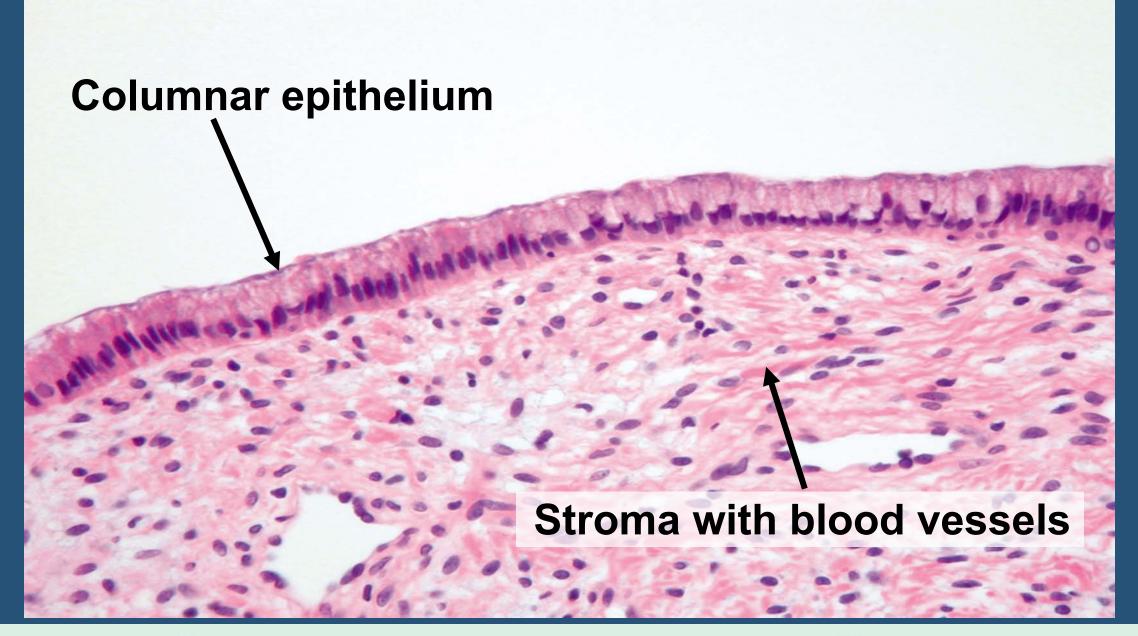


Columnar epithelium

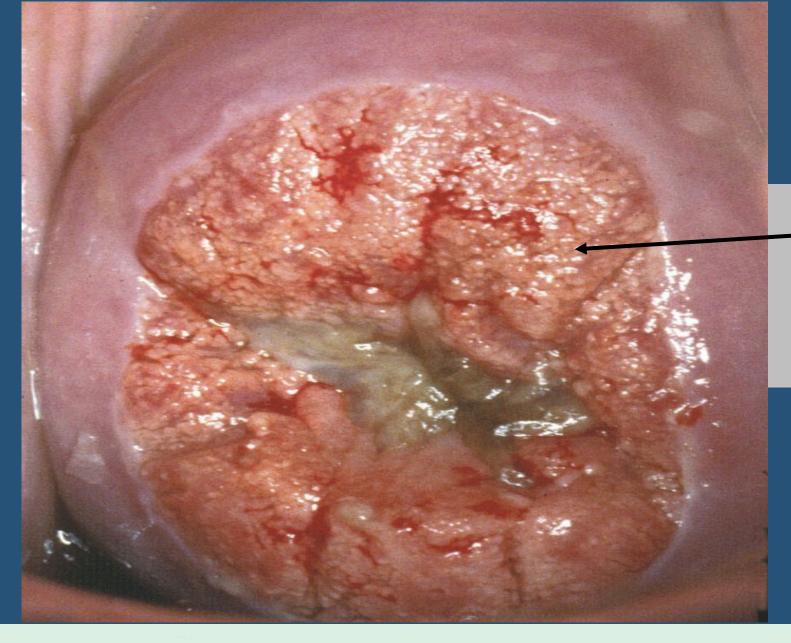
- Single layer tall columnar cells lining endocervical canal
 - Course proximally from SCJ through endocervical canal to internal os
 - Majority secrete mucus; may have cilia
- Covers villi that contain central loop capillaries
 - ***Vessels are poorly concealed: epithelium is "red" compared to squamous "pink"
- Endocervical cells invaginate into the stroma to depth of 5-8mm
 - Are called "glands" but are technically crypts
 - Infolding of epithelium creates texture on surface





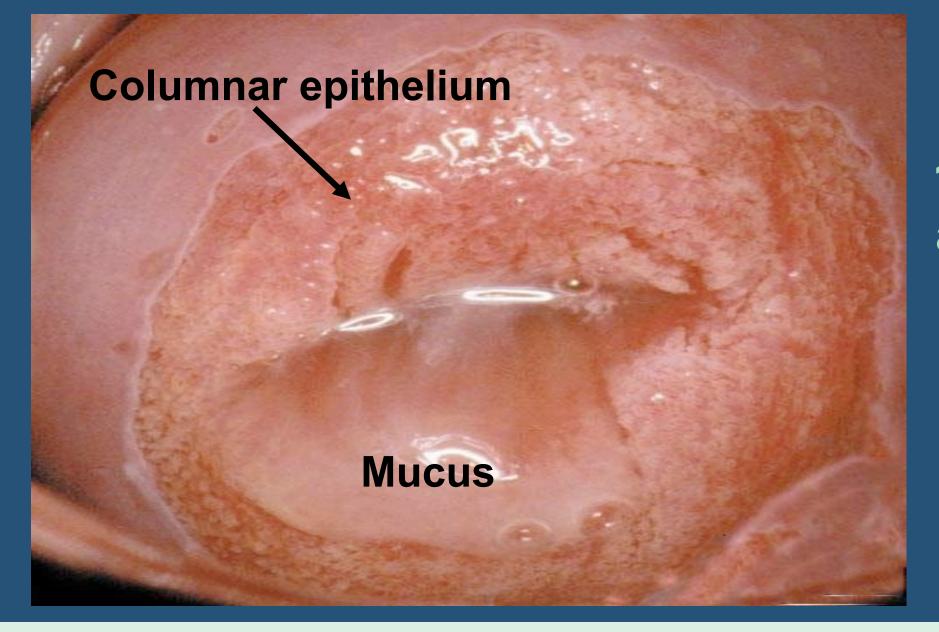






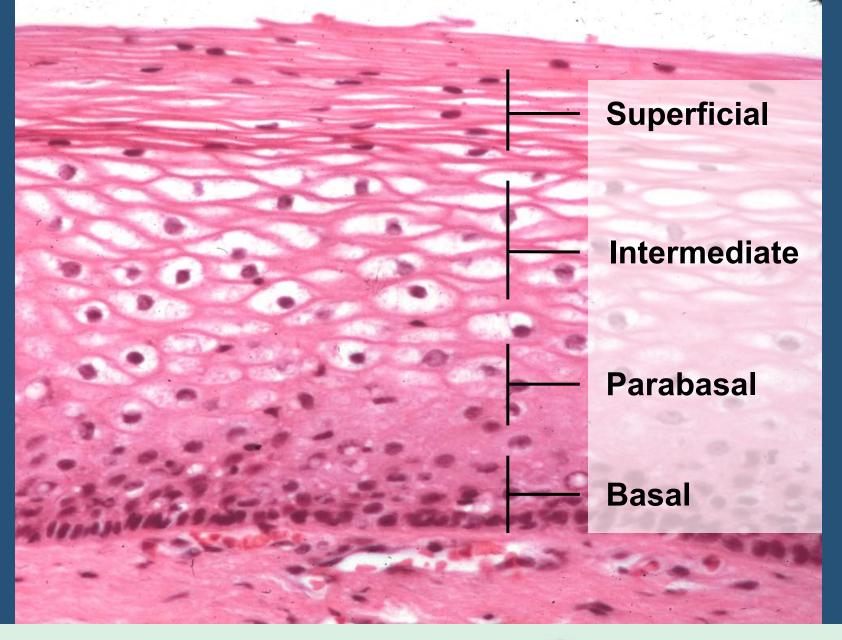
_Columnar epithelium

redder
than squamous



18-year-old adolescent

Stratified squamous epithelium divided into 4 layers



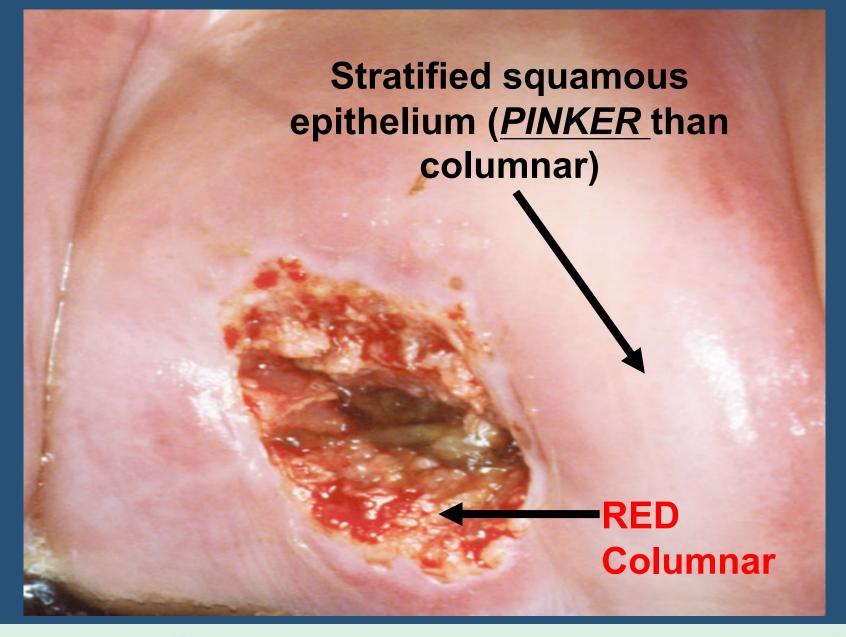


Stratified squamous epithelium

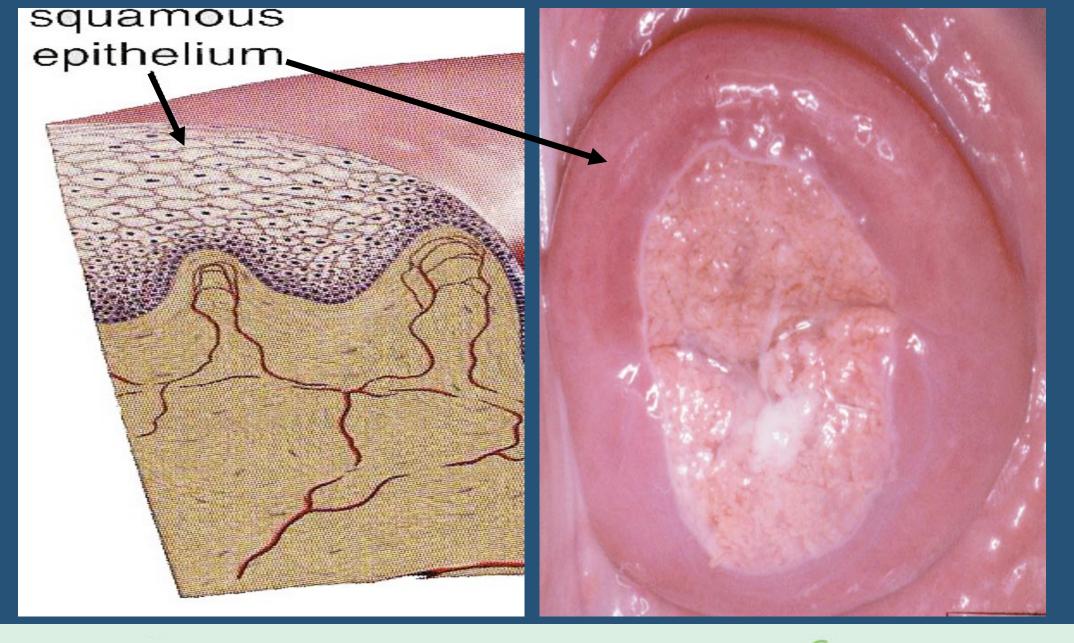
- Estrogen continuously remodels squamous epithelium in premenopausal women
 - Epithelial proliferation
 - Epithelial maturation
 - Epithelial desquamation
- Divided into layers
 - Basal / parabasal / intermediate / superficial
 - Regeneration from the basal layer under influence of estrogen



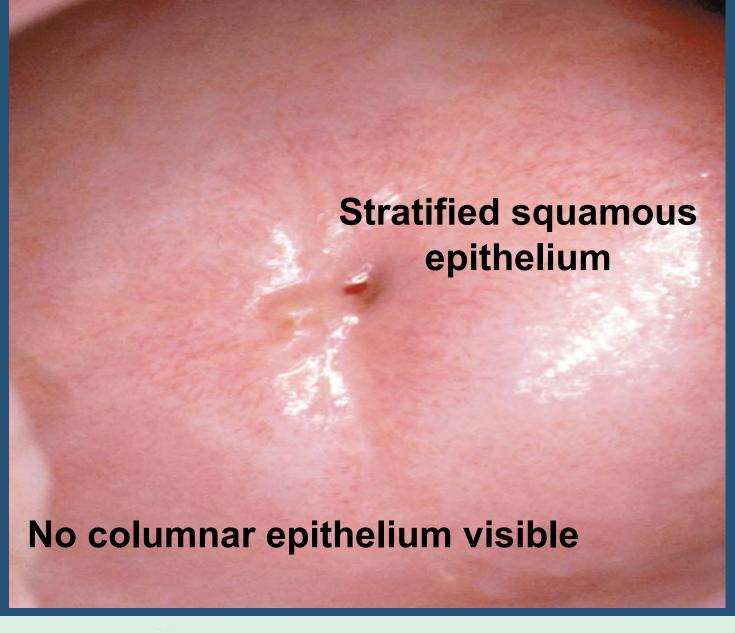




30-year-old Individual







65-year-old postmenopausal individual



Squamocolumnar junction (SCJ)

Interface between the columnar epithelium and squamous epithelium

- Original SCJ is defined at birth
 - Separates the original glycogenated squamous epithelium from the original columnar epithelium
 - Embryologically determined caudal extent of columnar epithelium
 - Cervical "transformation" begins here



Life cycle of the SCJ



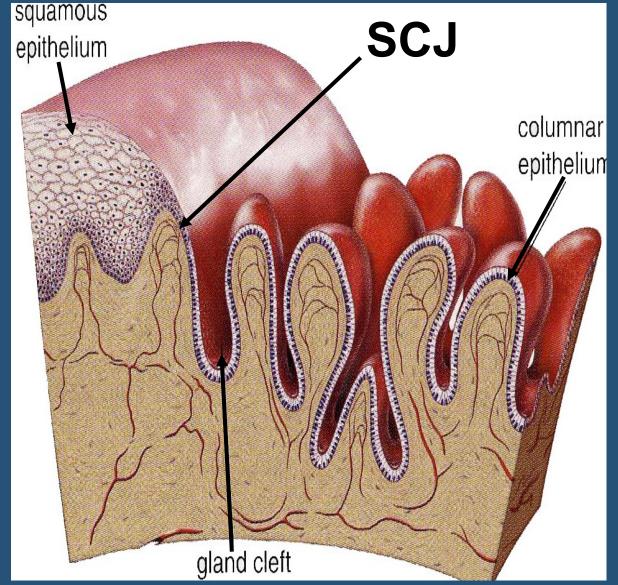
Adolescence



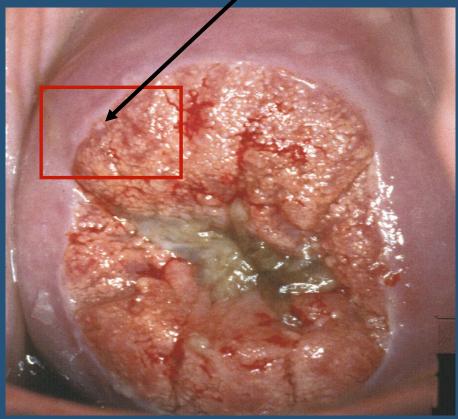
Reproductive years



Postmenopause

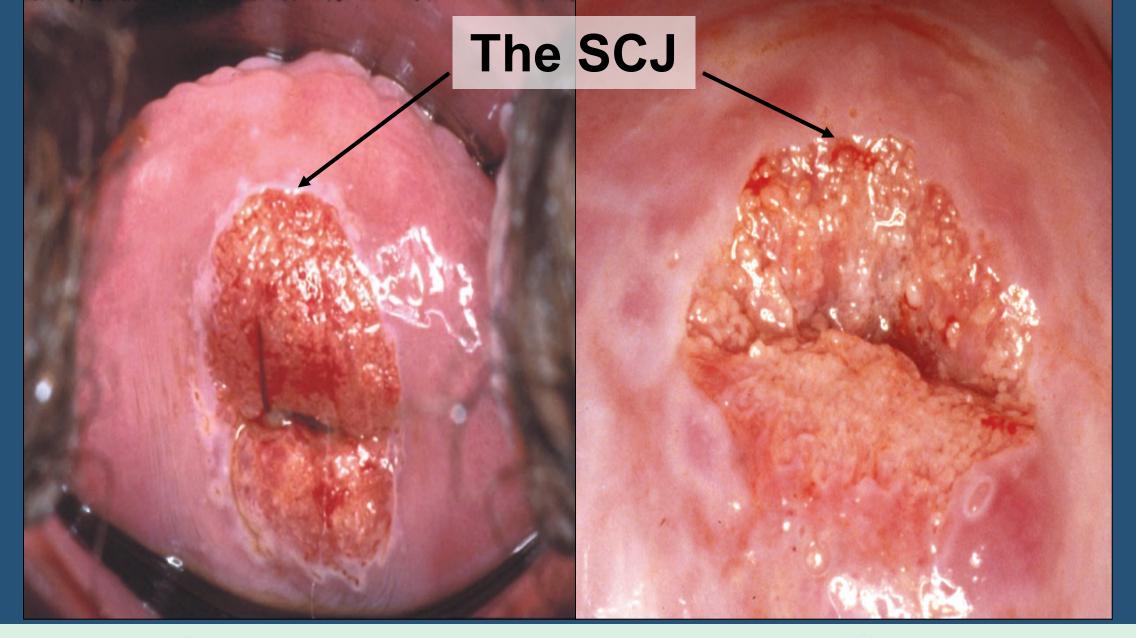


The SCJ



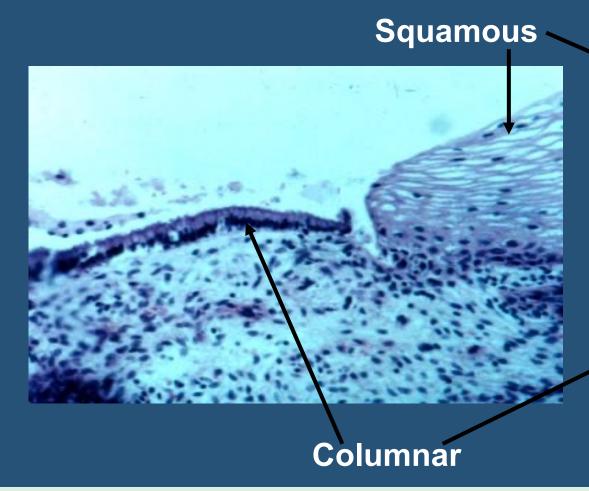
Mayeaux, E. J., & Cox, J. T. (2012). *Modern colposcopy: textbook & atlas*. Philadelphia: Wolters Kluwer Health/Lippincott Williams & Wilkins.

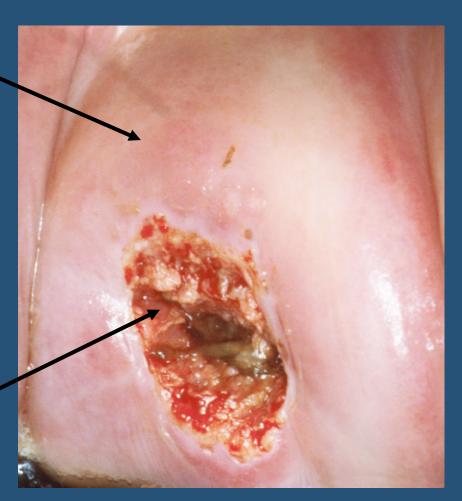






The evolution of the SCJ

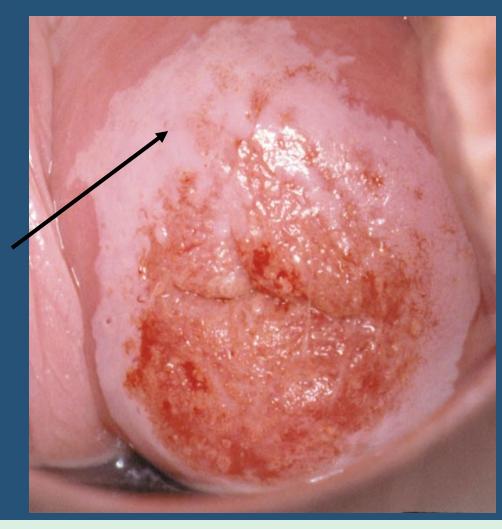






The SCJ "moves" centrally by the process of squamous metaplasia

- Metaplasia: transformation of one cell type into another
- Squamous metaplasia is the process of transformation of columnar into squamous epithelium
- Squamous metaplasia is a normal process of cervical maturation

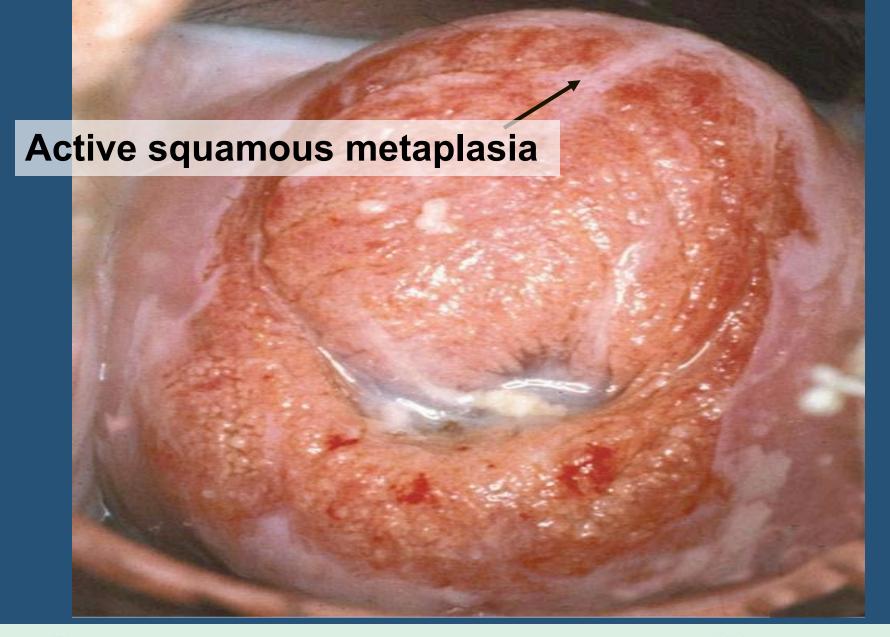


Mechanism of squamous metaplasia

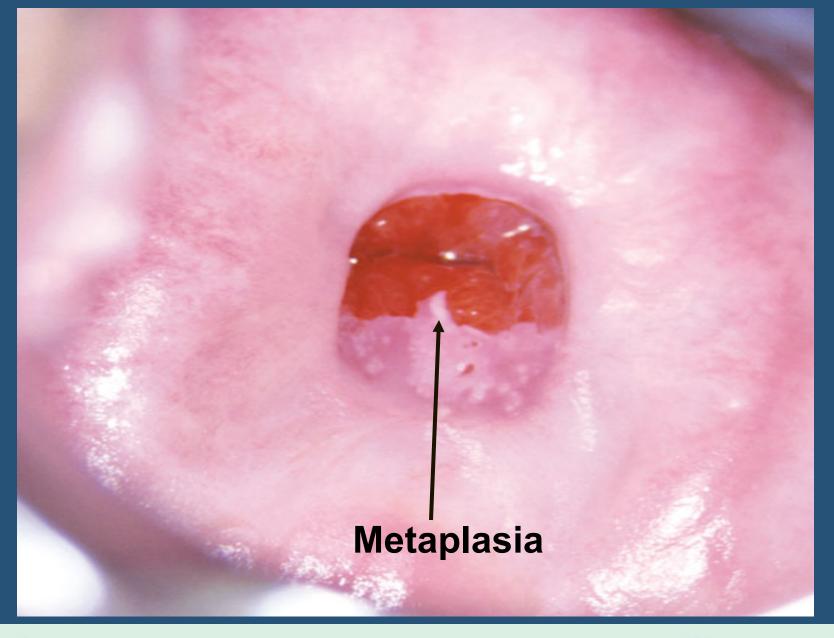
- Columnar cells changed into immature metaplastic cells
 - Become indistinguishable from squamous cells as they mature
- Transformation zone thought to arise from transformation of columnar epithelium through subcolumnar reserve cell hyperplasia
 - These reserve cells become metaplastic cells that then become the new squamous epithelium



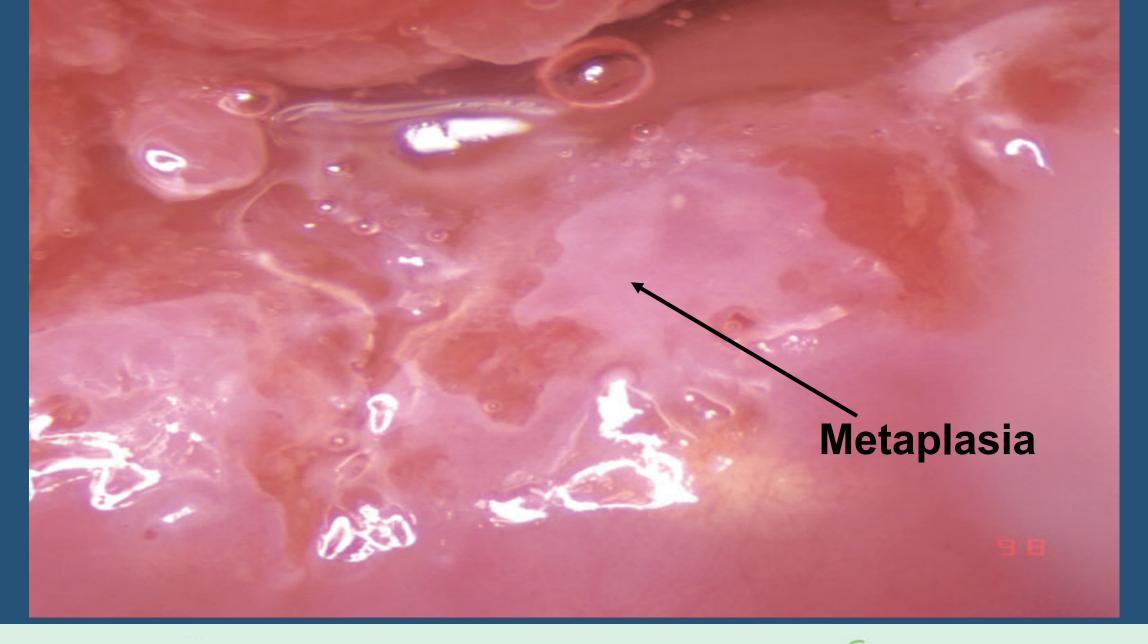














How squamous metaplasia occurs Mechanism of reserve cell hyperplasia

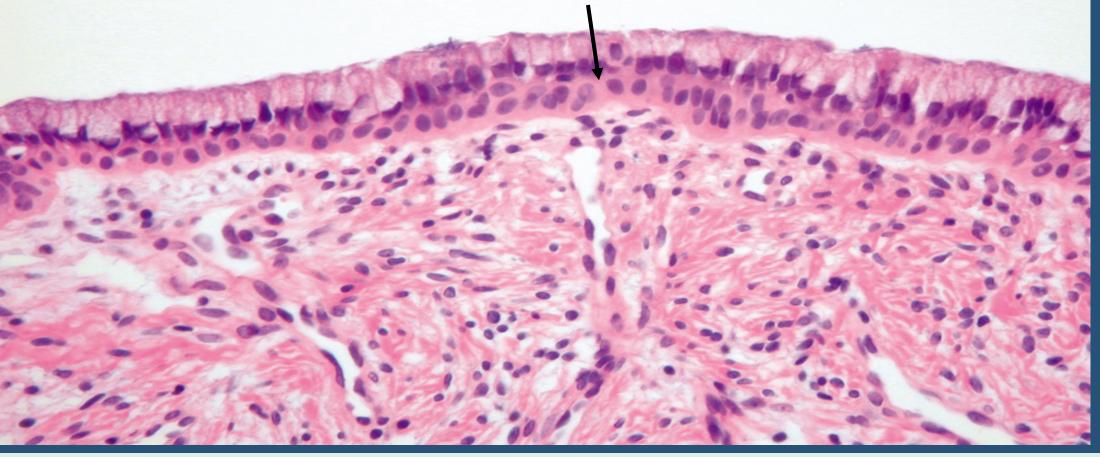
Estrogen lowers vaginal pH

- Estrogen levels increase in neonatal period and at puberty
- Cervical eversion mediated by estrogen
- Exposes endocervical columnar epithelium to more acidic vaginal environment
- "Acid burn" stimulates metaplasia of columnar epithelium



How squamous metaplasia occurs

Reserve cells below columnar cells

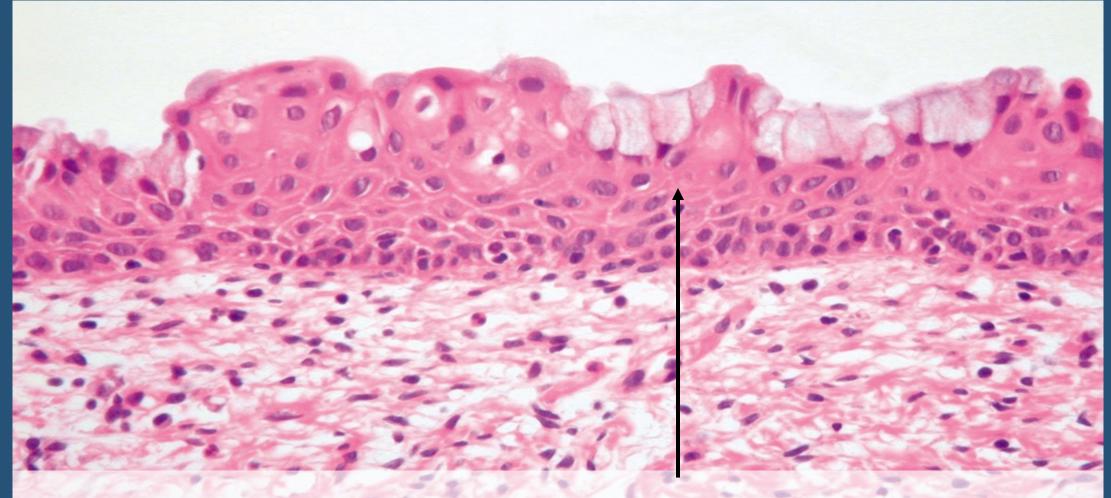




Factors inducing squamous metaplasia

- Etiologic factors
 - Mechanical irritation
 - Chronic inflammation
 - pH changes
 - Environmental conditions
- Probably begins when the original SCJ moves out of the os
 - Exposes delicate columnar cells to acidic vaginal environment



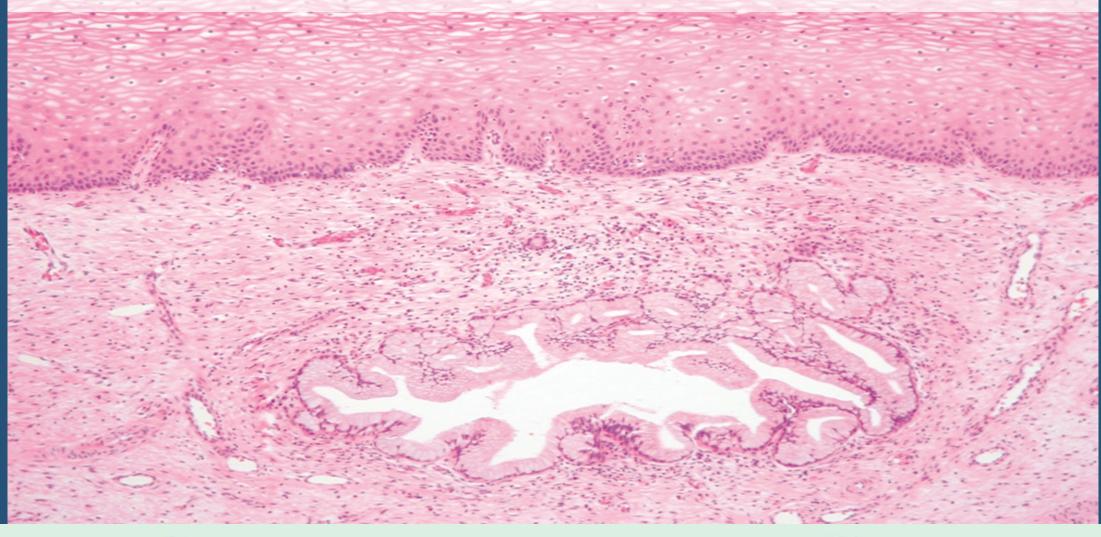


Proliferation of reserve cells - immature squamous cells directly below the remaining columnar cells





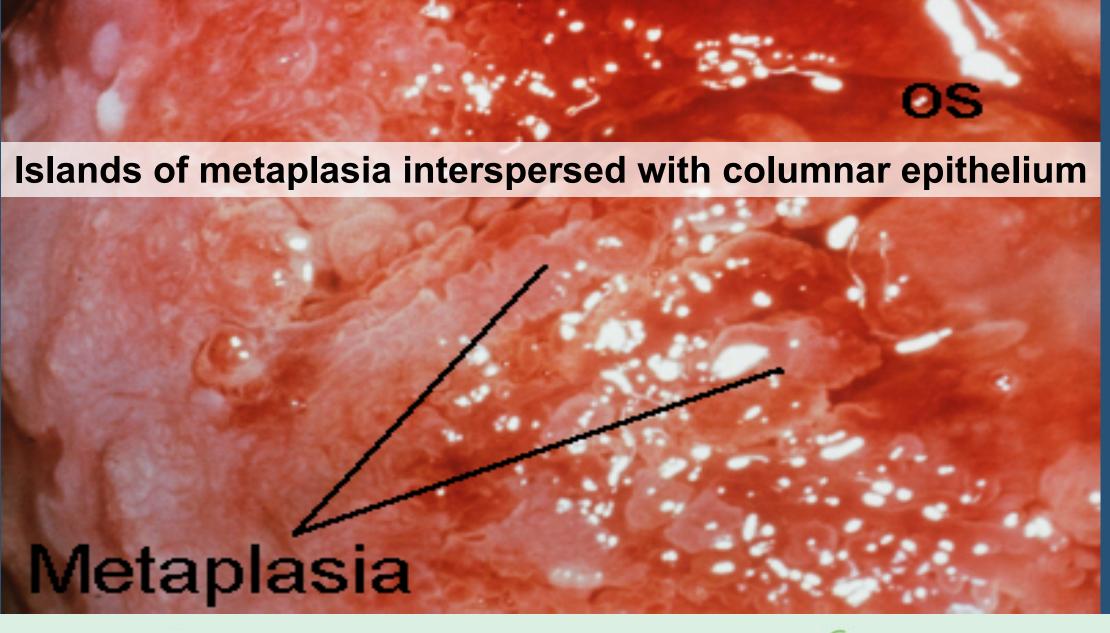
Mature stratified squamous epithelium



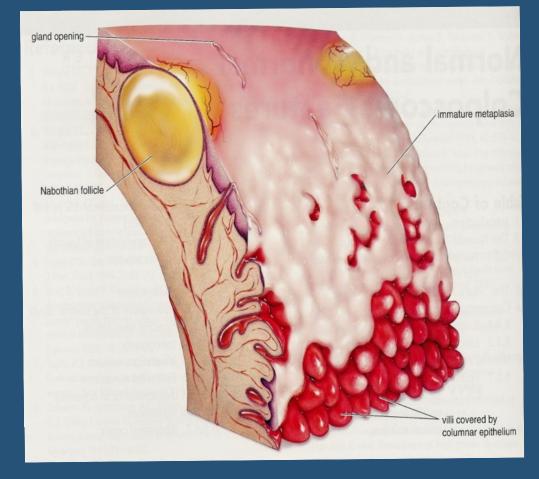


Squamous metaplasia

- Tips of columnar villi are traumatized by acidity of the vagina
- Immature cells gradually mature and the cells begin to produce glycogen
- Can be a random distribution on the cervix
 - Patchy, uneven areas of metaplasia
 - Immature tips have acetowhite appearance





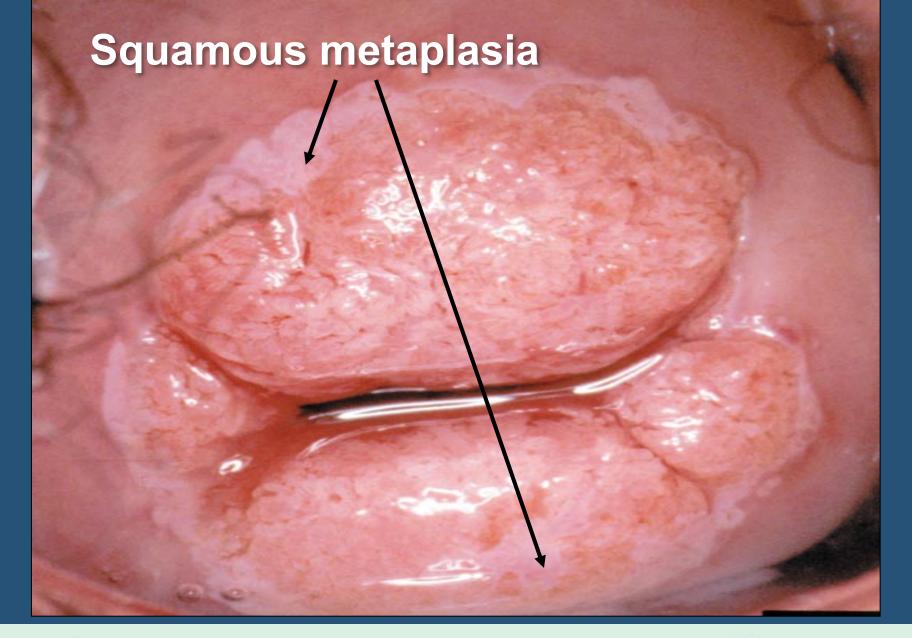


Normal transformation zone

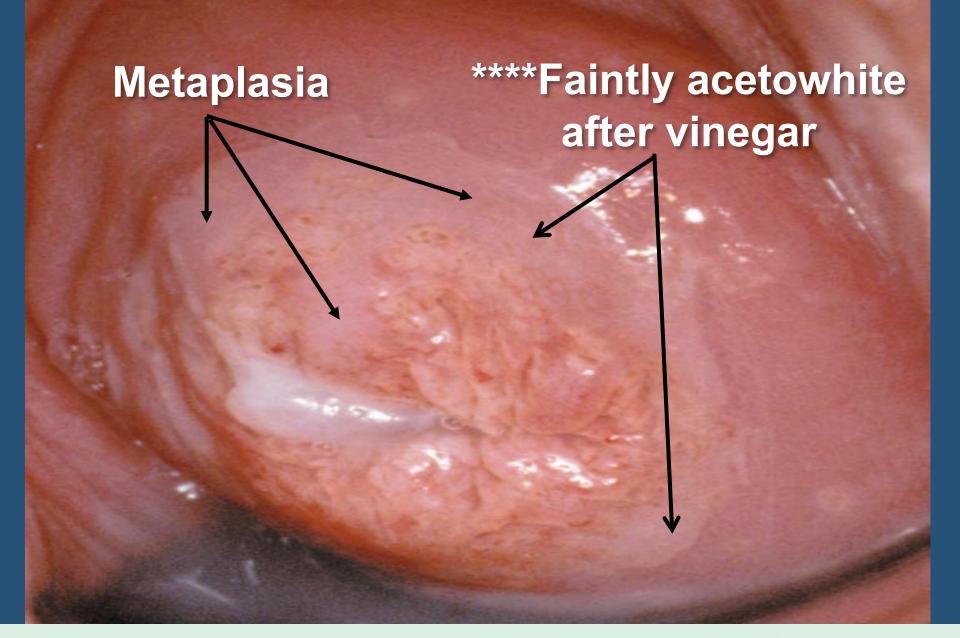


Mayeaux, E. J., & Cox, J. T. (2012). *Modern colposcopy: textbook & atlas*. Philadelphia: Wolters Kluwer Health/Lippincott Williams & Wilkins.



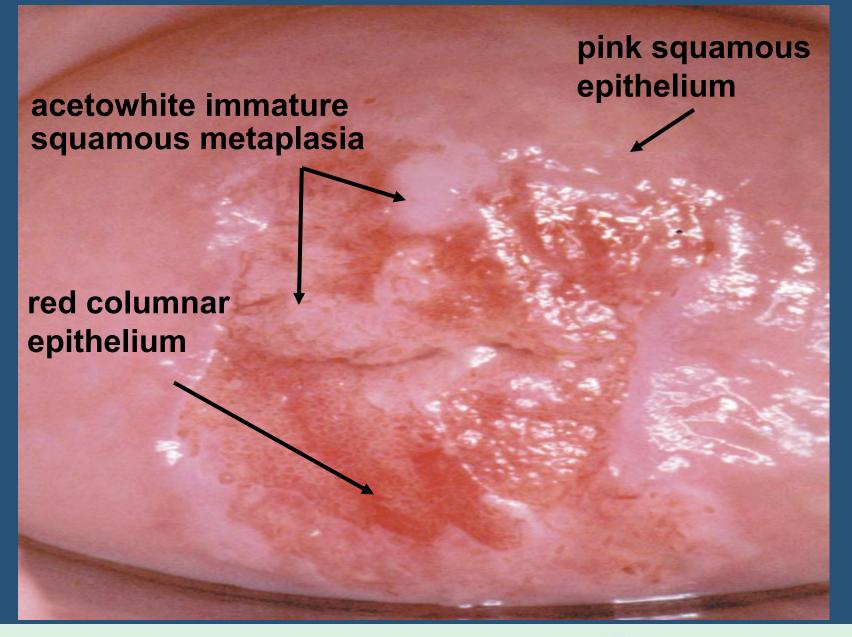








Colors of cervical epithelium

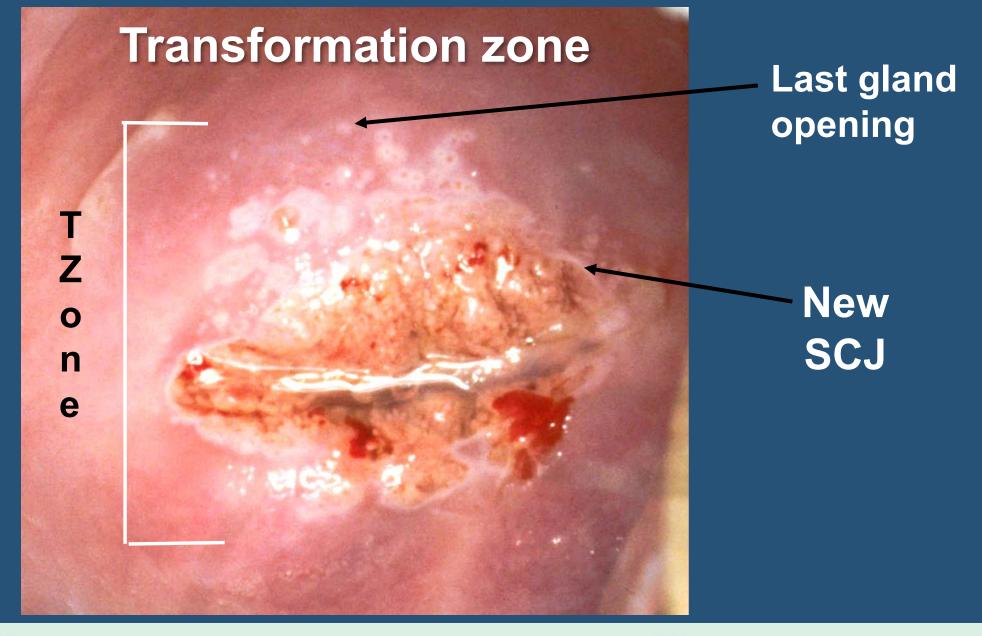




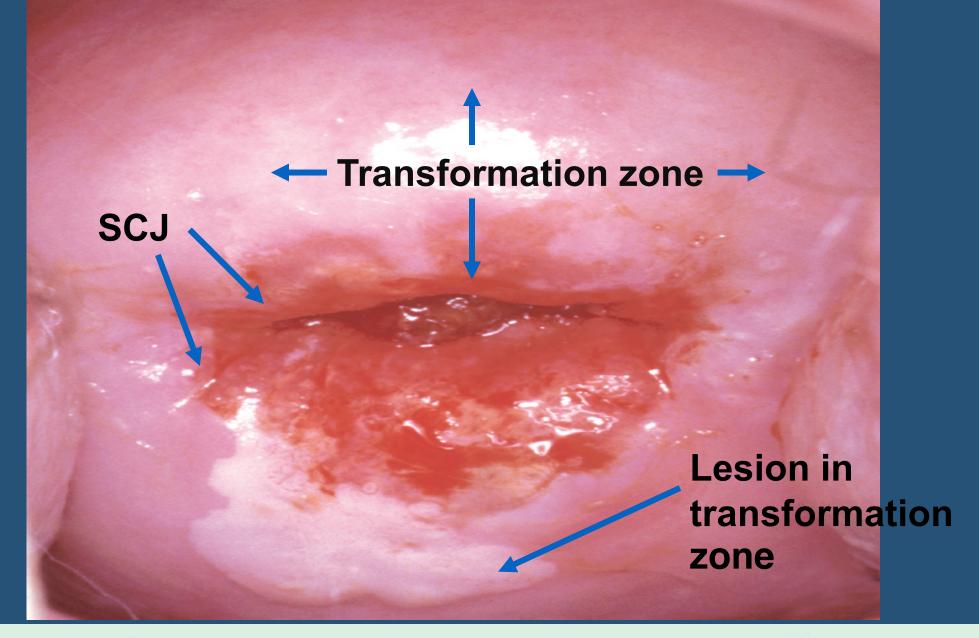
What is the transformation zone?

- Physical zone or area on the cervix
- Area between the original SCJ and the new SCJ
- TZ gets larger as the individual ages and remodeling or transformation occurs
- The new SCJ appears to "move" closer to the external os and then inside the endocervical canal
 - Process completed following menopause







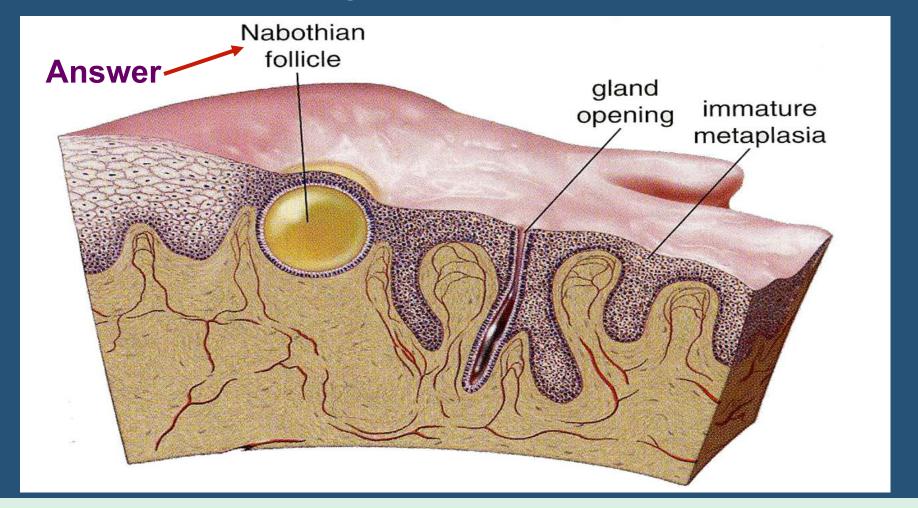




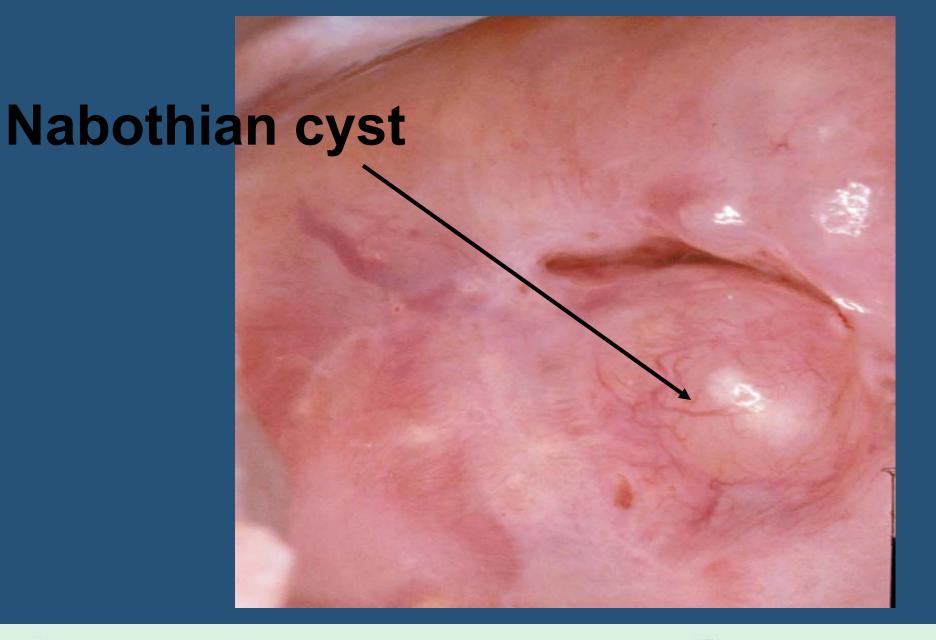
Nabothian cysts:

- Formed when the opening of the endocervical infoldings become blocked by metaplasia
- Accumulated mucus forms "cysts"
- Vessels accentuated over the cyst
- No treatment is necessary

What happens to those mucus secreting "glands" when they become blocked by squamous cells?



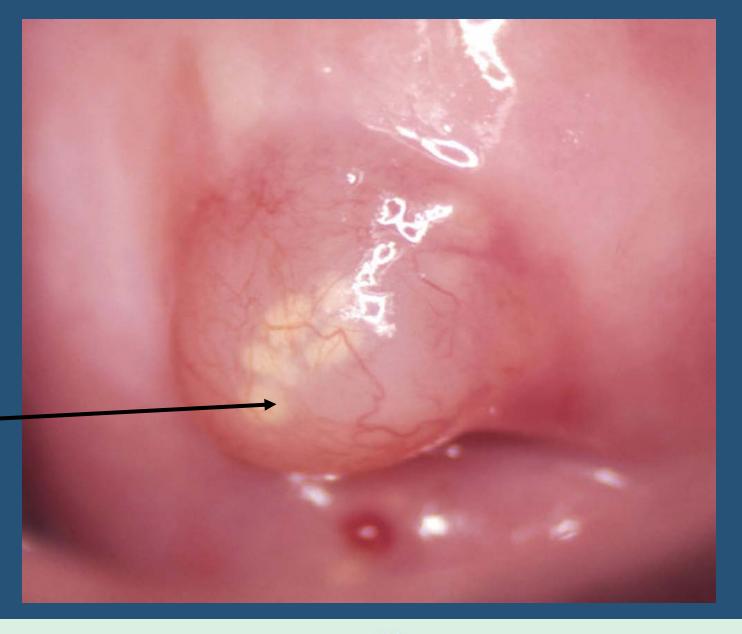




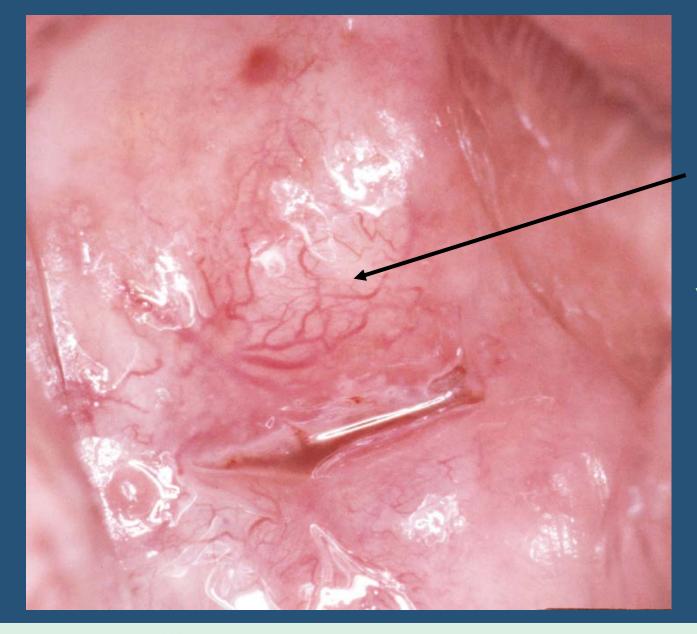


Nabothian cyst: good place to learn normal vessels

Branching vessels
Usually yellow







Nabothian cyst branching vessels

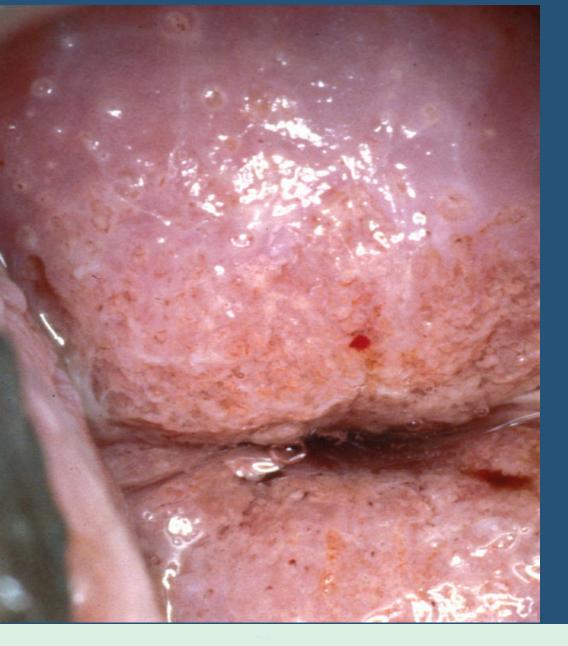


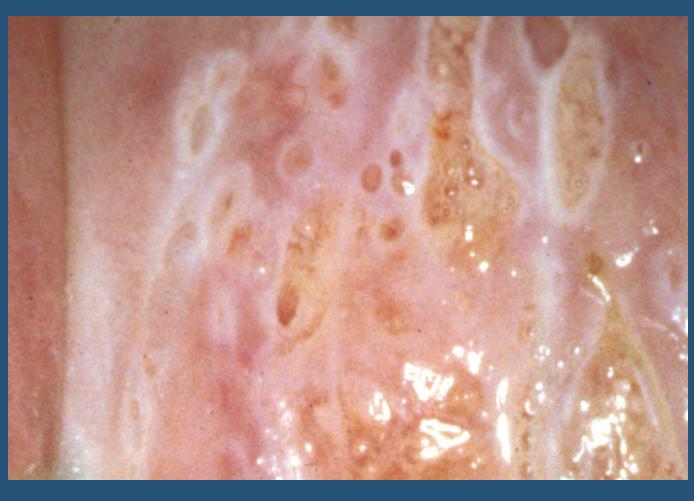
Multiple nabothian cysts



Islands of columnar epithelium

- Result from uneven process of squamous metaplasia
- Mini SCJ's
- If obliterate, may produce Nabothian cyst







"Gland" openings with mucin secretion

- Not true glands
- Infoldings of columnar epithelium

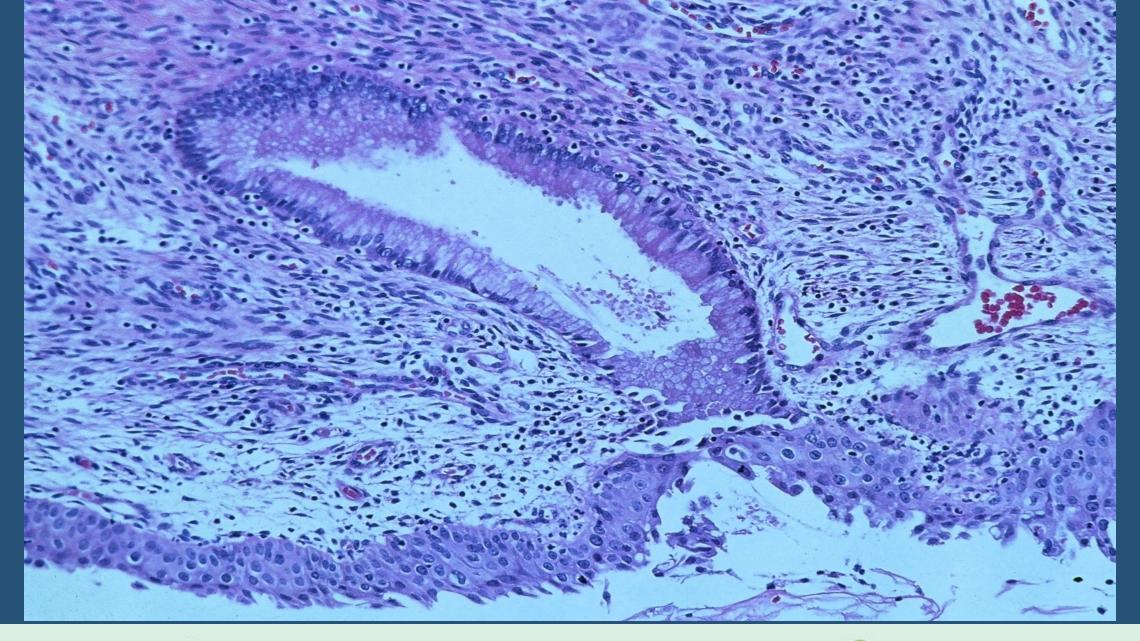






Image from Greenwald, Spitzer, Sedlis. ACOG Basic Colposcopy

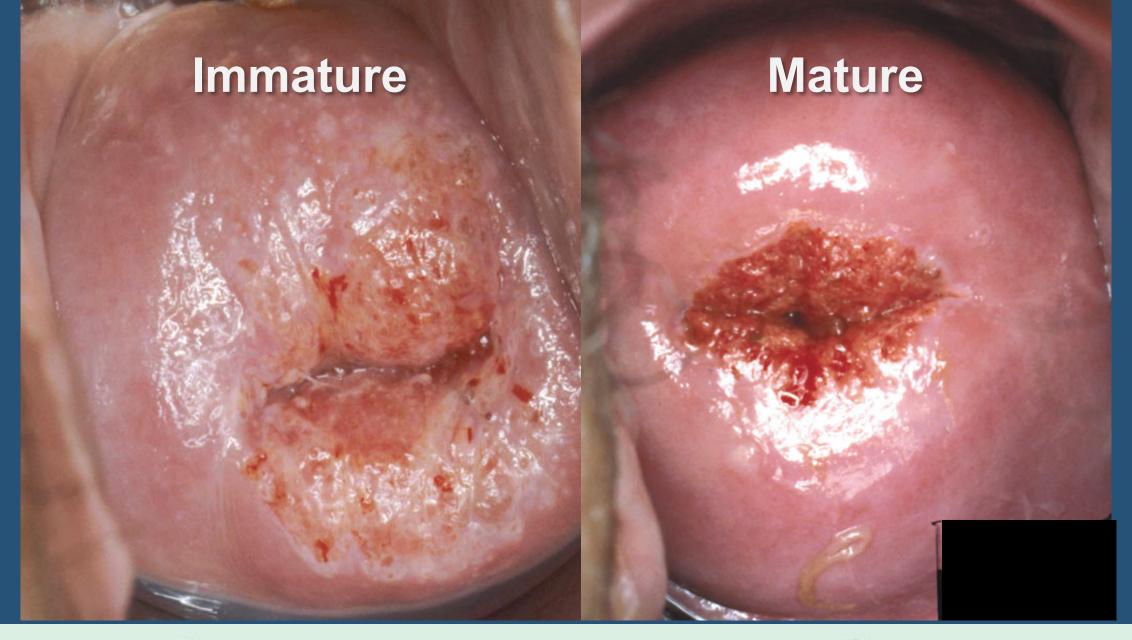




Immature metaplasia with multiple gland openings



- Mature squamous epithelium
- Final result of squamous metaplasia
 - Original SCJ and squamous epithelium interface becomes a squamous-squamous junction

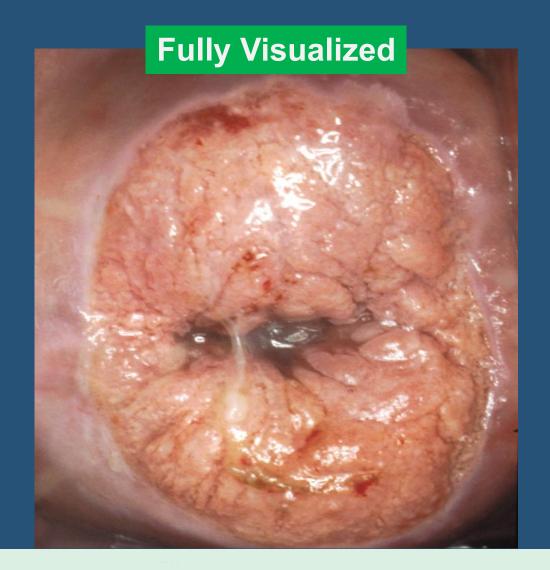


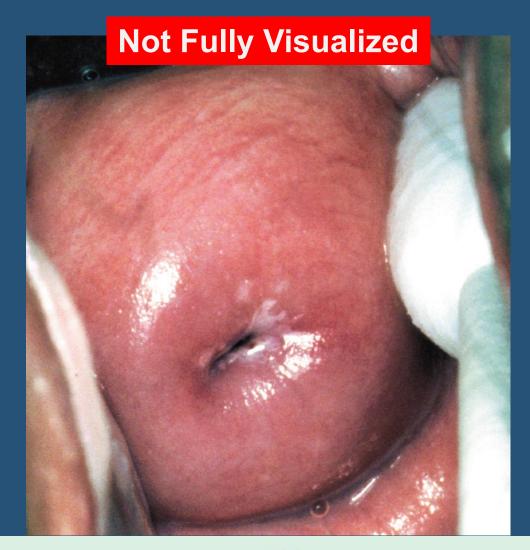


General assessment of the cervix and SCJ at the colposcopic exam: if fully visualized

- The entire new SCJ on the cervix can be visualized (360° of columnar epithelium)
- If any lesion is visible, must see it entirely
- May require manipulation or endocervical speculum for complete visualization

Is the general assessment of the SCJ and lesions fully visualized?





Summary

- Reviewed the epithelial features of the normal transformation zone
- Discussed the process of squamous metaplasia
- Explained the normal features of the transformation zone according to age
- Described features that define a fully visualized cervix and SCJ at colposcopy

