

# Pleomorphic Adenoma

- Slow-growing, painless mass
- Parotid: 90% in superficial lobe, most in tail of gland
- Minor salivary gland: lateral palate, submucosal mass
- Solitary vs. synchronous/metachronous neoplasms

# Pleomorphic Adenoma

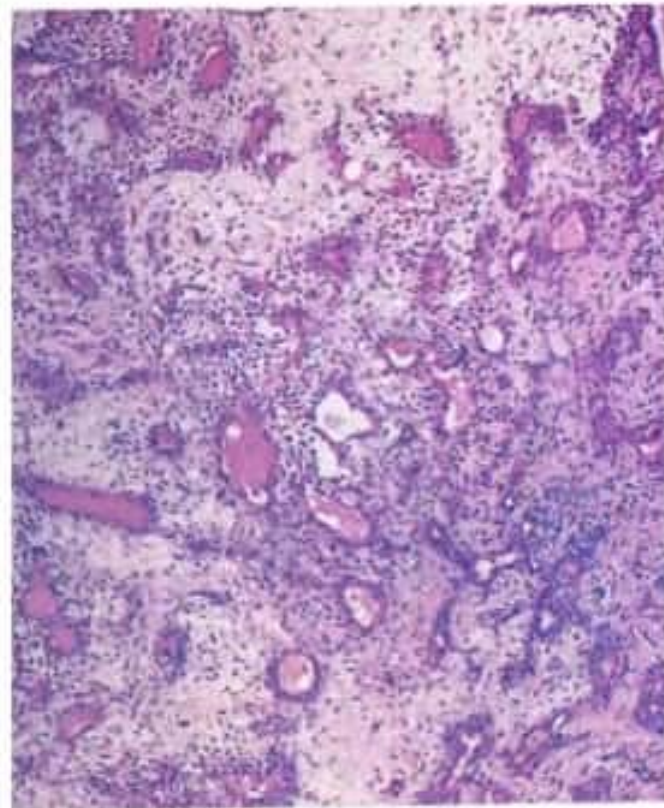
- Gross pathology
  - Smooth
  - Well-demarcated
  - Solid
  - Cystic changes
  - Myxoid stroma



# Pleomorphic Adenoma

- **Histology**

- Mixture of epithelial, myoepithelial and stromal components
- Epithelial cells: nests, sheets, ducts, trabeculae
- Stroma: myxoid, chondroid, fibroid, osteoid
- No true capsule
- Tumor pseudopods



# Pleomorphic Adenoma

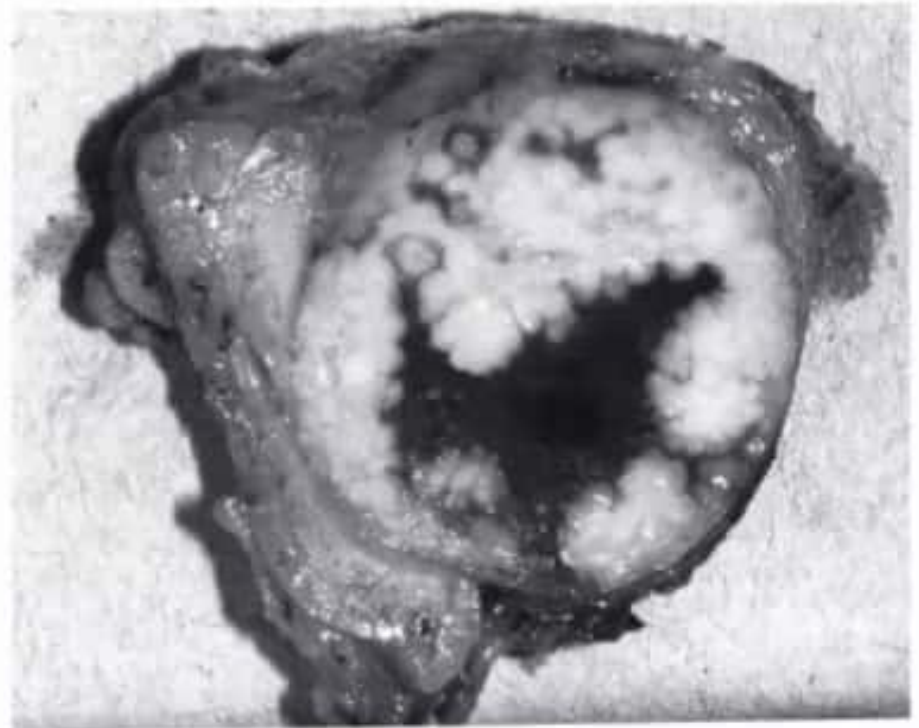
- Treatment: complete surgical excision
  - Parotidectomy with facial nerve preservation
  - Submandibular gland excision
  - Wide local excision of minor salivary gland
  
- Avoid enucleation and tumor spill

# Warthin's Tumor

- AKA: papillary cystadenoma lymphomatosum
- 6-10% of parotid neoplasms
- Older, Caucasian, males
- 10% bilateral or multicentric
- 3% with associated neoplasms
- Presentation: slow-growing, painless mass

# Warthin's Tumor

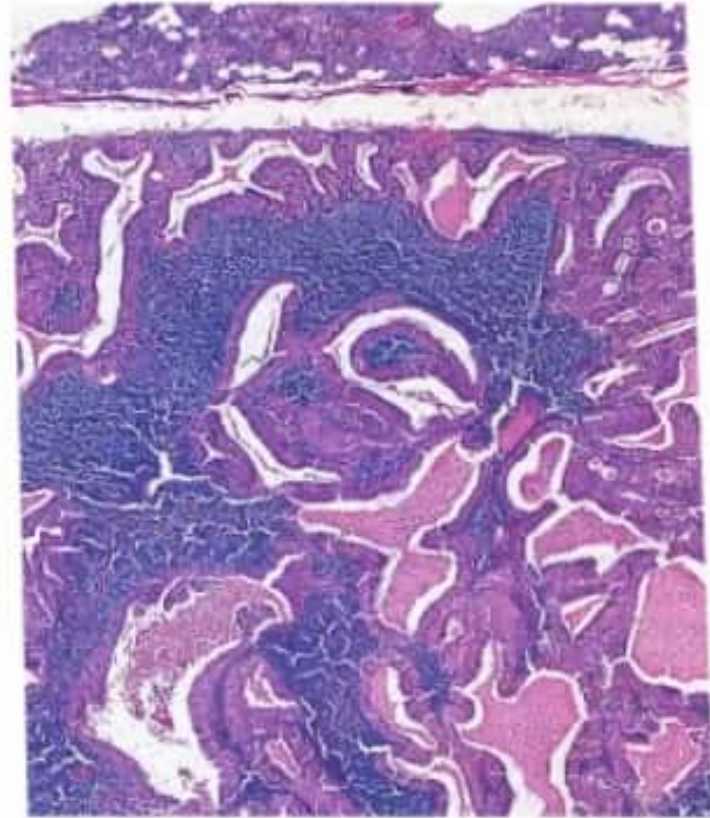
- Gross pathology
  - Encapsulated
  - Smooth/lobulated surface
  - Cystic spaces of variable size, with viscous fluid, shaggy epithelium
  - Solid areas with white nodules representing lymphoid follicles



# Warthin's Tumor

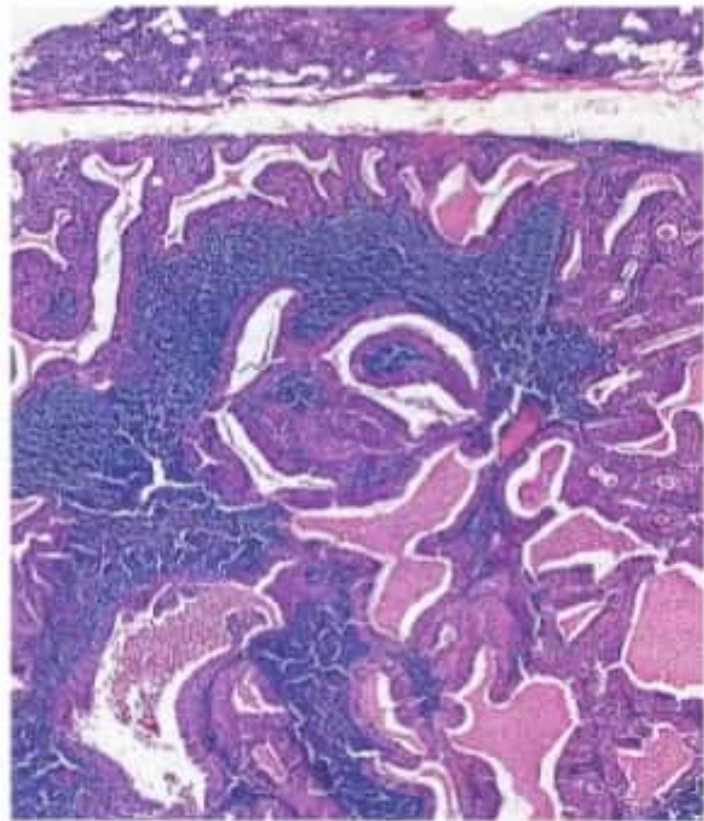
- Histology

- Papillary projections into cystic spaces surrounded by lymphoid stroma
- Epithelium: double cell layer
  - Luminal cells
  - Basal cells
- Stroma: mature lymphoid follicles with germinal centers



# Warthin's Tumor

- Histology
  - Papillary projections into cystic spaces surrounded by lymphoid stroma
  - Epithelium: double cell layer
    - Luminal cells
    - Basal cells
  - Stroma: mature lymphoid follicles with germinal centers





# Oncocytoma

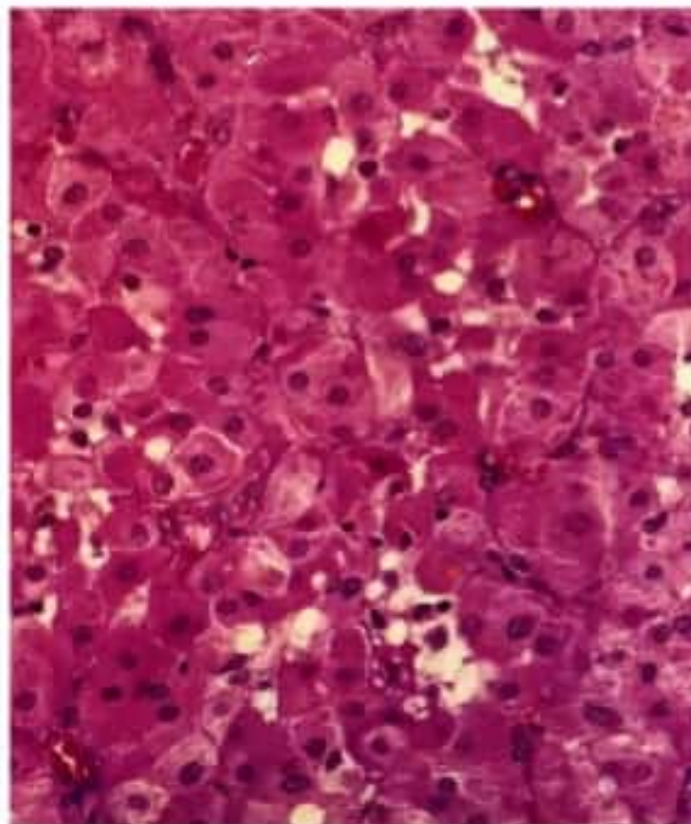
- Rare: 2.3% of benign salivary tumors
- 6<sup>th</sup> decade
- M:F = 1:1
- Parotid: 78%
- Submandibular gland: 9%
- Minor salivary glands: palate, buccal mucosa, tongue

# Oncocytoma

- Presentation
  - Enlarging, painless mass
- Technetium-99m pertechnetate scintigraphy
  - Mitochondrial hyperplasia

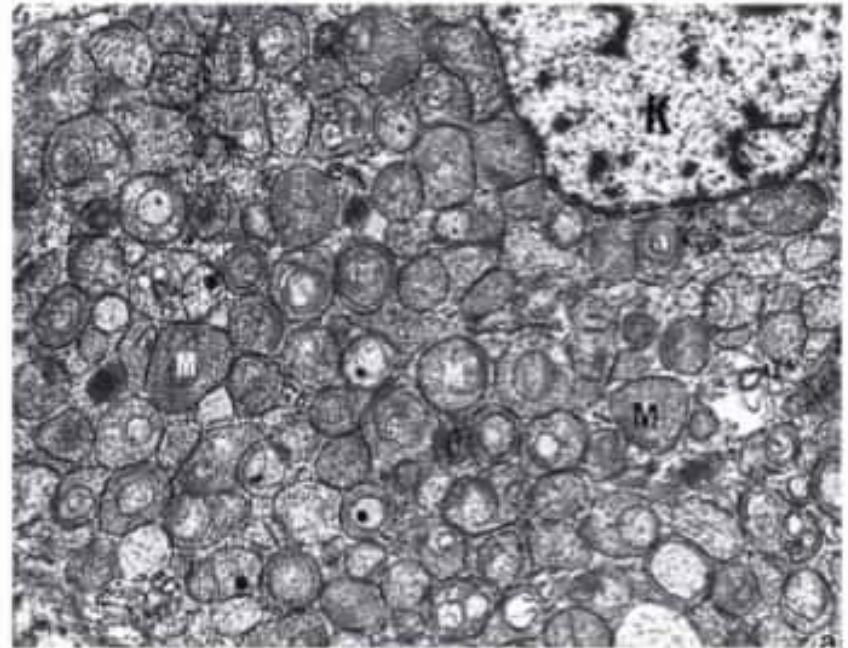
# Oncocytoma

- Gross
  - Encapsulated
  - Homogeneous, smooth
  - Orange/rust color
- Histology
  - Cords of uniform cells and thin fibrous stroma
  - Large polyhedral cells
  - Distinct cell membrane
  - Granular, eosinophilic cytoplasm
  - Central, round, vesicular nucleus



# Oncocytoma

- Electron microscopy:
  - Mitochondrial hyperplasia
  - 60% of cell volume

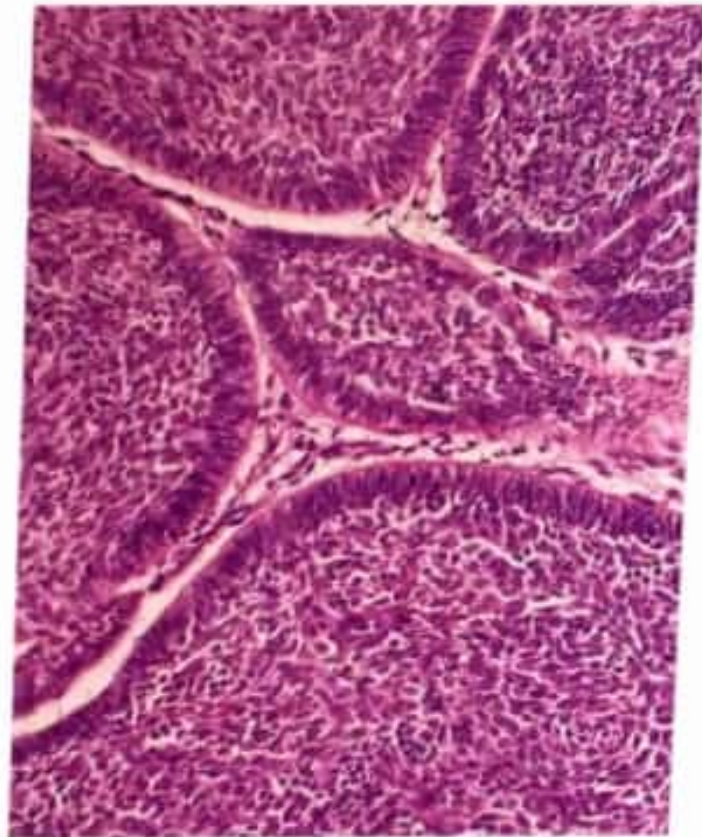


# Monomorphic Adenomas

- Types: Basal cell, canalicular, sebaceous, glycogen-rich, clear cell
- Basal cell is most common: 1.8% of benign epithelial salivary gland neoplasms
- 6<sup>th</sup> decade
- M:F = approximately 1:1
- Caucasian > African American
- Most common in parotid

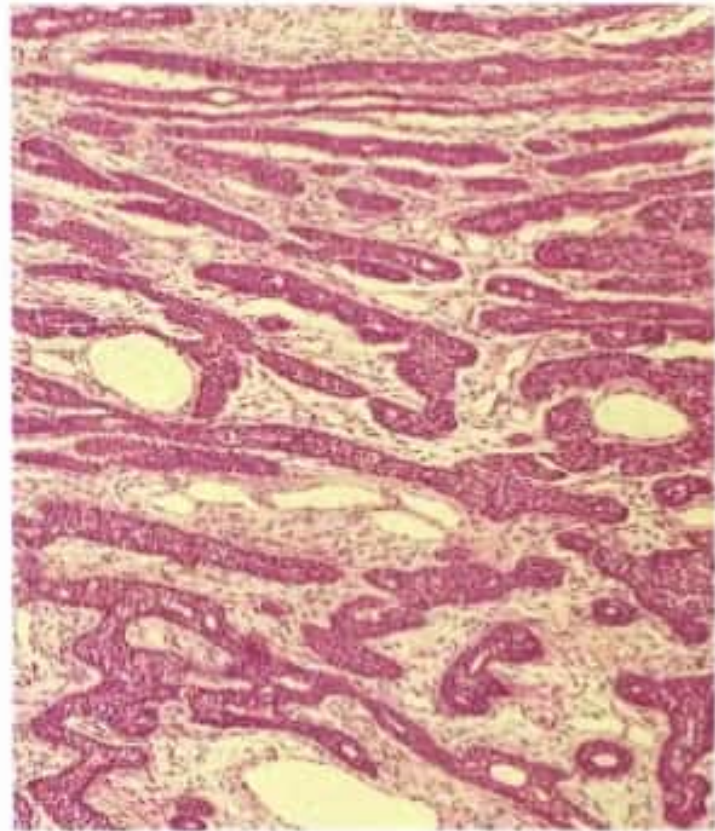
# Basal Cell Adenoma

- Solid
  - Most common
  - Solid nests of tumor cells
  - Uniform, hyperchromatic, round nuclei, indistinct cytoplasm
  - Peripheral nuclear palisading
  - Scant stroma



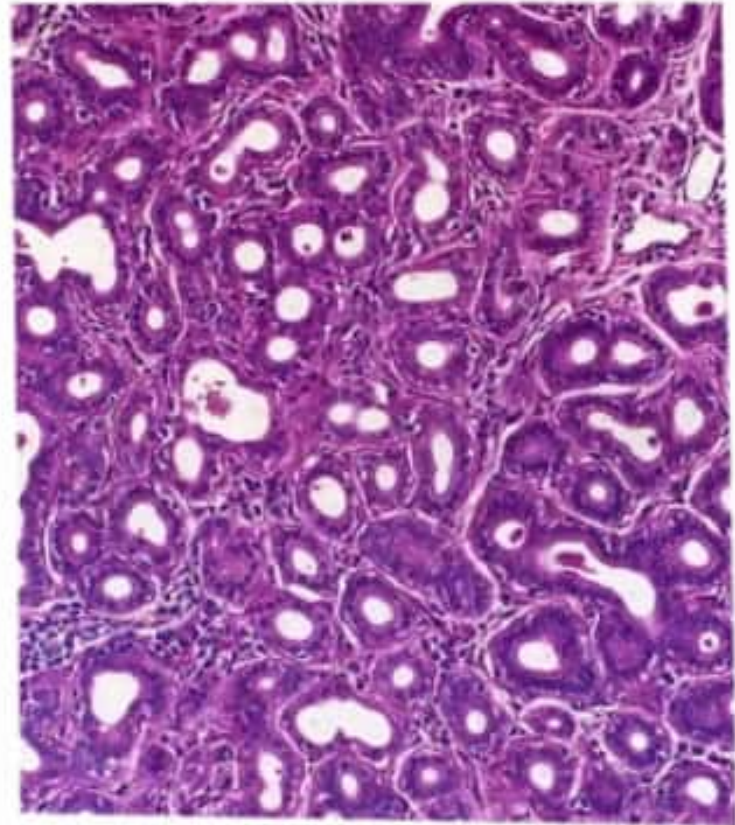
# Basal Cell Adenoma

- Trabecular
  - Cells in elongated trabecular pattern
  - Vascular stroma



# Basal Cell Adenoma

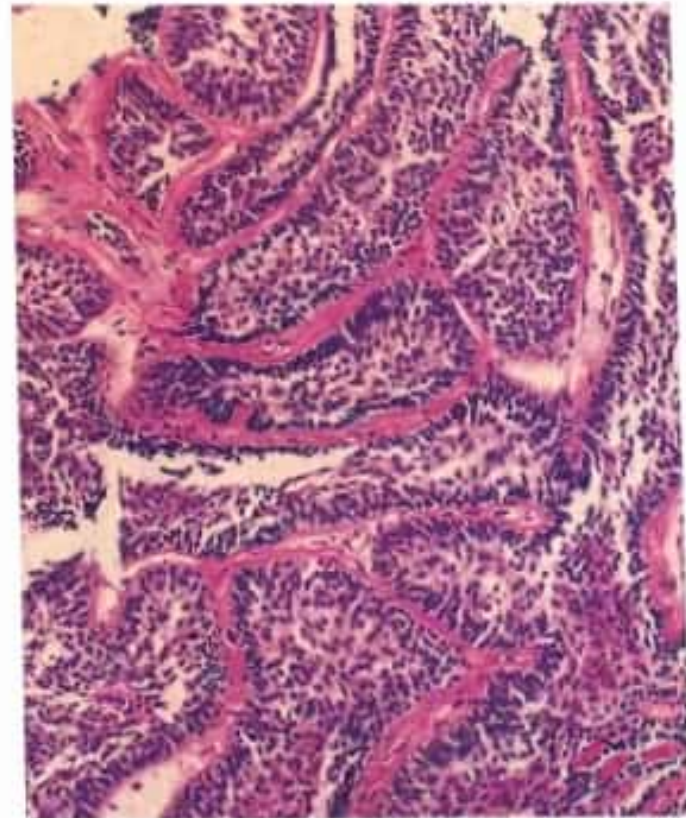
- Tubular
  - Multiple duct-like structures
  - Columnar cell lining
  - Vascular stroma





# Basal Cell Adenoma

- Membranous
  - Thick eosinophilic hyaline membranes surrounding nests of tumor cells
  - “jigsaw-puzzle” appearance



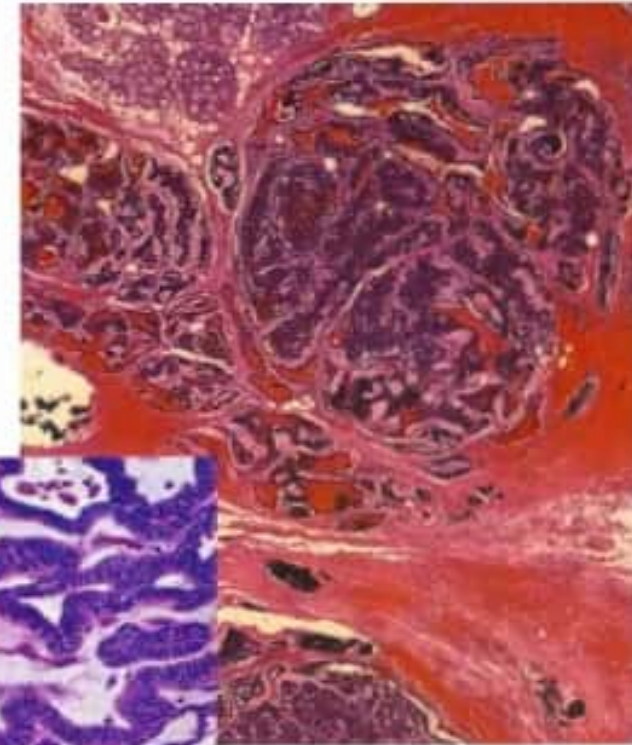
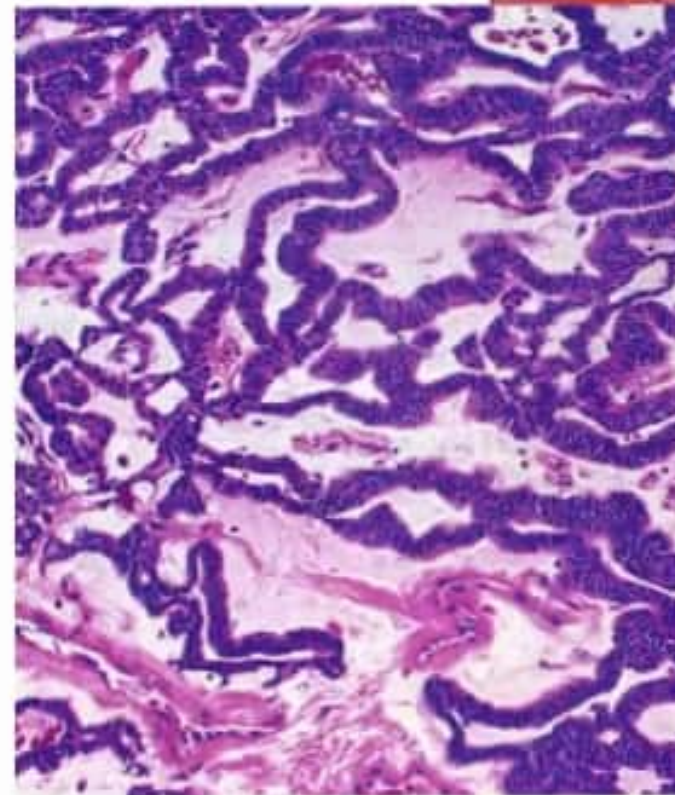
# Monomorphic Adenomas

- Canalicular adenoma
  - 7<sup>th</sup> decade
  - F:M – 1.8:1
  - Most common in minor salivary glands of the upper lip (74%)
  - Painless submucosal mass

# Canalicular Adenoma

- **Histology**

- Well-circumscribed
- Multiple foci
- Tubular structures lined by columnar or cuboidal cells
- Vascular stroma



# Myoepithelioma

- <1% of all salivary neoplasms
- 3<sup>rd</sup>-6<sup>th</sup> decades
- F>M
- Minor salivary glands > parotid > submandibular gland
- Presentation: asymptomatic mass

# Myoepithelioma

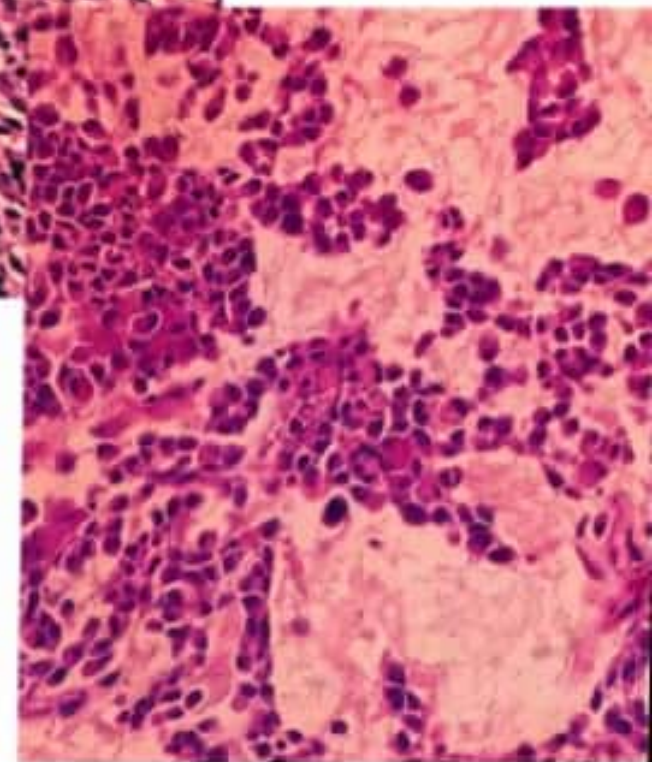
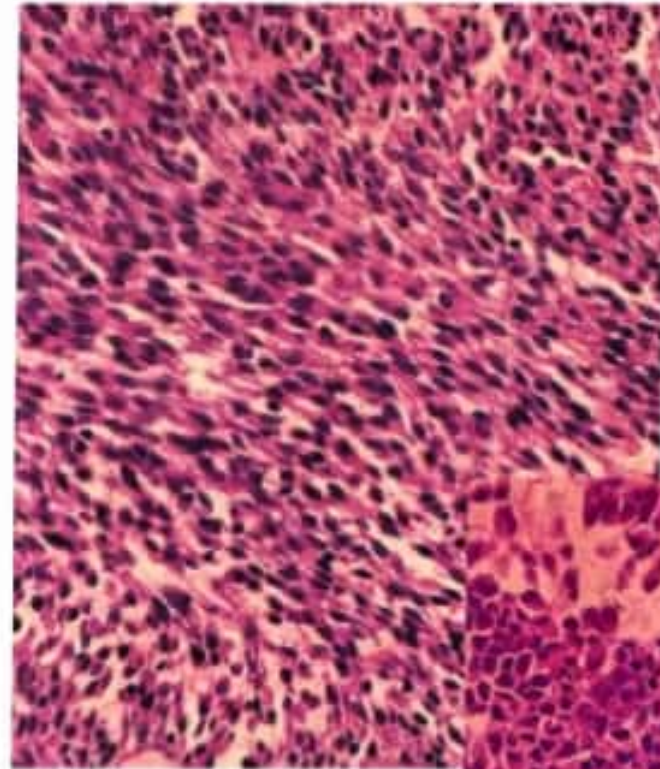
- Histology

- Spindle cell

- More common
    - Parotid
    - Uniform, central nuclei
    - Eosinophilic granular or fibrillar cytoplasm

- Plasmacytoid cell

- Polygonal
    - Eccentric oval nuclei



# MALIGNANT NEOPLASMS

1. Mucoepidermoid Carcinoma
2. Adenoid Cystic Carcinoma
3. Acinic Cell Carcinoma
4. Adenocarcinoma
5. Malignant Mixed Tumors: Carcinoma Ex-Pleomorphic Adenoma, Carcinosarcoma
6. Squamous Cell Carcinoma
7. Polymorphous Low-Grade Adenocarcinoma
8. Clear Cell Carcinoma
9. Epithelial-Myoepithelial Carcinoma
10. Undifferentiated Carcinoma

# MALIGNANT TUMORS

## Mucoepidermoid Carcinoma

- Most common salivary gland malignancy in children
- 5-9% of salivary neoplasms
- Parotid 45-70% of cases
- Palate 18%
- 3<sup>rd</sup>-8<sup>th</sup> decades, peak in 5<sup>th</sup> decade
- F>M
- Caucasian > African American

# Mucoepidermoid Carcinoma

- Presentation
  - Low-grade: slow growing, painless mass
  - High-grade: rapidly enlarging, +/- pain
  
- \*\*Minor salivary glands: may be mistaken for benign or inflammatory process
  - Hemangioma
  - Papilloma
  - Tori



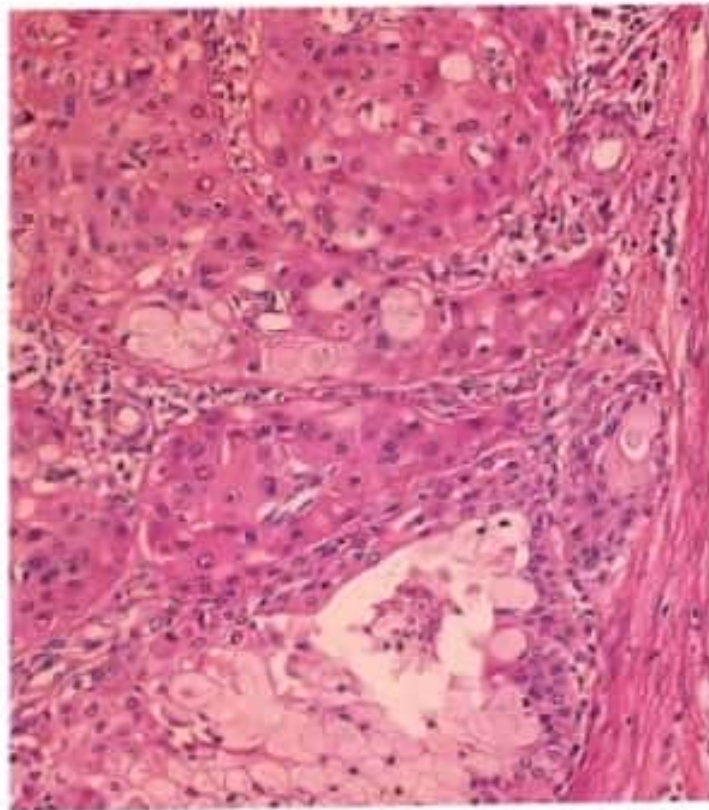
# Mucoepidermoid Carcinoma

- Gross pathology
  - Well-circumscribed to partially encapsulated to unencapsulated
  - Solid tumor with cystic spaces



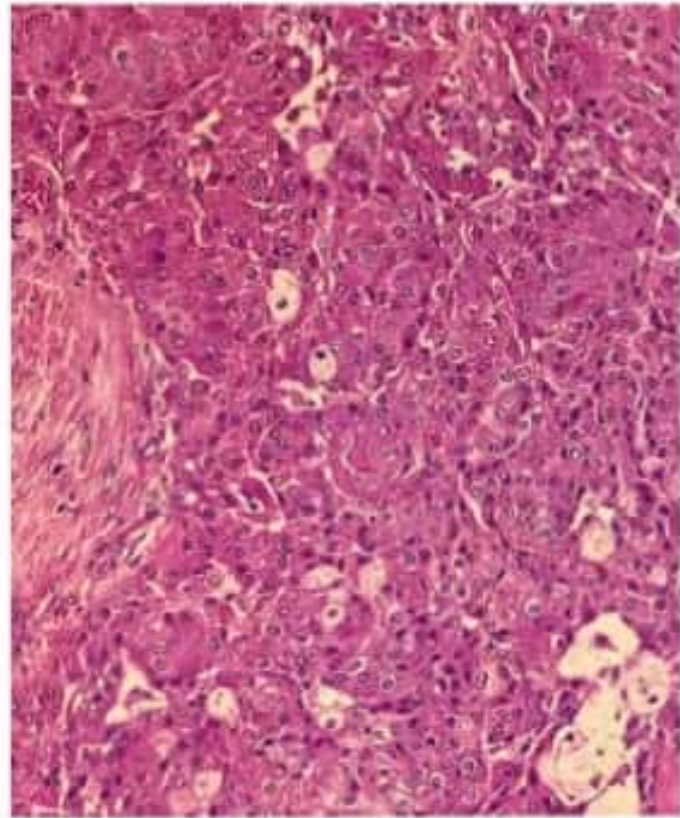
# Mucoepidermoid Carcinoma

- Histology—Low-grade
  - Mucus cell > epidermoid cells
  - Prominent cysts
  - Mature cellular elements



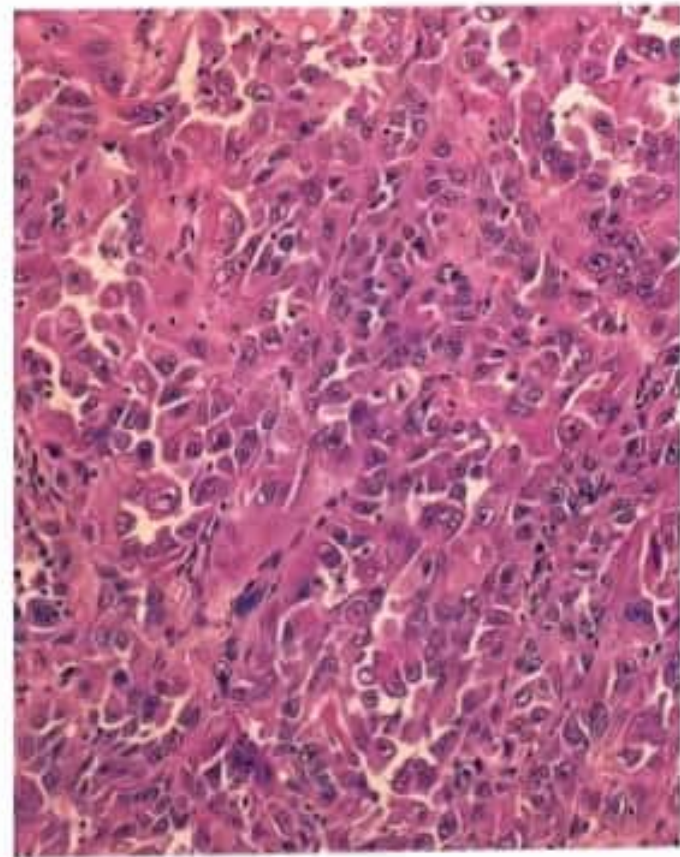
# Mucoepidermoid Carcinoma

- Histology—Intermediate-grade
  - Mucus = epidermoid
  - Fewer and smaller cysts
  - Increasing pleomorphism and mitotic figures



# Mucoepidermoid Carcinoma

- Histology—High-grade
  - Epidermoid > mucus
  - Solid tumor cell proliferation
  - Mistaken for SCCA
    - Mucin staining



# Mucoepidermoid Carcinoma

- Treatment
  - Influenced by site, stage, grade
  - Stage I & II
    - Wide local excision
  - Stage III & IV
    - Radical excision
    - +/- neck dissection
    - +/- postoperative radiation therapy

# Adenoid Cystic Carcinoma

- Overall most common malignancy
- Most common in submandibular, sublingual and minor salivary glands
- M = F
- 5<sup>th</sup> decade
- Presentation
  - Asymptomatic enlarging mass
  - Pain, paresthesias, facial weakness/paralysis

# Adenoid Cystic Carcinoma

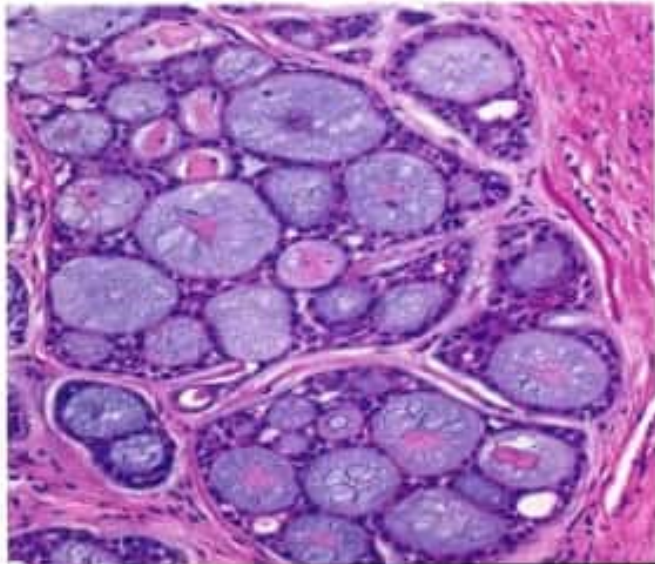
- Gross pathology
  - Well-circumscribed
  - Solid, rarely with cystic spaces
  - infiltrative



# Adenoid Cystic Carcinoma

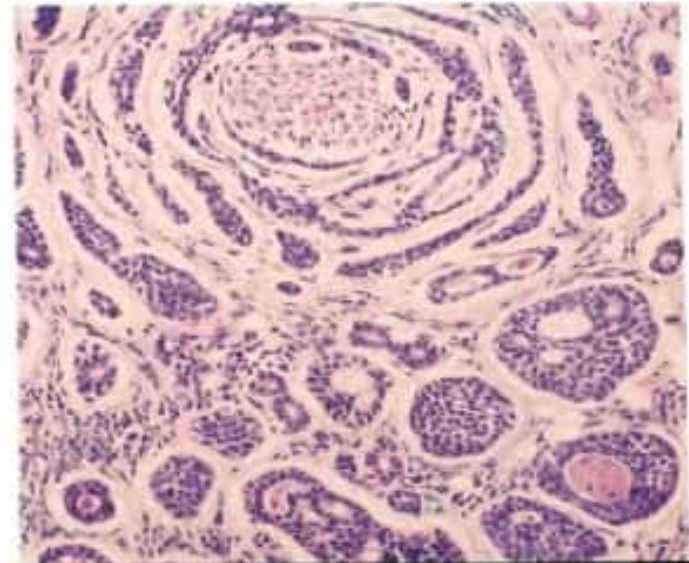
- Histology—tubular pattern

- Layered cells forming duct-like structures
- Basophilic mucinous substance



- Histology—solid pattern

- Solid nests of cells without cystic or tubular spaces





# Adenoid Cystic Carcinoma

- Treatment
  - Complete local excision
  - Tendency for perineural invasion: facial nerve sacrifice
  - Postoperative XRT
- Prognosis
  - Local recurrence: 42%
  - Distant metastasis: lung
  - Indolent course: 5-year survival 75%, 20-year survival 13%

# Acinic Cell Carcinoma

- 2<sup>nd</sup> most common parotid and pediatric malignancy
- 5<sup>th</sup> decade
- F>M
- Bilateral parotid disease in 3%
- Presentation
  - Solitary, slow-growing, often painless mass

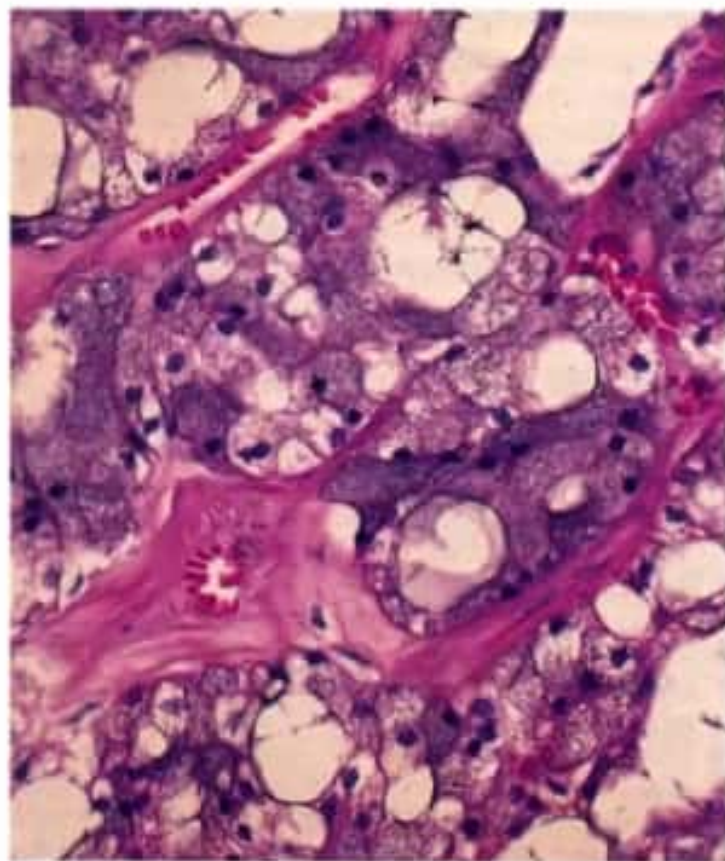
# Acinic Cell Carcinoma

- Gross pathology
  - Well-demarcated
  - Most often homogeneous



# Acinic Cell Carcinoma

- Histology
  - Solid and microcystic patterns
    - Most common
    - Solid sheets
    - Numerous small cysts
  - Polyhedral cells
  - Small, dark, eccentric nuclei
  - Basophilic granular cytoplasm



# Acinic Cell Carcinoma

- Treatment
  - Complete local excision
  - +/- postoperative XRT
- Prognosis
  - 5-year survival: 82%
  - 10-year survival: 68%
  - 25-year survival: 50%

# Acinic Cell Carcinoma

- Treatment
  - Complete local excision
  - +/- postoperative XRT
- Prognosis
  - 5-year survival: 82%
  - 10-year survival: 68%
  - 25-year survival: 50%

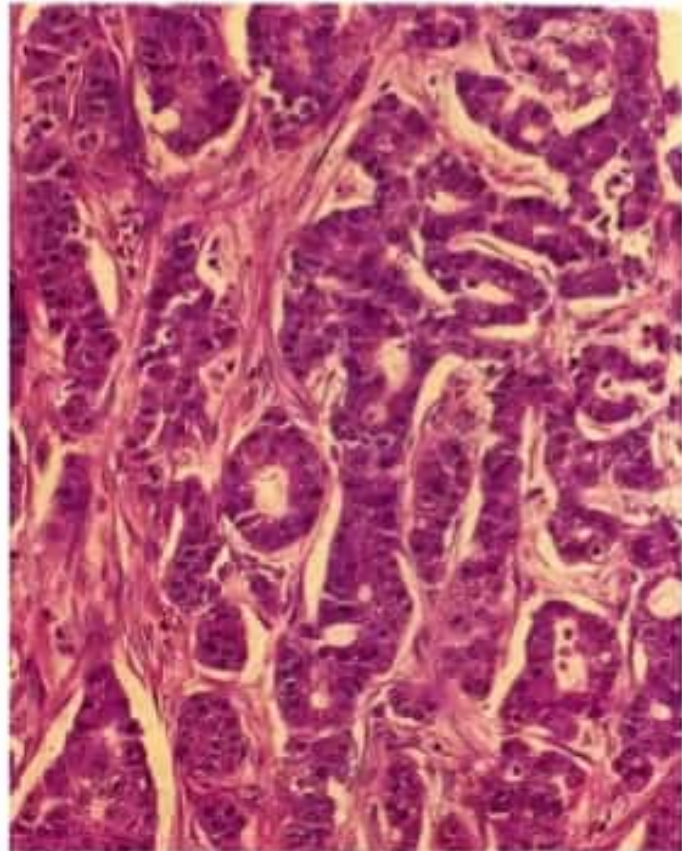
# Adenocarcinoma

- Rare
- 5<sup>th</sup> to 8<sup>th</sup> decades
- F > M
- Parotid and minor salivary glands
- Presentation:
  - Enlarging mass
  - 25% with pain or facial weakness



# Adenocarcinoma

- Histology
  - Heterogeneity
  - Presence of glandular structures and absence of epidermoid component
  - Grade I
  - Grade II
  - Grade III





# Adenocarcinoma

- Treatment
  - Complete local excision
  - Neck dissection
  - Postoperative XRT
- Prognosis
  - Local recurrence: 51%
  - Regional metastasis: 27%
  - Distant metastasis: 26%
  - 15-year cure rate:
    - Stage I = 67%
    - Stage II = 35%
    - Stage III = 8%

# Malignant Mixed Tumors

- Carcinoma ex-pleomorphic adenoma
  - Carcinoma developing in the epithelial component of preexisting pleomorphic adenoma
- Carcinosarcoma
  - True malignant mixed tumor—carcinomatous and sarcomatous components
- Metastatic mixed tumor
  - Metastatic deposits of otherwise typical pleomorphic adenoma

# Malignant Mixed Tumors

- Carcinoma ex-pleomorphic adenoma
  - Carcinoma developing in the epithelial component of preexisting pleomorphic adenoma
- Carcinosarcoma
  - True malignant mixed tumor—carcinomatous and sarcomatous components
- Metastatic mixed tumor
  - Metastatic deposits of otherwise typical pleomorphic adenoma

# Malignant Mixed Tumors

- Carcinoma ex-pleomorphic adenoma
  - Carcinoma developing in the epithelial component of preexisting pleomorphic adenoma
- Carcinosarcoma
  - True malignant mixed tumor—carcinomatous and sarcomatous components
- Metastatic mixed tumor
  - Metastatic deposits of otherwise typical pleomorphic adenoma

# Carcinoma Ex-Pleomorphic Adenoma

- 2-4% of all salivary gland neoplasms
- 4-6% of mixed tumors
- 6<sup>th</sup>-8<sup>th</sup> decades
- Parotid > submandibular > palate
- Risk of malignant degeneration
  - 1.5% in first 5 years
  - 9.5% after 15 years
- Presentation
  - Longstanding painless mass that undergoes sudden enlargement

# Carcinoma Ex-Pleomorphic Adenoma

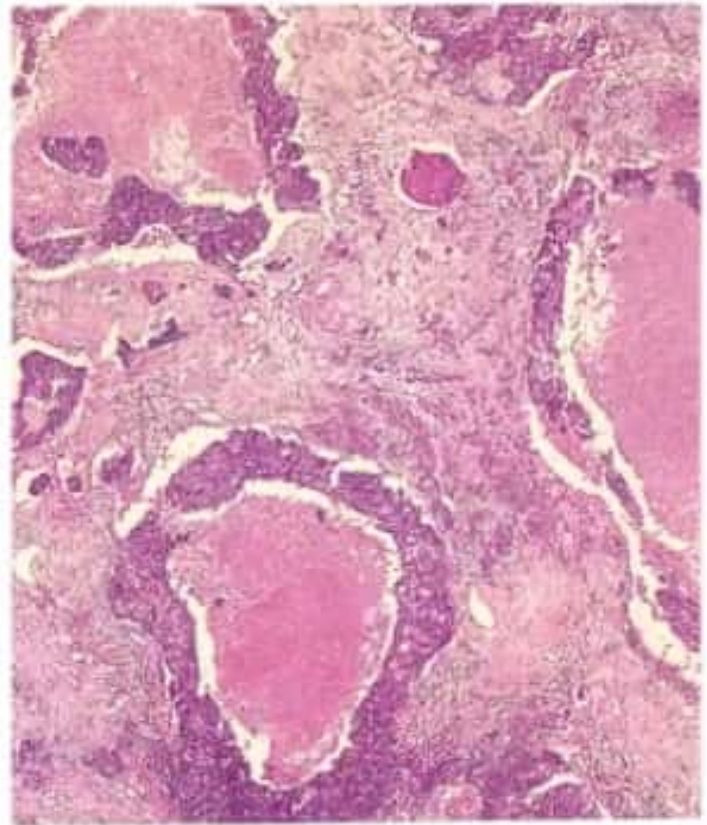
- Gross pathology
  - Poorly circumscribed
  - Infiltrative
  - Hemorrhage and necrosis



# Carcinoma Ex-Pleomorphic Adenoma

- **Histology**

- Malignant cellular change adjacent to typical pleomorphic adenoma
- Carcinomatous component
  - Adenocarcinoma
  - Undifferentiated



# Carcinoma Ex-Pleomorphic Adenoma

- Treatment
  - Radical excision
  - Neck dissection (25% with lymph node involvement at presentation)
  - Postoperative XRT
- Prognosis
  - Dependent upon stage and histology



# Carcinosarcoma

- Rare: <.05% of salivary gland neoplasms
- 6<sup>th</sup> decade
- M = F
- Parotid
- History of previously excised pleomorphic adenoma, recurrent pleomorphic adenoma or recurring pleomorphic treated with XRT
- Presentation

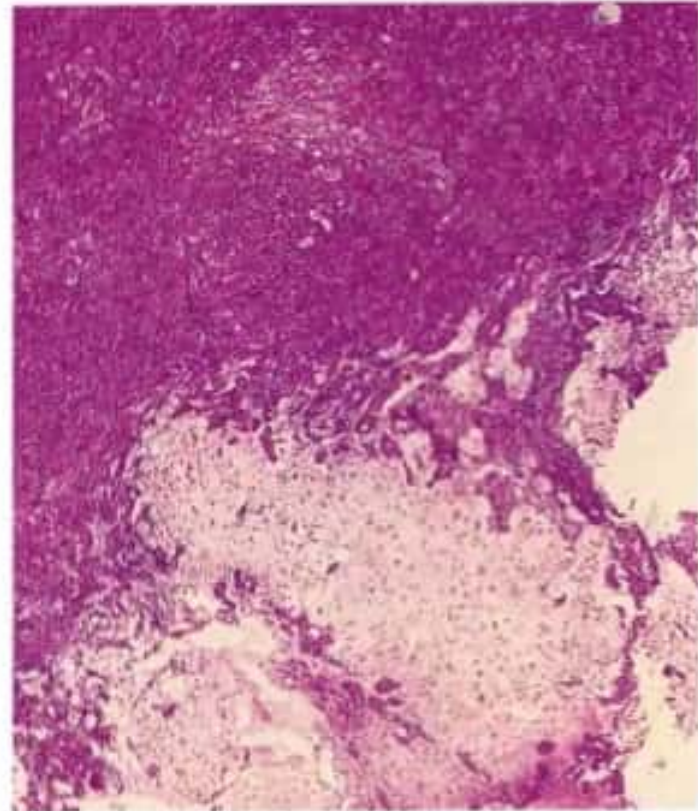
# Carcinosarcoma

- Gross pathology
  - Poorly circumscribed
  - Infiltrative
  - Cystic areas
  - Hemorrhage, necrosis
  - Calcification



# Carcinosarcoma

- **Histology**
  - Biphasic appearance
  - Sarcomatous component
    - Dominant
    - chondrosarcoma
  - Carinomatous component
    - Moderately to poorly differentiated ductal carcinoma
    - Undifferentiated



# Carcinosarcoma

- Treatment
  - Radical excision
  - Neck dissection
  - Postoperative XRT
  - Chemotherapy (distant metastasis to lung, liver, bone, brain)
- Prognosis
  - Poor, average survival less than 2 ½ years

# Squamous Cell Carcinoma

- 1.6% of salivary gland neoplasms
- 7<sup>th</sup>-8<sup>th</sup> decades
- M:F = 2:1
- MUST RULE OUT:
  - High-grade mucoepidermoid carcinoma
  - Metastatic SCCA to intraglandular nodes
  - Direct extension of SCCA

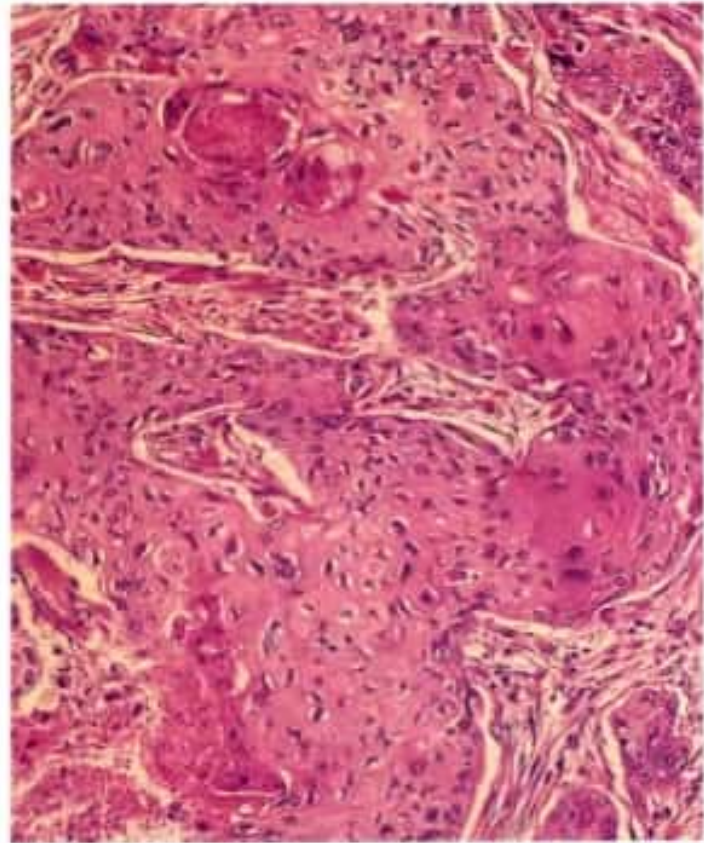
# Squamous Cell Carcinoma

- Gross pathology
  - Unencapsulated
  - Ulcerated
  - fixed



# Squamous Cell Carcinoma

- Histology
  - Infiltrating
  - Nests of tumor cells
  - Well differentiated
    - Keratinization
  - Moderately-well differentiated
  - Poorly differentiated
    - No keratinization



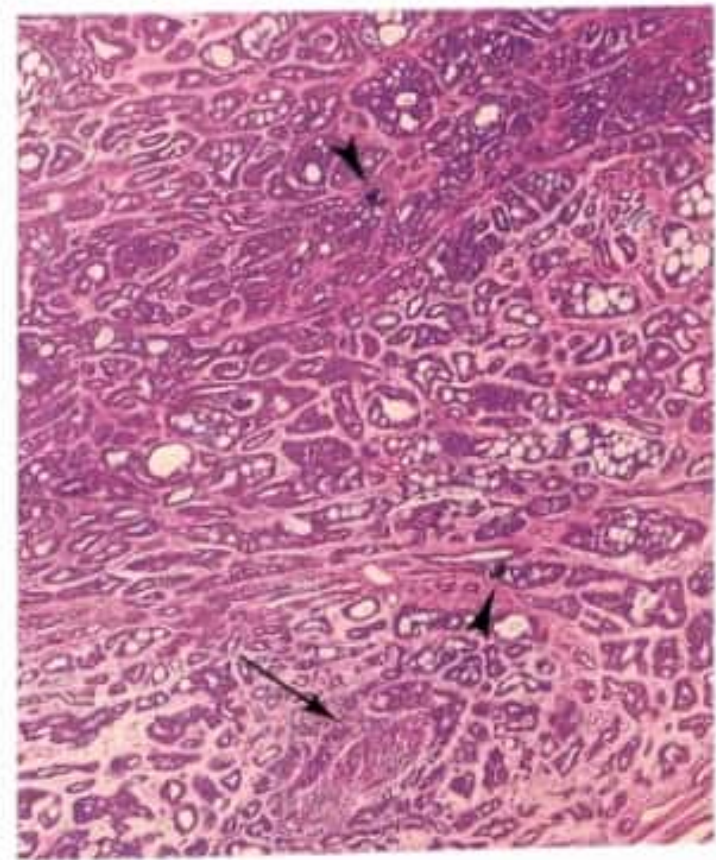
# Squamous Cell Carcinoma

- Treatment
  - Radical excision
  - Neck dissection
  - Postoperative XRT
- Prognosis
  - 5-year survival: 24%
  - 10-year survival: 18%



# Polymorphous Low-Grade Adenocarcinoma

- 2<sup>nd</sup> most common malignancy in minor salivary glands
- 7<sup>th</sup> decade
- F > M
- Painless, submucosal mass
- Morphologic diversity
  - Solid, glandular, cribriform, ductular, tubular, trabecular, cystic



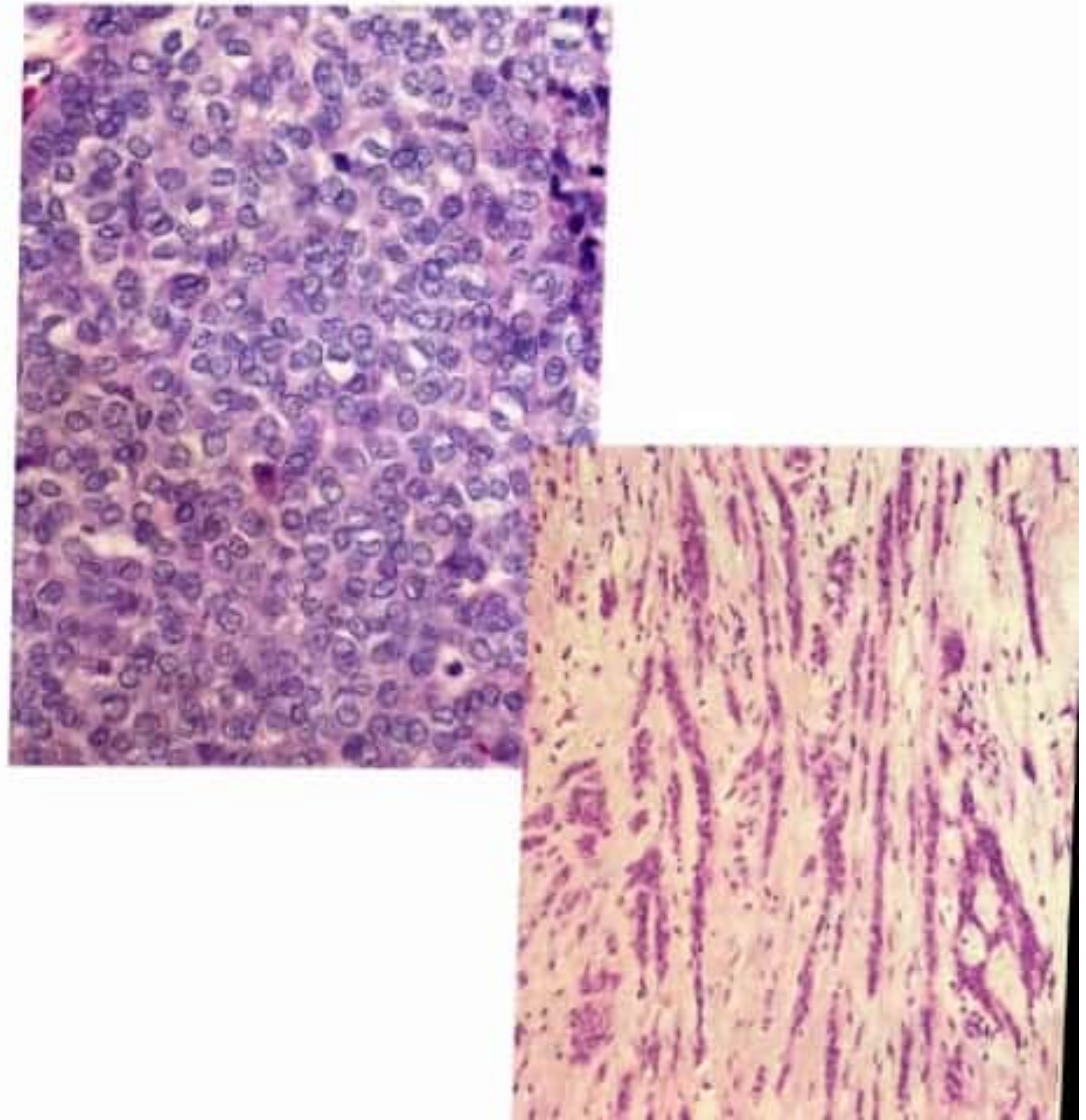
# Polymorphous Low-Grade Adenocarcinoma

- Histology

- Isomorphic cells, indistinct borders, uniform nuclei
- Peripheral “Indian-file” pattern

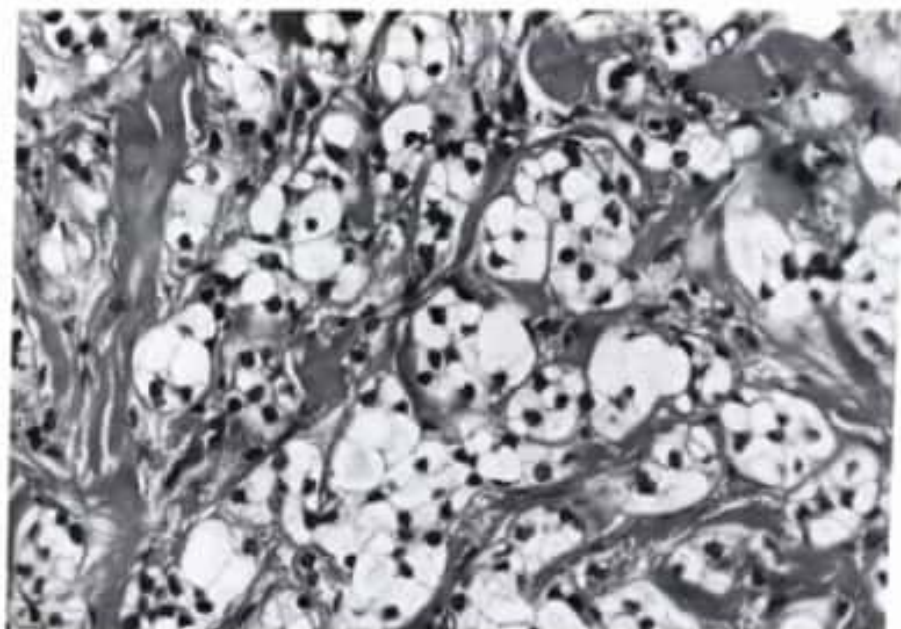
- Treatment

- Complete yet conservative excision



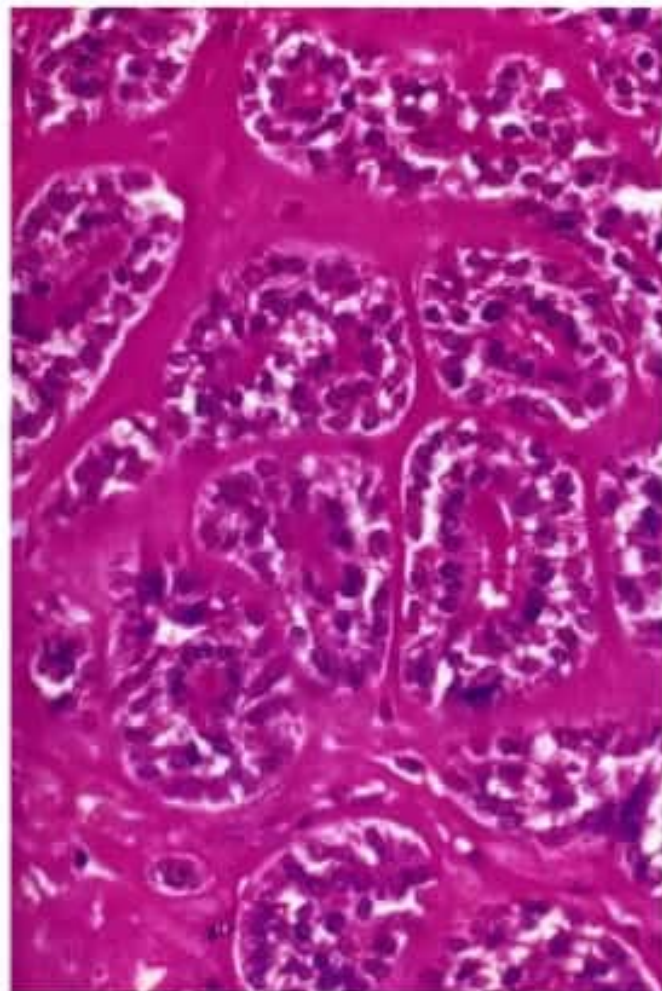
# Clear Cell Carcinoma

- AKA glycogen-rich
- Palate and parotid
- 6<sup>th</sup>-8<sup>th</sup> decade
- M = F
- Histology
  - Uniform, round or polygonal cells
  - Peripheral dark nuclei
  - Clear cytoplasm
- Treatment
  - Complete local excision



# Epithelial-Myoepithelial Carcinoma

- < 1% of salivary neoplasms
- 6<sup>th</sup>-7<sup>th</sup> decades, F > M, parotid
- ? Increased risk for 2<sup>nd</sup> primary
- Histology
  - Tumor cell nests
  - Two cell types
  - Thickened basement membrane
- Treatment
  - Surgical excision



# Undifferentiated Carcinoma

- **Lymphoepithelial**

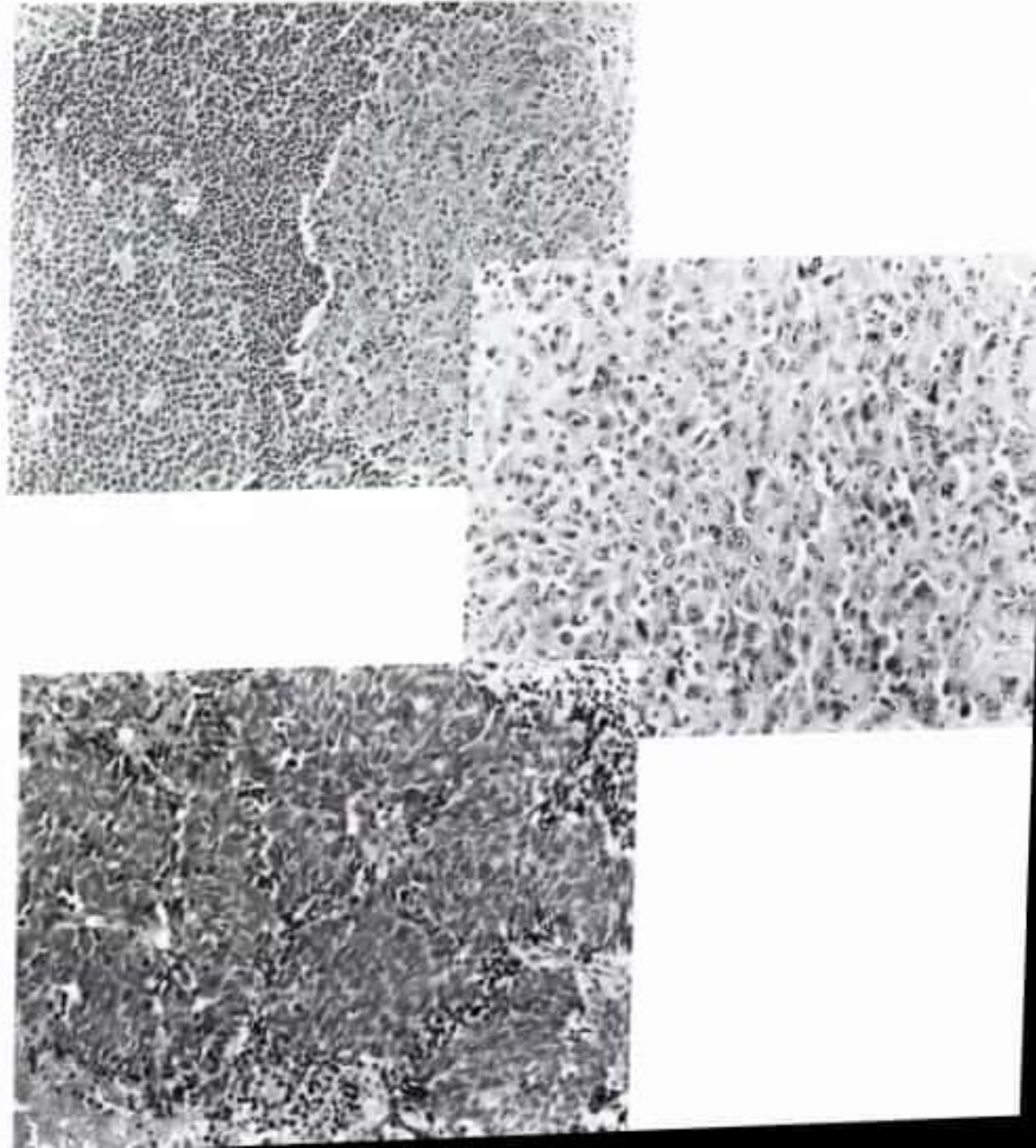
- Eskimos: parotid, F > M, familial
- Asian: submandibular, M > F

- **Large-cell**

- Bimodal peaks
- M > F
- Parotid

- **Small-cell**

- 6<sup>th</sup>-7<sup>th</sup> decades
- M:F = 1.6:1
- parotid



# Controversial Issues

- **Management of the N0 Neck:**

- Recurrence in the neck = low likelihood of salvage
- Parotid: clinical neck disease, 16%
  - N- disease = 74% 5-year survival
  - N+ disease = 9% 5-year survival
- Submandibular: clinical neck disease, 8%
  - N- disease = 41% 5-year survival
  - N+ disease = 9% 5-year survival

# Management of the N0 Neck

- **Increase risk of occult neck metastasis:**
  - \*\*High-grade malignancies
  - \*\*Advanced primary tumor stage (T3-T4)
  - High risk histology
    - Undifferentiated, SCCA, adenocarcinoma, high-grade mucoepidermoid, salivary duct carcinoma
  - Tumor size > 3cm
  - Patient > 54 years of age
  - Facial paralysis
  - Extracapsular, perilymphatic spread

# Management of the N0 Neck

- **Neck Dissection**

- Advantages

- Pathologic staging

- Improved counseling and prediction of prognosis

- Disadvantages

- Longer OR time, increase complications, increased cost

- Functional deficits, cosmetic effects

- Type

- Parotid: levels II-IV

- Submandibular: levels I-III



# Management of the N0 Neck

- **Radiation Therapy**

- Advantage

- Avoids surgical sequelae

- Disadvantages

- Radiation effect on normal tissue
    - Radiation induced malignancies

- Proponents argument: the same factors that increase the risk of occult neck disease also increase the risk for local recurrence and necessitate postoperative XRT to the primary so it is reasonable to treat the neck with XRT as well

# Management of the N0 Neck

- **Radiation Therapy**

- Advantage

- Avoids surgical sequelae

- Disadvantages

- Radiation effect on normal tissue
    - Radiation induced malignancies

- Proponents argument: the same factors that increase the risk of occult neck disease also increase the risk for local recurrence and necessitate postoperative XRT to the primary so it is reasonable to treat the neck with XRT as well

# Fine-Needle Aspiration Biopsy - Facts

- Efficacy is well established
- Accuracy = 84-97%
- Sensitivity = 54-95%
- Specificity = 86-100%
- Safe, well tolerated

# Fine-Needle Aspiration Biopsy

- **Opponents argument (advise against):**
  - Doesn't change management
    - Surgery regardless of reported diagnosis
  - Obscuring final pathologic diagnosis
  - Frequency of "inadequate" sampling, requires multiple biopsies, prolongs course until definitive treatment, increases cost

# Fine-Needle Aspiration Biopsy

- **Proponent's argument (advise for/recommend):**
  - Important to distinguish benign vs. malignant nature of neoplasm
  - Preoperative patient counseling
  - Surgical planning
  - Differentiate between neoplastic and non-neoplastic processes
    - Avoid surgery in large number of patients

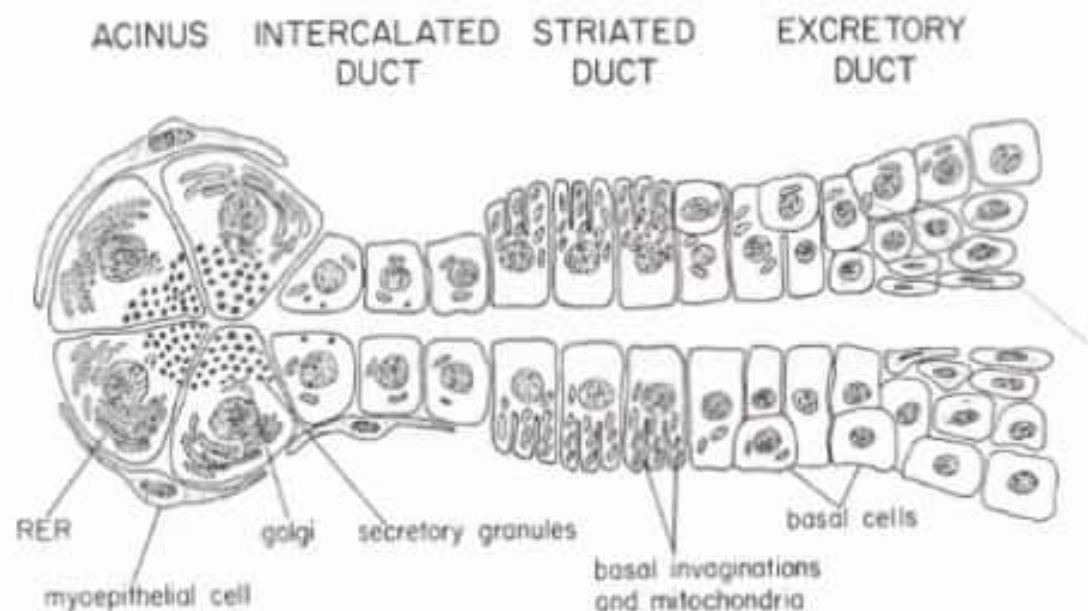
# Tumorigenesis: Bicellular Theory

- Intercalated Ducts

- Pleomorphic adenoma
- Warthin's tumor
- Oncocytoma
- Acinic cell
- Adenoid cystic

- Excretory Ducts

- Squamous cell
- Mucoepidermoid



# Tumorigenesis

- Contradictory evidence:
  - Luminal cells are readily capable of replication
  - Acinar cells participate in gland regeneration
  - Immunohistochemical staining of S-100 protein
    - Present in many salivary gland neoplasms
    - Not present in normal ductal cells

# Conclusions

- Hugely diverse histopathology
- Accurate pathologic diagnosis does influence management
- Relatively rare malignancies
- Utilize preoperative studies when indicated