



Tumors of lung and pleura

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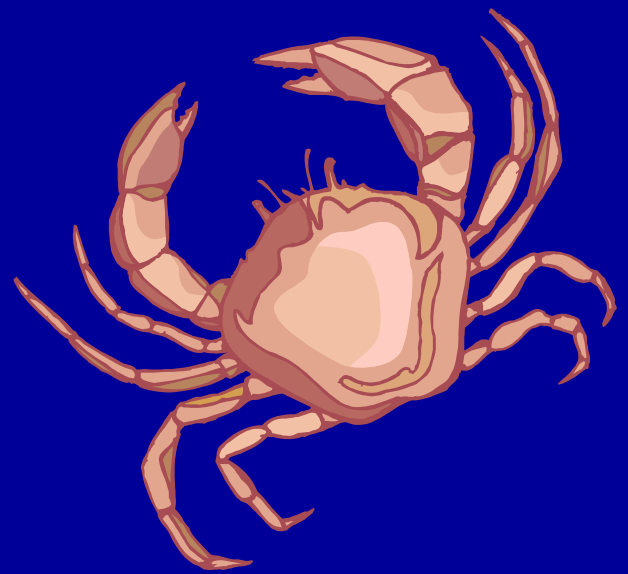
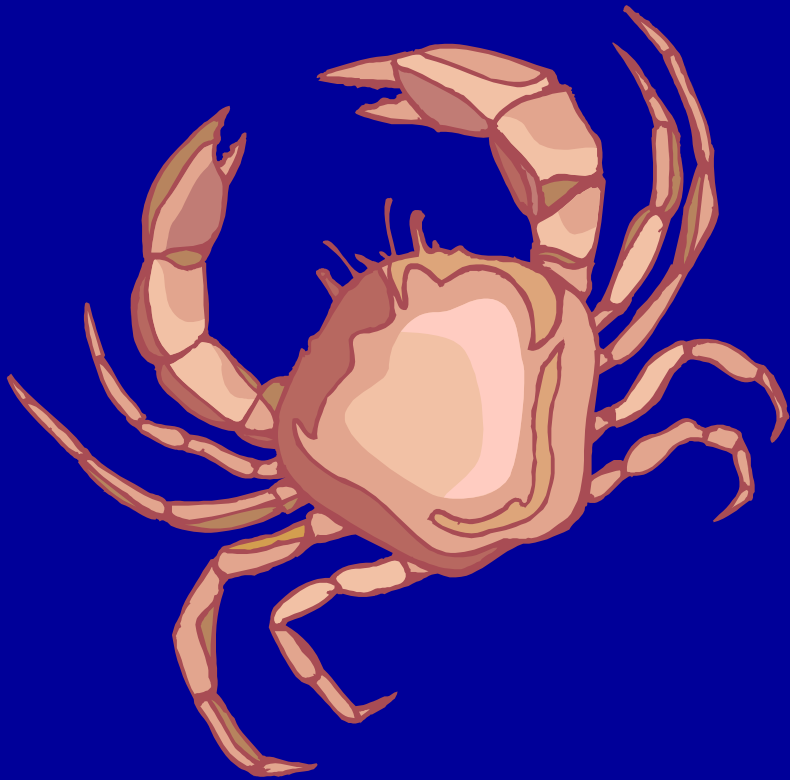
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TUMORS OF LUNG AND PLEURA



Primary Tumors of Lung

- **Carcinoma (90%-95%)**
- **Bronchial carcinoid (5%)**
- **Mesenchymal and other miscellaneous tumors (2%-5%)**

LUNG CANCER

- Near 1/3 mortality from cancer
 - 84% will die within 5 years
 - Most will be diagnosed after age 40 (98%)
 - Cigarette smoking – main risk factor (87%)
 - 75% are centrally located
-

CARCINOMA

Frequent invasion to neighboring organs - •
Pleura, mediastinum.

Involvement of regional lymph nodes – •
50%.

Distant spread – lymphatic or hematogenic. •

Metastases as presenting symptoms – •
frequent.

Bronchogenic carcinoma

Frequent location of metastases: •

Adrenals – more than 50% •

Liver – 30-50% •

Brain – 20% •

Bone – 20% •

Clinical vs histological classification

- **Squamous cell carcinoma (M32%-F25%)**
- **Adenocarcinoma (M37%-F47%)**
- **Large cell carcinoma (M18%-F10%)**
- **Small cell carcinoma (M14%-F18%)**

Non small cell carcinoma

Non squamous non small cell carcinoma (EGFR)

Small cell carcinoma

10% - combined carcinoma

Precursor lesions

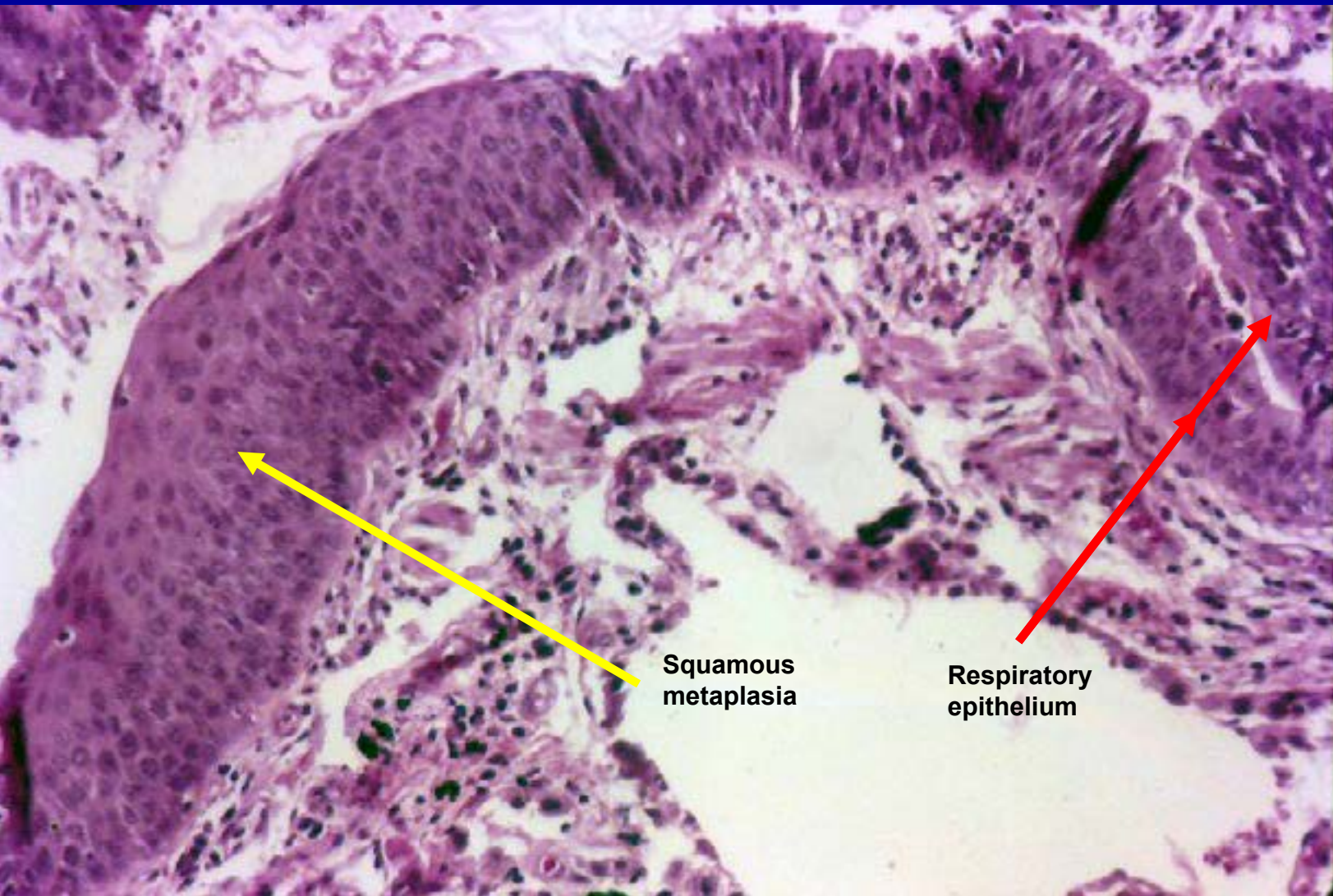
- Squamous dysplasia and carcinoma in situ
- Atypical adenomatous hyperplasia
- Diffuse idiopathic pulmonary neuroendocrine cell hyperplasia

A precursor condition does not always imply a cancer.
Its transformation is rarely preconceived.

DEVELOPMENT OF BRONCHIAL CARCINOMA

- **Area of cytological atypia**
 - **Thickening of bronchial mucosa**
 - **Warty excrescence and erosion**
 - a. Fungating into bronchial lumen
 - b. Infiltration along peribronchial wall
 - c. Producing intraparenchymal mass
-

Development of squamous cell carcinoma





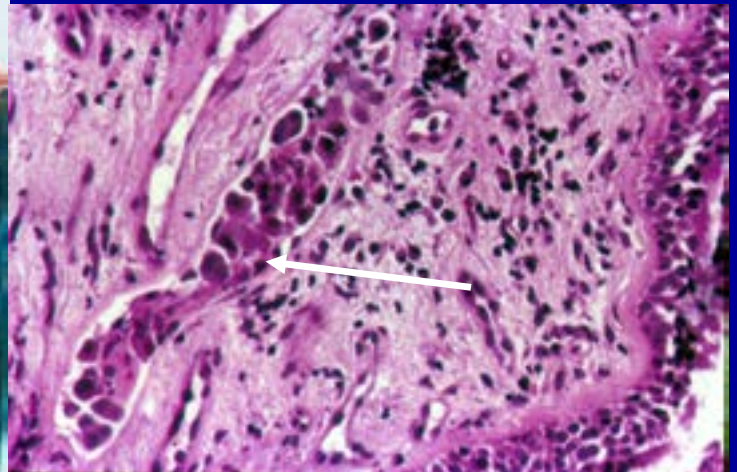
Cytological atypia / dysplasia

Bronchogenic carcinoma

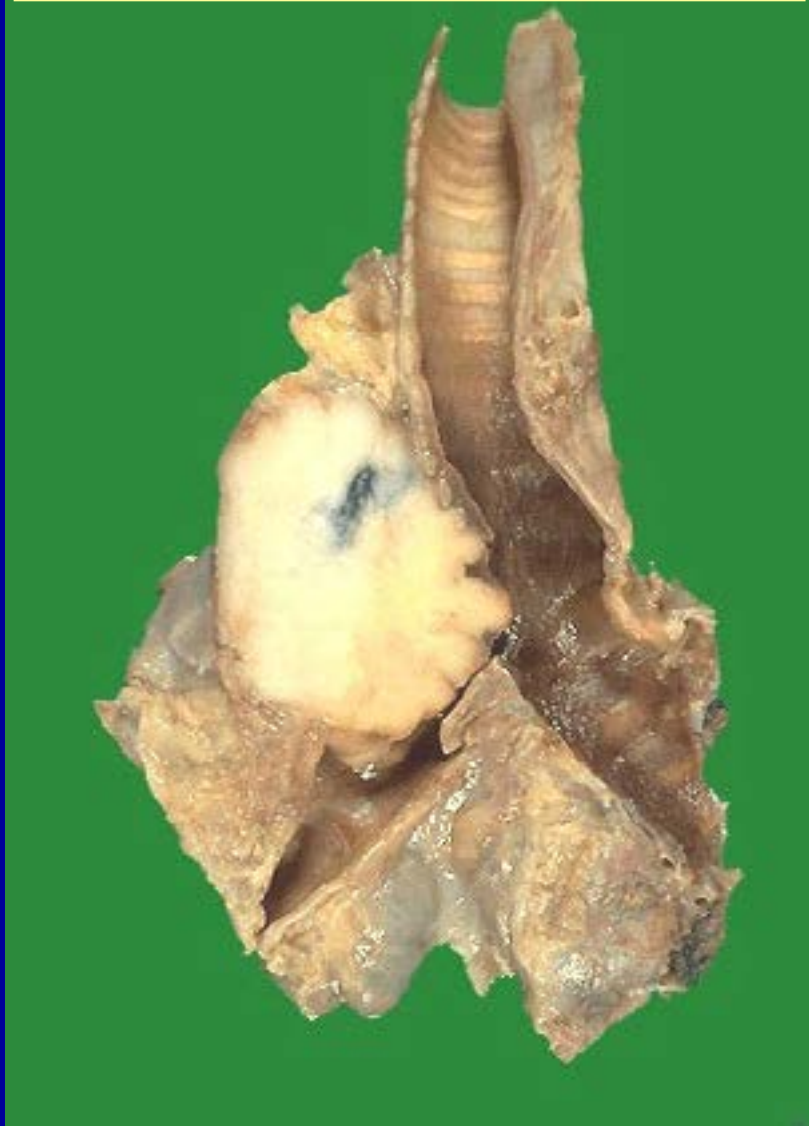




**Peribronchial
infiltration**



Squamous cell carcinoma



90%
central

**Frequent
P53
mutations**

**IHC
supporting**

P63

CK 5-6

**overexpressio
80%EGFR-
mutations are
rare**

*Cigarette
smoking*

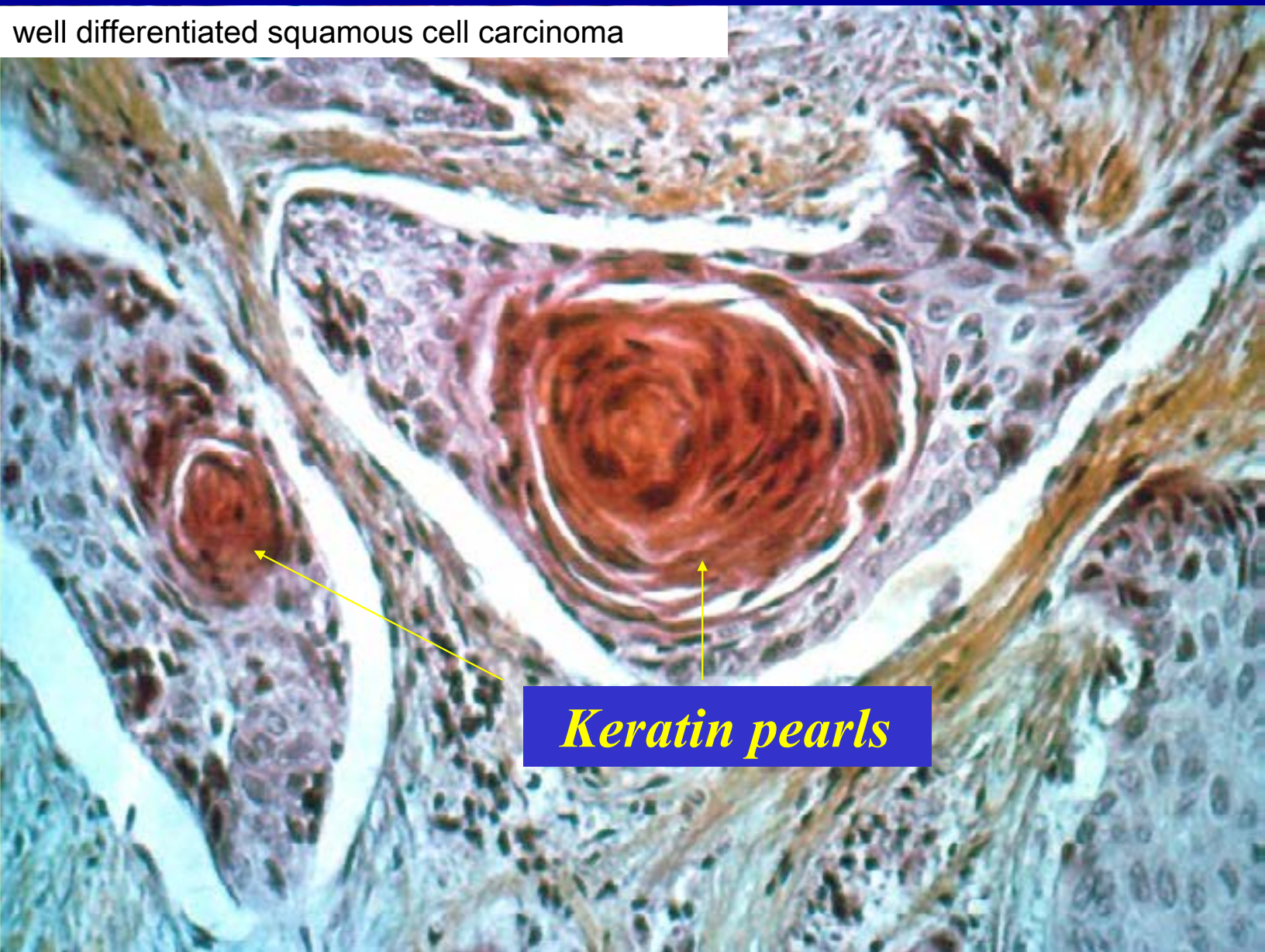
Squamous cell carcinoma

Histopathological characteristics:

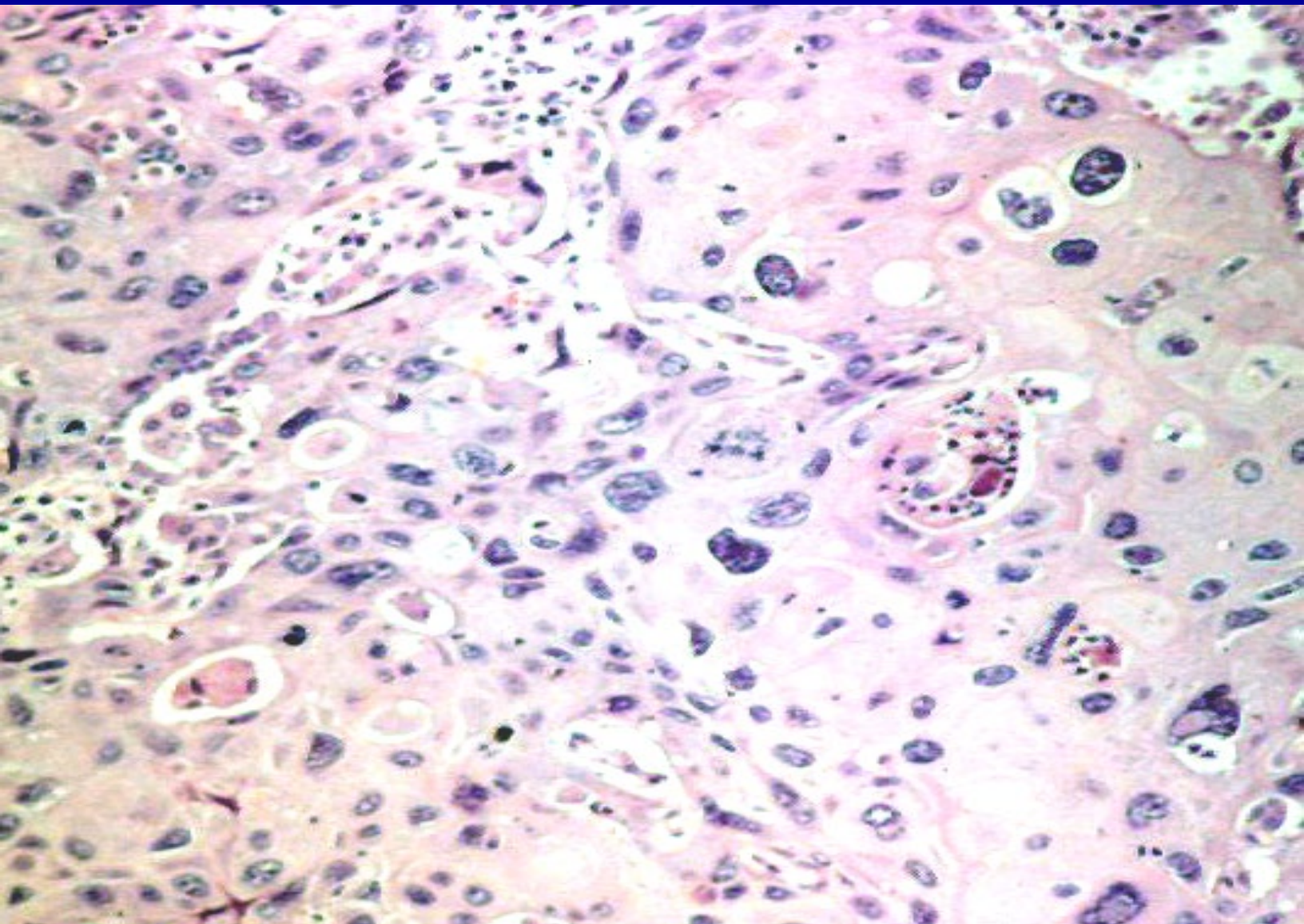
Keratin formation.

Marked intercellular bridges
(desmosomes).

well differentiated squamous cell carcinoma



Keratin pearls



Poorly differentiated squamous cell carcinoma



Desmosomes

Adenocarcinoma

- **Usual Adenocarcinoma**
- **Bronchioloalveolar carcinoma**

No link with smoking ❖

Good prognosis ❖

Adenocarcinoma (Usual)

mutations

K-RAS

EGFR

*Mostly in
women and
non-smokers*

TTF1 NAPSIN
A

Supporting
IHC

peripheral 75%



Next to scar

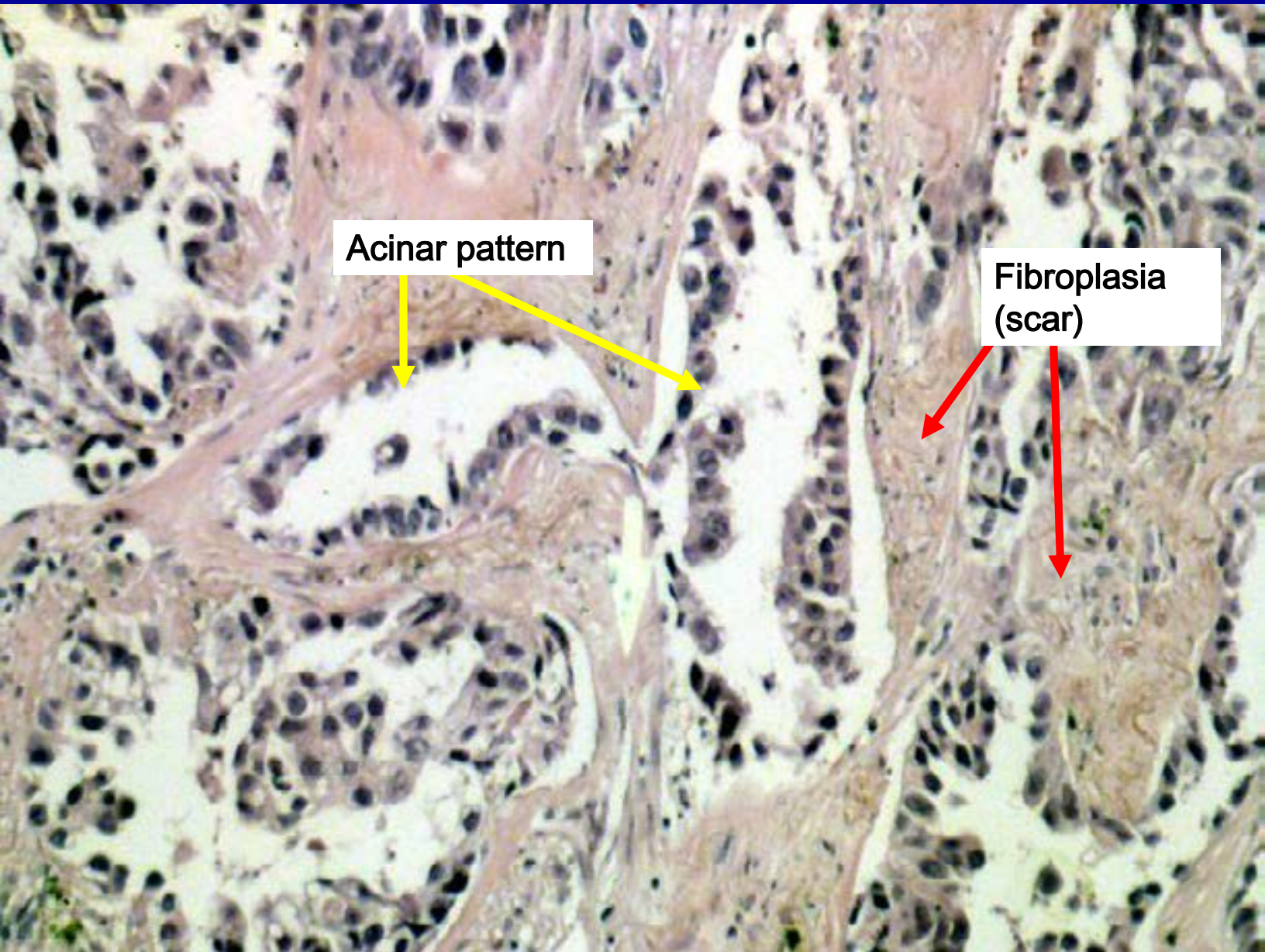
Usual Adenocarcinoma

ACINAR –

PAPILLARY –

SOLID –

Glandular differentiation or mucus formation. –

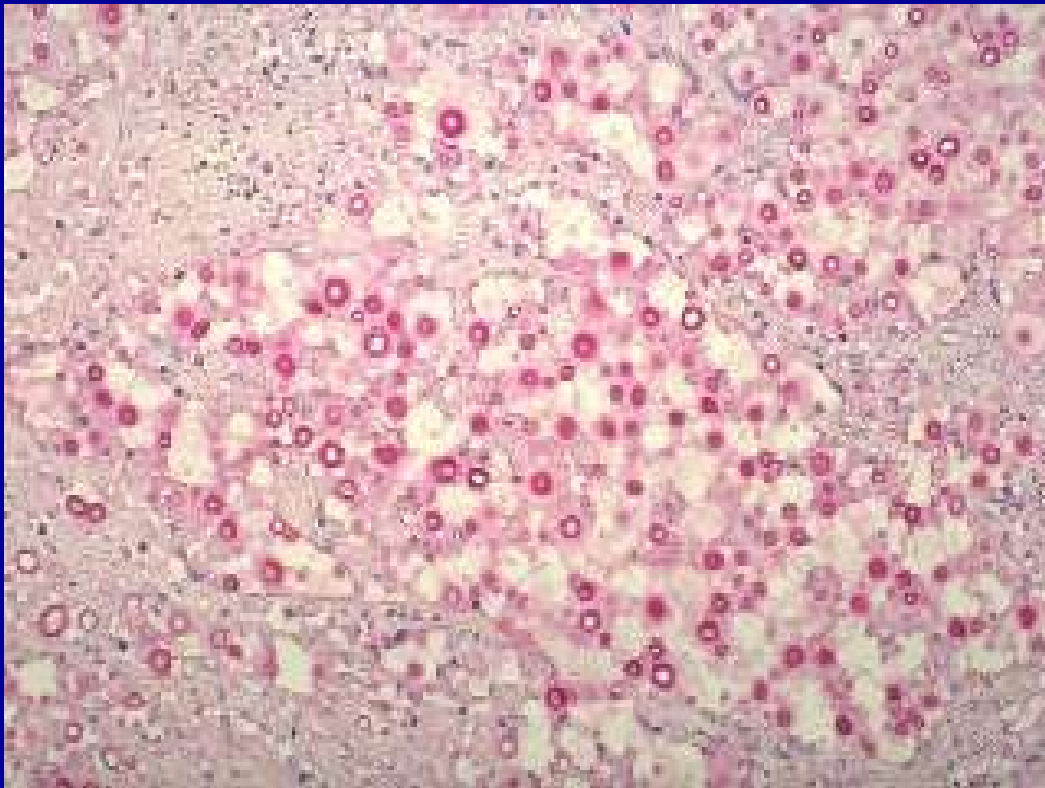


Acinar pattern

Fibroplasia (scar)

Mucous staining- (Mucicarmine)

80%Positive



Bronchioloalveolar carcinoma

1-9% of lung cancers –

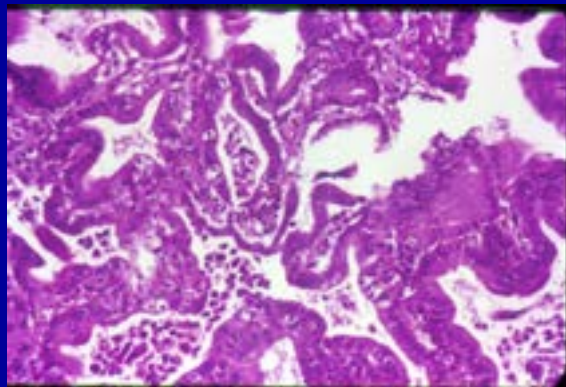
Nodular formation –

Or pneumonia-like consolidation. –

HISTOPATHOLOGY: –

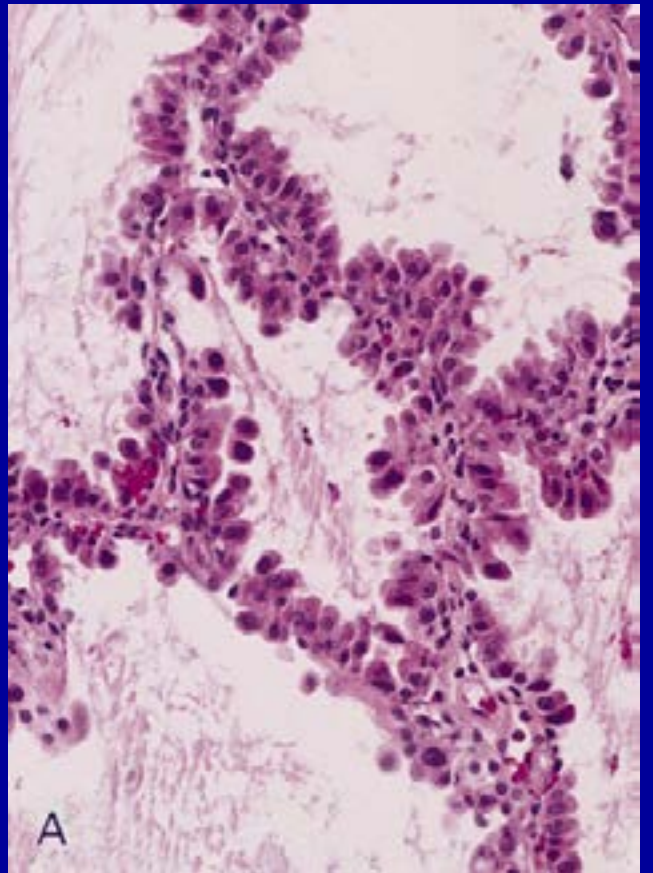
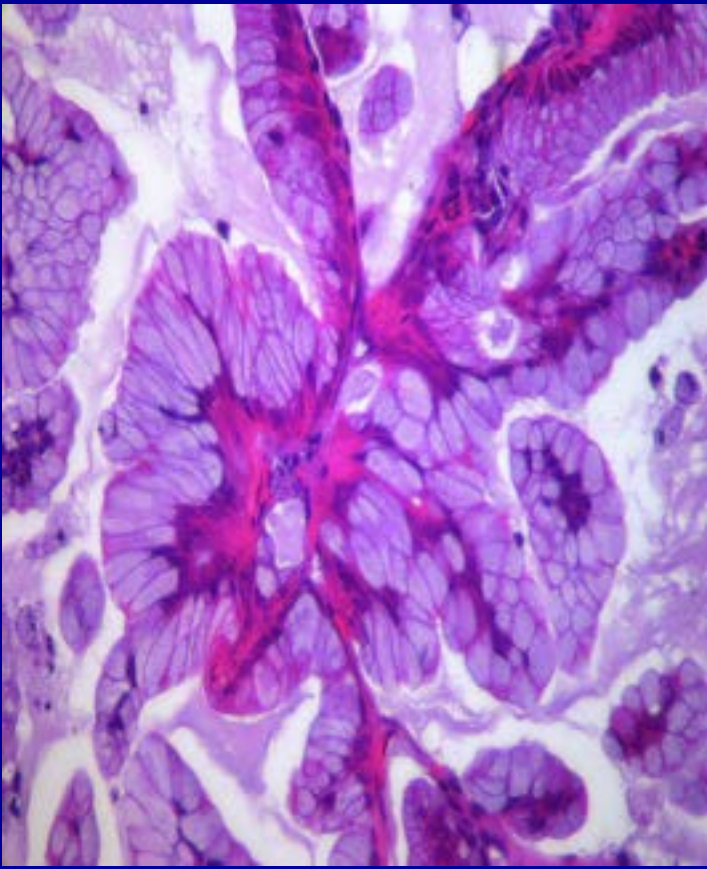
Spreads along preserved alveoli (Lepidic) –

Composed of cuboidal or columnar cells. –



Bronchioloalveolar carcinoma

- **Non mucinous** (adenocarcinoma in situ)
 - Columnar, peg shaped, cuboidal
 - Peripheral lung nodule
- **Mucinous**
 - Tall columnar with cytoplasmatic and intra alveolar mucin
 - Aerogenously spread forming satellite tumors
 - May resemble lobar pneumonia



A

Non-Small Cell Lung Cancer (NSCLC)

NSCLC differs from Small Cell Lung Cancer by a variety of histomorphological features, by a diverse biological activity, and therefore, by a changing response to chemotherapy, to biological and to radiotherapy. •

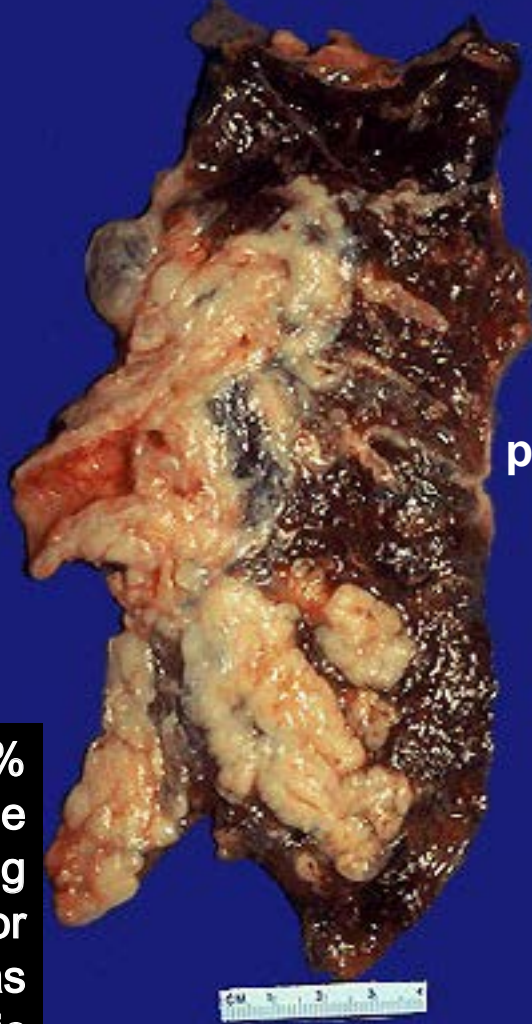
A standard NSCLC case, displaying metastases, a positive expression for PD-L1, and no abnormal EGFR or ALK expression, may respond best to biological therapy with nivolumab, and with ipilimumab. •

SMALL CELL LUNG CANCER (SCLC)

Small cell lung cancer is highly sensitive to radiotherapy. However, this mode of treatment is insufficient, due to the very high stage of this malignancy at diagnosis. Therefore, it is highly advised to append treatment with EGFR tyrosine kinase inhibitor, as well as with an antiangiogenic mode of therapy.



Small cell carcinoma



Central
or
peripheral

75%
hormone
producing
or
paraneoplastic
syndromes

99% -Link with smoking

Small cell carcinoma

(POORLY DIFFERENTIATED NEUROENDOCRINE CARCINOMA)

Microscopic features:

Small epithelial cells, scant cytoplasm, fine granular nuclear chromatin

High mitotic activity

Extensive necrosis

Electron microscopic studies:

Dense-core neurosecretory granules in some of tumor cells

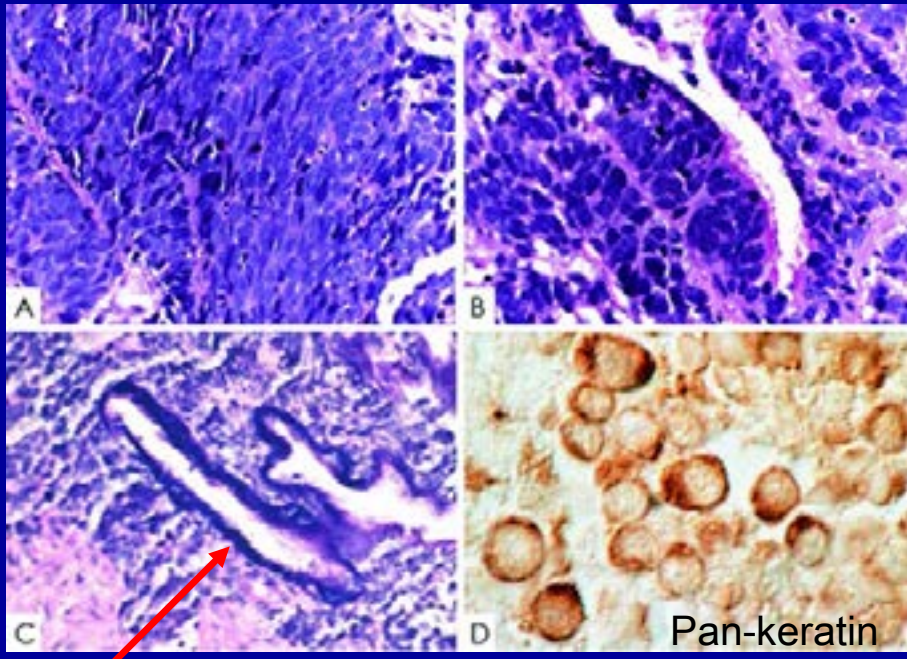
Molecular studies :

Mutant expression: 80-100% RB, 50-80% p53

High levels of BCL-2 (anti apoptotic) - 90%

Reduction of BAX (proapoptotic gene) expression

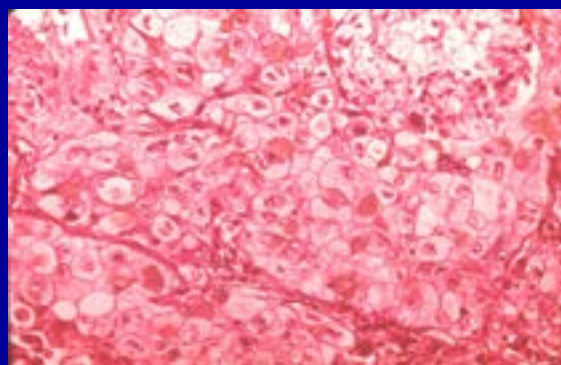
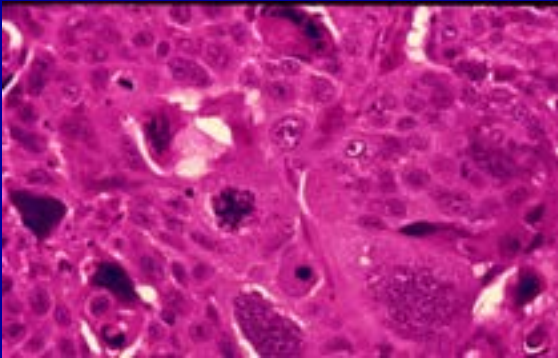
Small cell carcinoma



Azzopardi effect

Large cell carcinoma

Composed of large, undifferentiated, malignant cells.
A neuroendocrine variant is described.



Bronchial Carcinoid

No relation with smoking; good prognosis; mostly below age 40.

May present as a nodule in the bronchial wall.

Classification:

Typical carcinoid (well differentiated neuroendocrine carcinoma)

Atypical carcinoid (moderately differentiated neuroendocrine carcinoma)

Bronchial Carcinoid

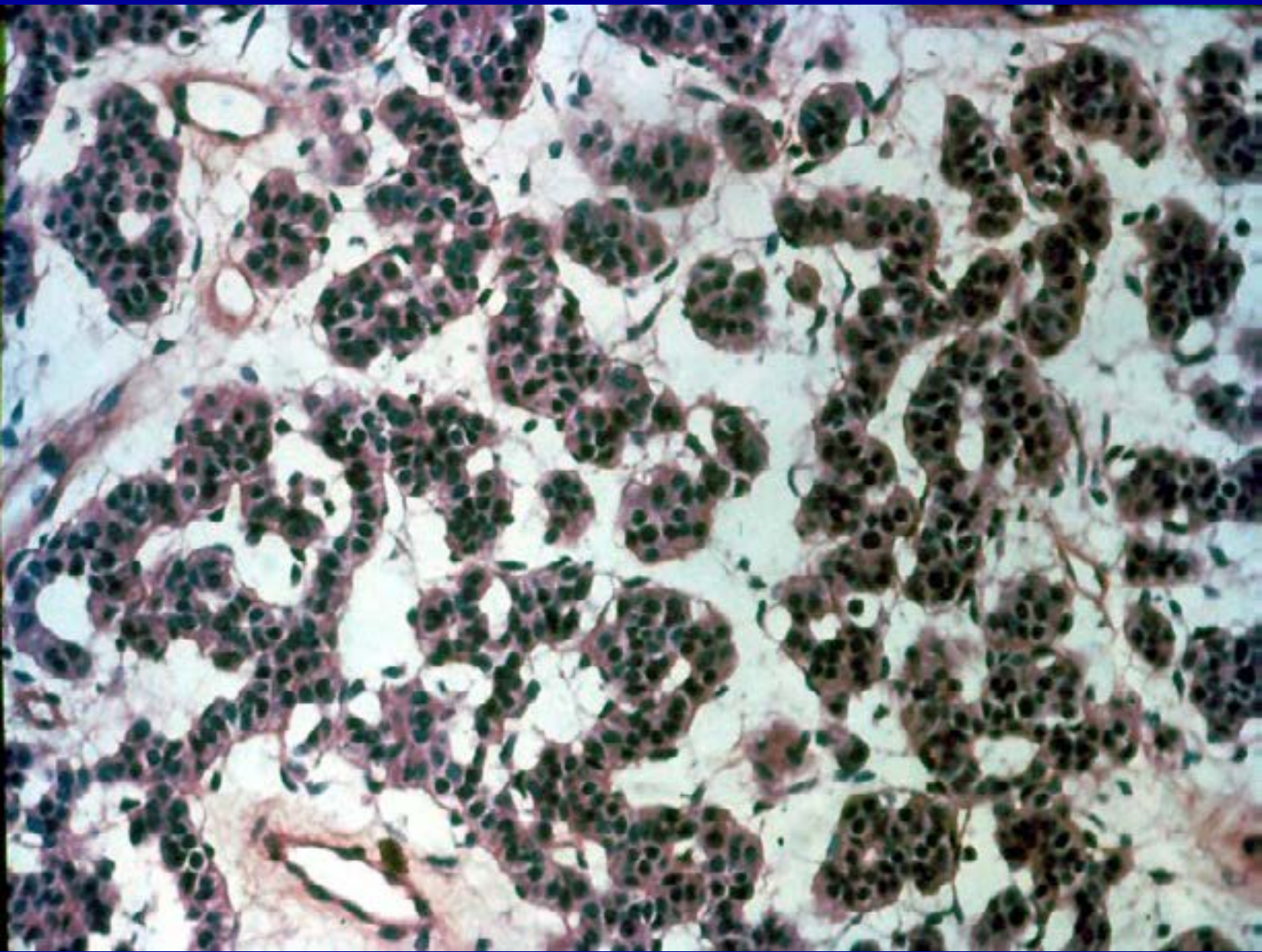
MICROSCOPIC FEATURES:

Organoid (growing organ-like).

Surrounded by delicate connective tissue.

Nests or cords, with uniform cells.

Positive for neuroendocrine markers.

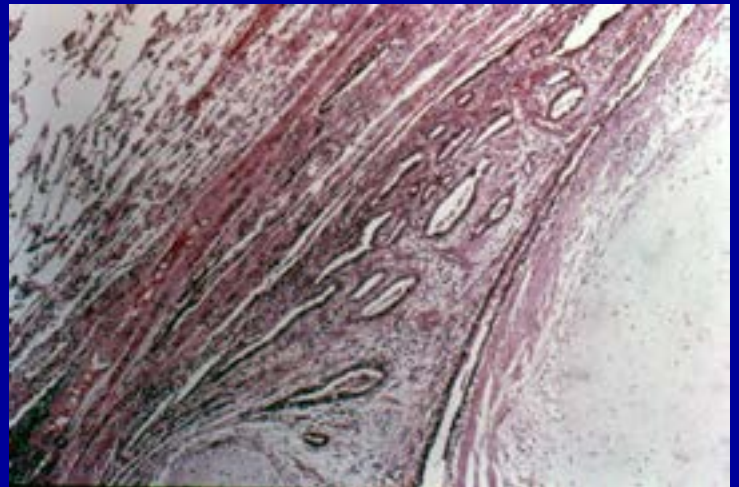
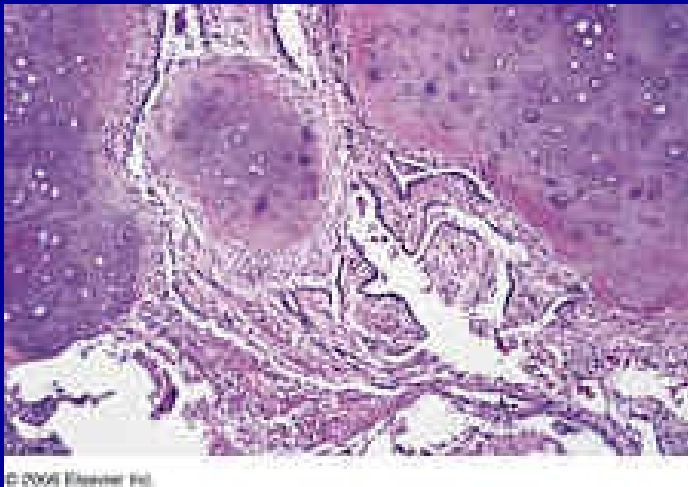


Neuroendocrine carcinoma	5y survival (%)	10y survival (%)
Typical Carcinoid	87	87
Atypical carcinoid	56	35
Large cell neuroendocrine carcinoma	27	9
Small cell carcinoma	9	5

Pulmonary Hamartoma



Pulmonary Hamartoma

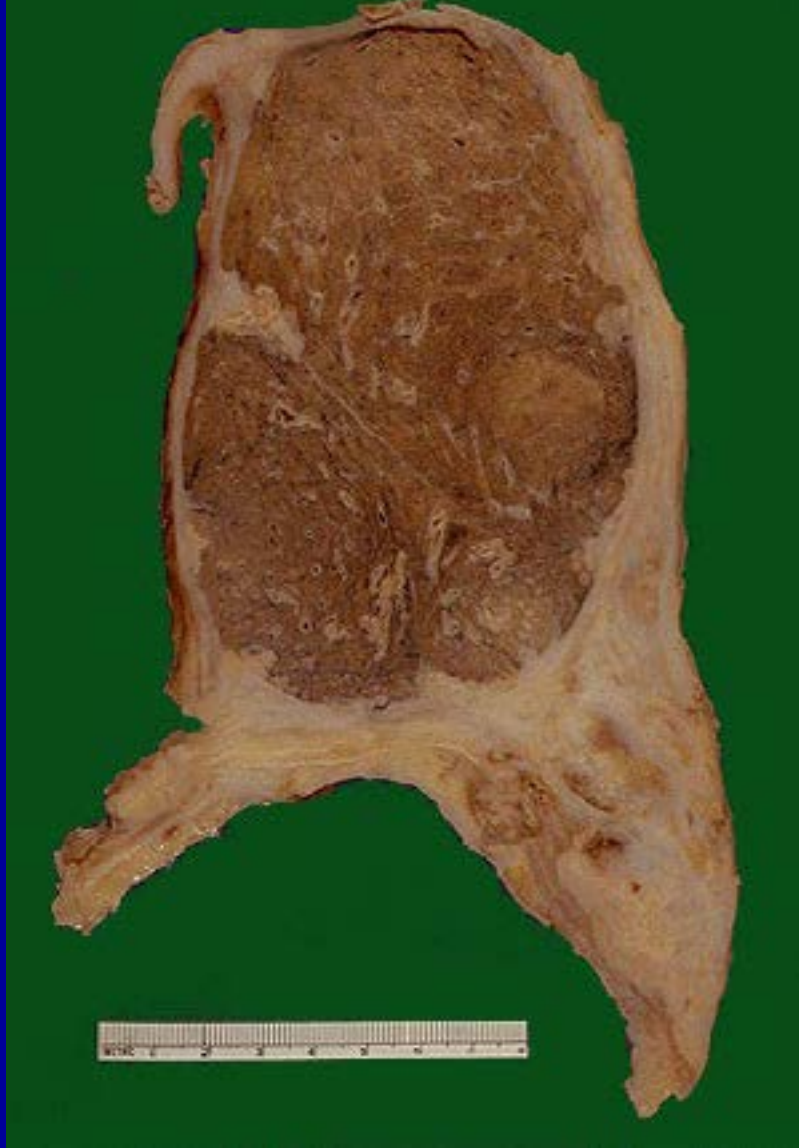


Pleural Tumors

MALIGNANT MESOTHELIOMA



50% die within one year



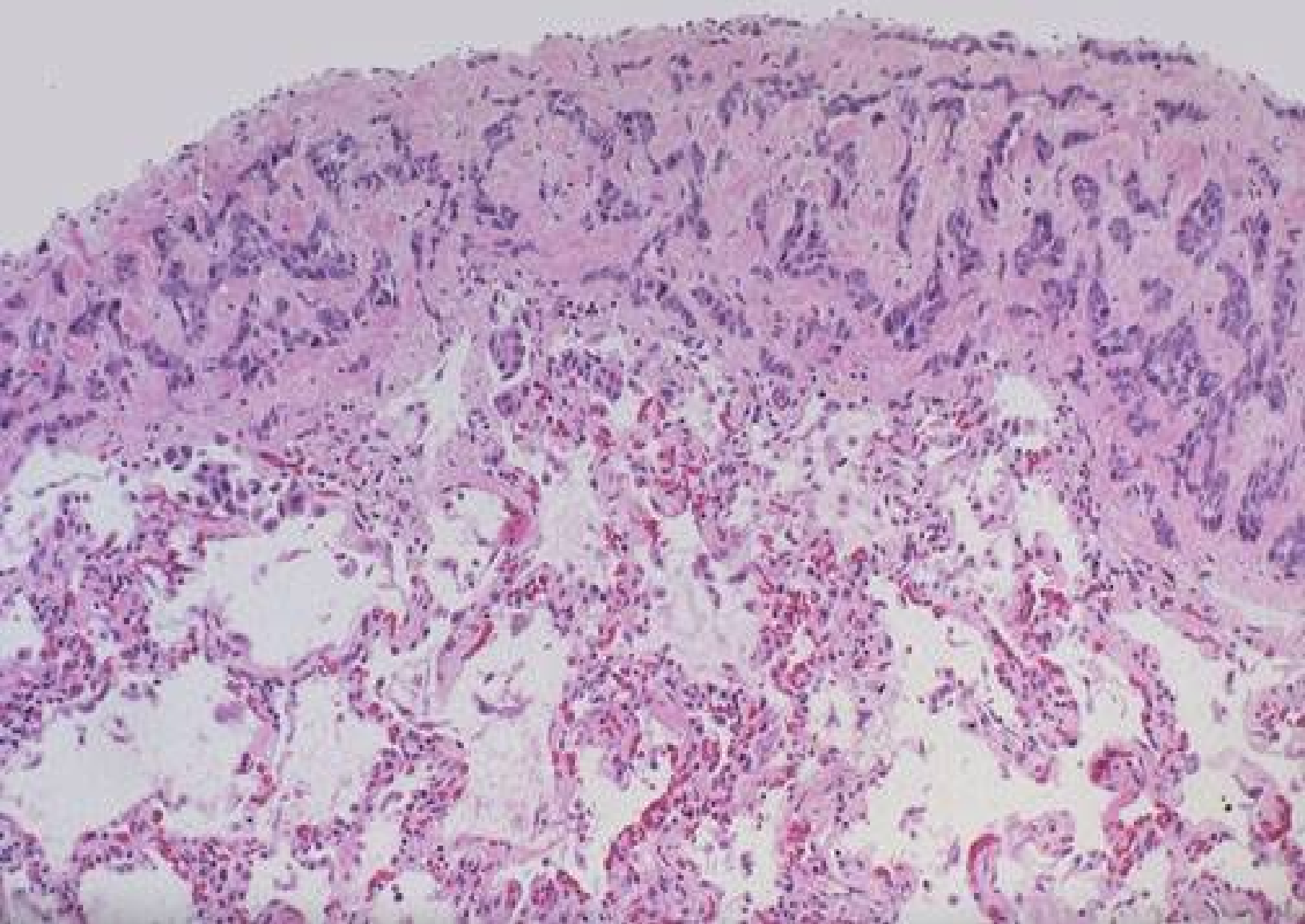
MALIGNANT MESOTHELIOMA

Microscopic types:

- **Mesenchymal (20%)** (sarcomatoid) – spindle cells
- **Epithelioid (60%)** – forming tubular and papillary structures
- **Mixed (20%)**

E.M examination:

- Long microvilli (gold standard)
-



MALIGNANT MESOTHELIOMA

