CYTOPATHOLOGY

outlines:

- 1- Definition of cytopathology.
- 2- Differences between cytopathology and histopathology.
- 3- Advantage and disadvantage of cytopathology.
- 4- Sampling techniques.
- 5- Indications of cytopathology.

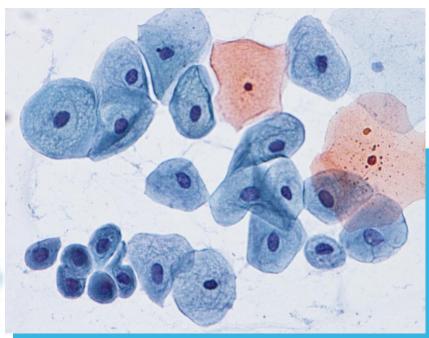
DEFINITION

Scientific study of cells obtained from tissues or body secretions to identify disease.

The individual cells reflect the normal and abnormal morphology of the tissue from which they are derived.

Hmm....I examine the structure of few separated cells NOT an intact tissue.





TYPES

Based on sampling techniques, cytology is classified into the following:

- 1. Exfoliative Cytology.
- 2. Abrasive Cytology.
- 3. Aspiration Cytology.

Sampling techniques

The collection of exfoliated cells "Exfoliative Cytology".

Brushing or similar abrasive techniques "Abrasive Cytology"

Aspiration biopsy

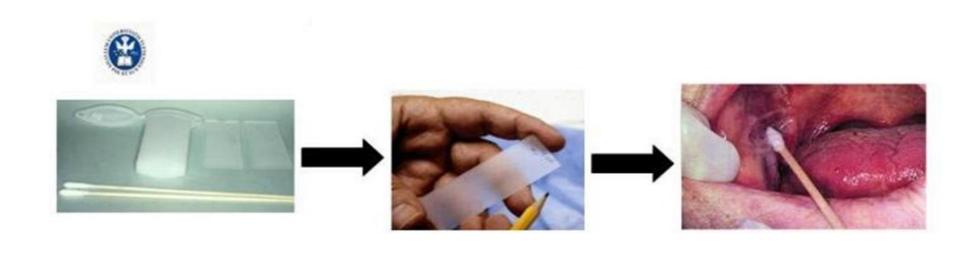
EXFOLIATIVE CYTOLOGY(NATURAL SPONTANEOUS EXFOLIATION)

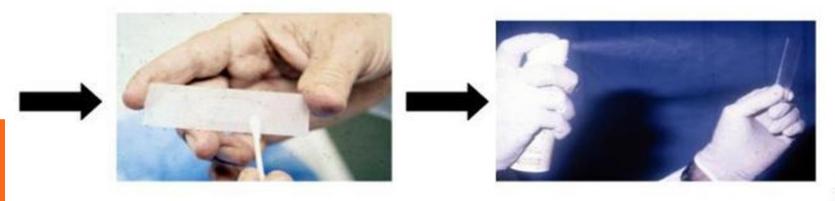
Based on spontaneous shedding of cells derived from the lining of an organ into a cavity.

- It is the simplest of the three sampling techniques.
- Contents of the sample are derived from several sources.

Examples: vaginal smear, sputum, urine, effusion.

The material is collected spontaneously or by a syringe or a cotton swab.







ABRASIVE CYTOLOGY(ARTIFICIAL ENHANCED EXFOLIATION)

- Cells are obtained directly from the surface of the target of interest.
- Samples are taken by :

scraping, : from cervix(pap smear), vagina, oral cavity(Buccal mucosal smear), and skin lesions.

brushing, washing. and lavage: bronchi, GIT, and urinary tract

Examples: cervical scraper, endoscopy, and gastric lavage.

Samples can be obtained from superficial or deep lesions.

The purpose of this procedure is to dislodge cells.... enrich the sample with cells obtained directly from the surface of the target of interest.

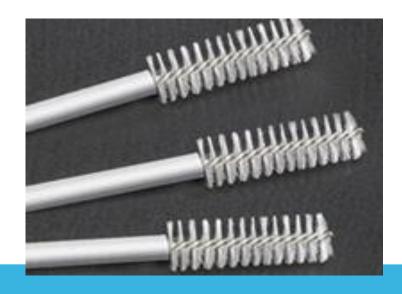


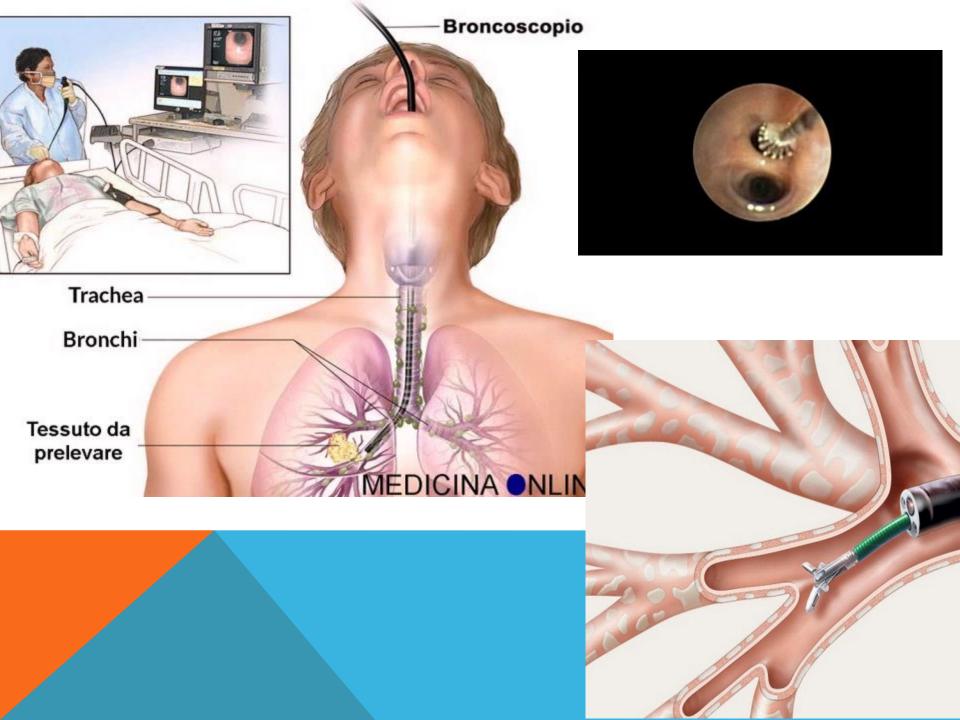












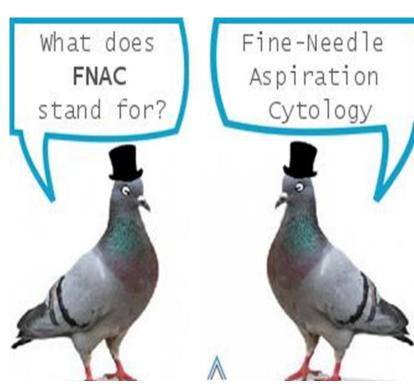
3- Fine Needle Aspiration Cytology (FNAC):

- FNAC provides many advantages to the surgeons being an easy, reliable, cost effective diagnostic technique which could give rapid results.
- The procedure could be performed in outpatient setting without anesthesia.
- ➤ It is usually not more painful than a venipuncture and can be repeated immediately if the acquired material is inadequate.

FNAC

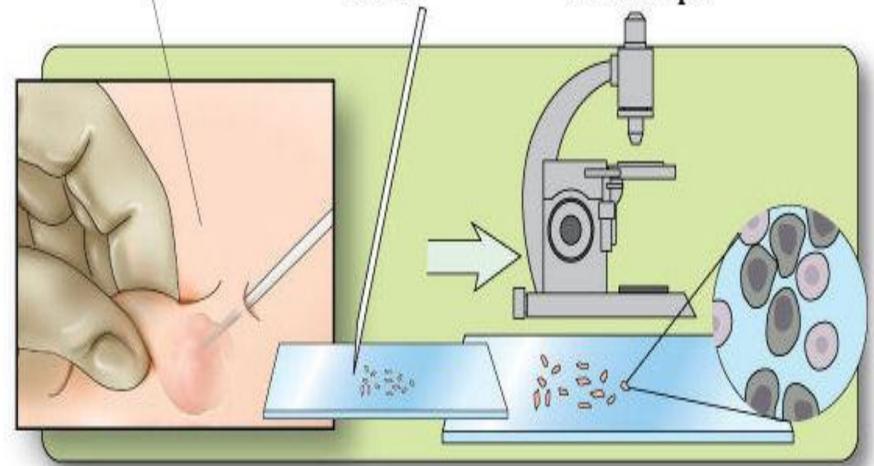
Samples are obtained from solid tissue: body organs, tumors & other swell e.g lymph node, breast and thyroid A needle with a syringe is used. Simple, safe, rapid, cost effective Virtually every organ in the body is

accessible to this method.

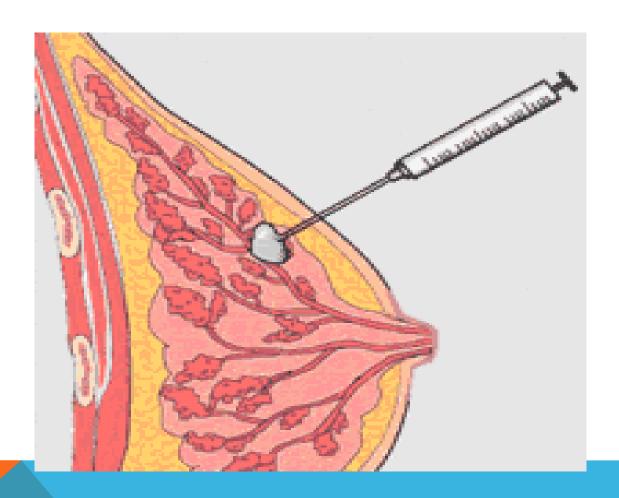


FNAC consists of:
\square Using a needle and syringe to remove material from a mass.
☐ Smearing it on a glass slide.
☐ Applying a routine stain.
\square Examining it under the microscope.

Thin needle draws out small bits of tissue Tissue samples placed on glass slides Tissue samples examined under microscope



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Fixatives used in Cytopathology:

- 1-95% ethyl alcohol (for Pap smear & FNA).
- 2-Spray fixatives (does not result in lysis of RBC & better reserved of nuclear details).
- 3-Carnoy's fixative (lysis of RBC).
- 4-Other (Formaline)

Stains used in cytopathology:

1. Papnicolaou stain:

- Mainly used in Exfoliative cytopathology.
- Good & better demonstration of nuclear details.
- 2. Hematoxyline & Eosin (H& E):
- Mainly used in FNA.
- 3. Leishman & Giemsa stains:
- Mainly in FNA.

Characteristics of ideal stain in cytopathology:

- 1-Evaluation of architectural pattern of tissue fragments.
- 2- Proper evaluation of nuclear morphology (most important characteristic).
- 3- Proper evaluation of cytoplasmic morphology.
- 4- Evaluation of background (blood, secretions, mucin.....)

Staining methods in Cytopathology

- 1.Air dried Giemsa staining:
- Air drying follows by staining with Giemsa).
- Well demonstrated cytoplasmic details.
- Exaggerated cells & nuclear size.
- Poorly seen individual cells.
- 2. Wet -fixed Pap staining:
- Alcohol fixation follows by staining with Pap or H & E.
- Excellent demonstration of nuclear details.
- Normal size of cell & nucleus.
- Clearly seen individual cells.

.Air – dried Giemsa staining:

Wet -fixed Pap staining:

Air drying

staining with Giemsa).

Well demonstrated

cytoplasmic details

Exaggerated cells &

calle

nuclear size.

Poorly seen individual

Alcohol fixation

staining with Pap or H & E

Excellent demonstration of

nuclear details Normal size of cell & nucleus.

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Differences between cytopathology and histopathology

➤ It differs importantly from histopathology in that cytology examines only discrete cells or small groups of cells, but histopathology examines intact tissues.

Histopathology

 Deals with the form and the structure of the tissue.

II. Evaluation usually begins with a tissue biopsy so it is more invasive and traumatic.

Cytopathology

I. Deals with the changes within the nucleus and cytoplasm of cells.

II. Fine needles with small gauge are usually preferred so it is less invasive and traumatic.

III. Basic stain is H&E

III. Basic stain is Pap stain (however H&E could be used as well)

IV. Diagnosis obtained after days.

IV. Rapid diagnosis that could be obtained within minutes.

V. Expensive means of diagnosis do not allow for repetition

V. Inexpensive simple means of diagnosis which allows frequent repetition of cellular sampling.

Advantages of diagnostic cytology:

Superior morphologic detail.



- Ability to characterize the cellular components of various fluids.
- Rapid: Speed of sample collection and diagnosis.
- Inexpensive: Excellent cost effectiveness.
- Simple : Minimally invasive diagnostic procedure not requiring surgery or specialized anesthesia.

Advantages of diagnostic cytology:

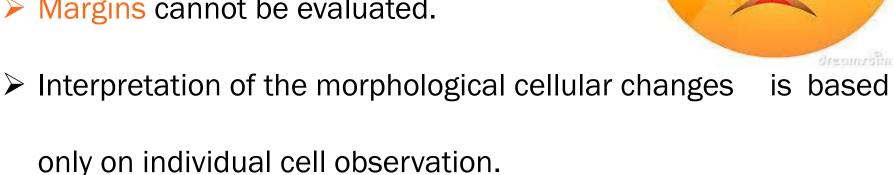
> It is better in evaluating the infectious disease



- No injury to tissue allowing repeated sampling
- ➤ It is better for hormonal assay.
- Cytopathological smear cover a wider surface than that involved in surgical biopsy.

The disadvantage of diagnostic cytology:

- > Tumors cannot be graded.
- Margins cannot be evaluated.



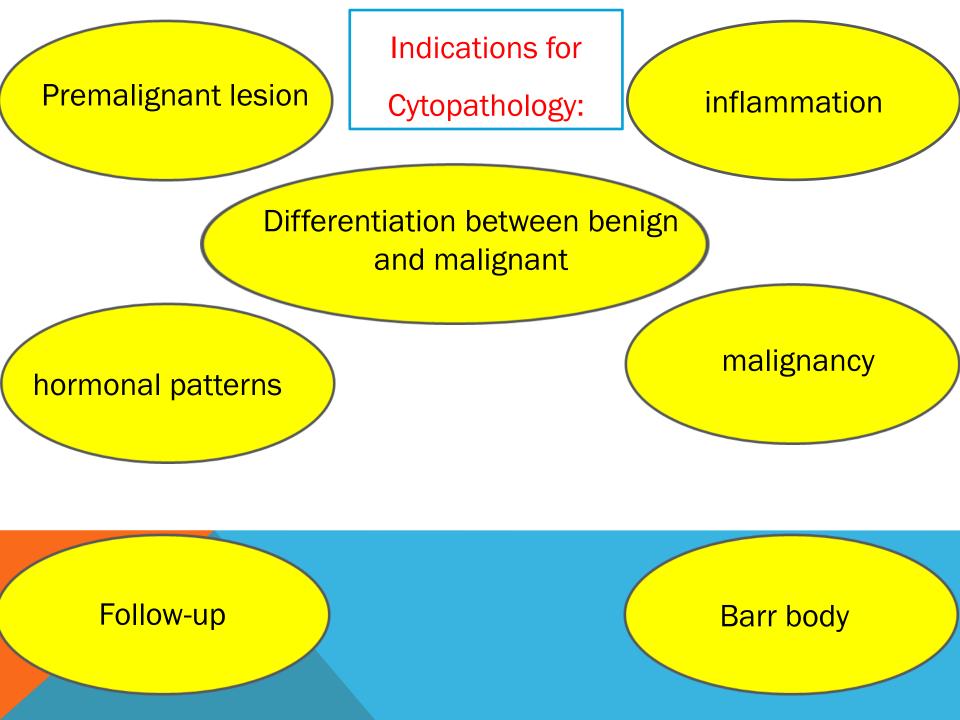
- information Limited about structural arrangement (architecture) of the cells within the lesion.
- Less definitive diagnosis as to the specific tumor type or the distribution of an inflammatory infiltrate.

➤ Very bad negative test: A problem related to the small sample size and "blind" nature of many common collection techniques, is the possibility that the specimen may not represent the primary lesion.

The following points should be kept in mind when starting a cytological evaluation:

➤ 1) Cytology is an aid to clinical diagnosis, and results must be interpreted in the context of the overall case. The cytologic diagnosis must be based on the entire clinical evidence available, rather than on changes in individual cells.

2) Cytology can be but is not always a substitute for histopathology.



Indications for Cytopathology:

- 1- Differentiation between benign and malignant lesions.
- 2- Diagnosis of malignancy and the identification of the neoplastic cells in primary, metastatic (secondary).
- 3- Diagnosis of premalignant diseases.
- 4- Detection of inflammation and certain types of pathogenic agents.

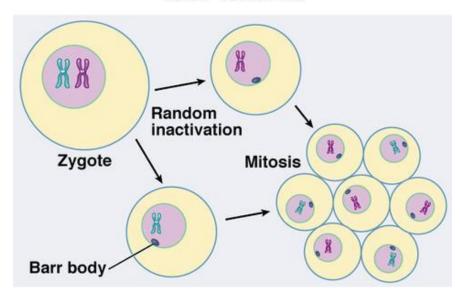
- 5- Study of the hormonal patterns through the examination of the squamous cells in vaginal smears; which are under the influence of ovarian hormones.
- 6- Follow-up and monitoring of response to chemotherapy and irradiation, the latter producing certain cellular features which could be diagnostic on cytological examination.

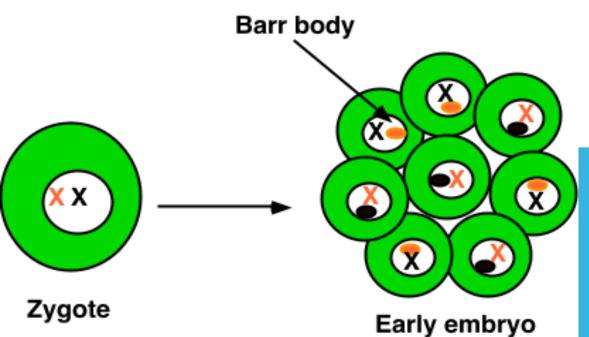
7- The identification of sex chromosome: if a newborn presents with ambiguous genitalia, one can not tell whether the sex is male or female.

The presence of a dark dot attached to the nuclear membrane from inside (Barr body +ve) indicates that a sex chromosome is present, i.e., the genotype of the baby is XX (\mathcal{P}).

Barr Bodies

Barr body





<u>Cytopathology can be further subdivided</u> into:

- Gynecological Cytopathology, include cervicovaginal cytopathology.....etc.
- Non gynecological Cytopathology include cytopathology of all other organs.
- Fine Needle Aspiration (FNA): include FNA of breast, FNA of thyroid.....etc.

Gynecological cytopathology:

Two types of epithelia are present in female genital organs

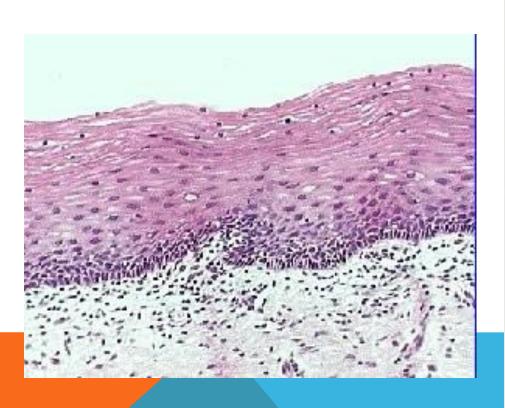
Columnar epithelium: lined uterus & endocervix.

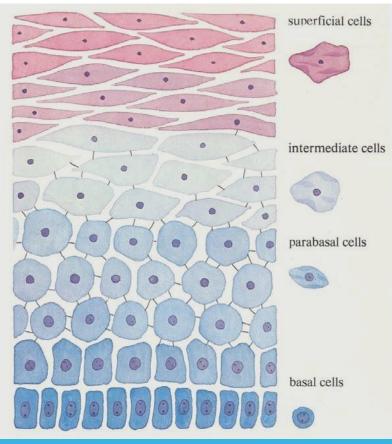
Squamous epithelium: lined ectocervix & vagina.

Squamous Metaplasia of endocervical epithelium (columnar) into Squamous epithelium is usual process occurring in all women, as a result of hormonal effects. (Changes more in cytoplasm of cells more than nuclei).

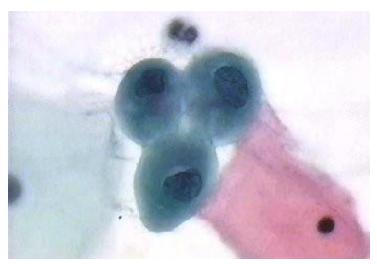
For reporting of cervico- vaginal cytology, Bethesda system (1991) is currently used.

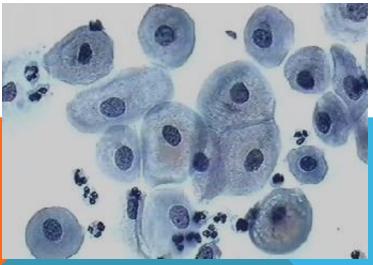
SQUAMOUS EPITHELIUM

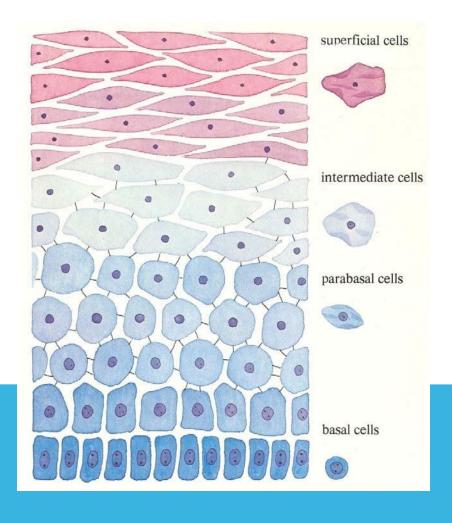




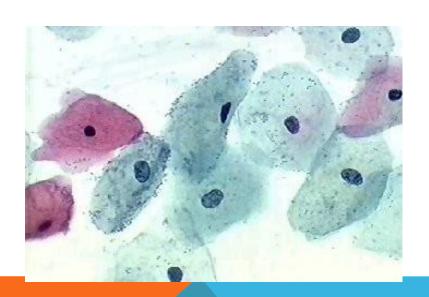
PARABASAL CELLS

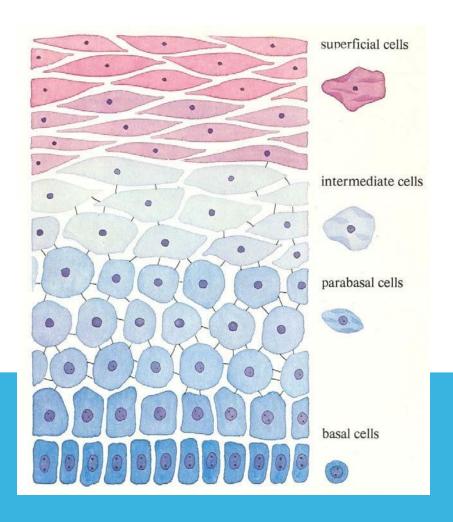




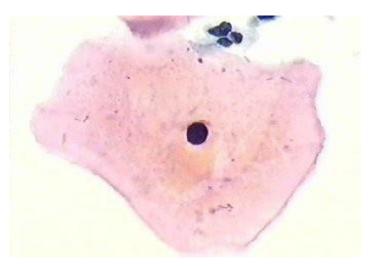


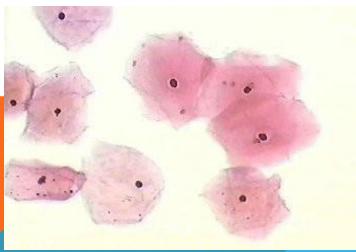
Intermediate CELLS

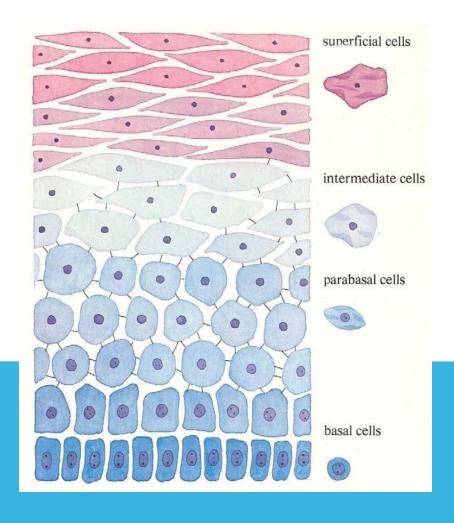




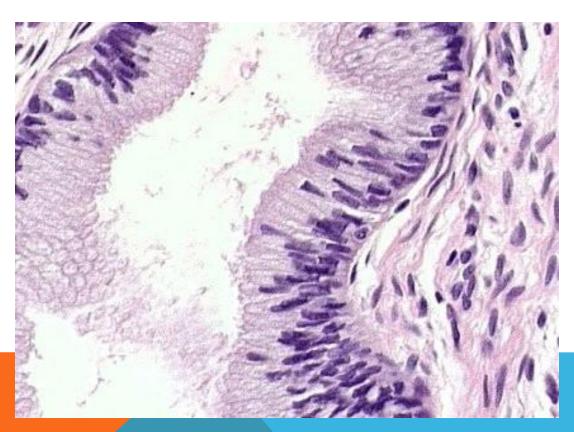
Superficial CELLS

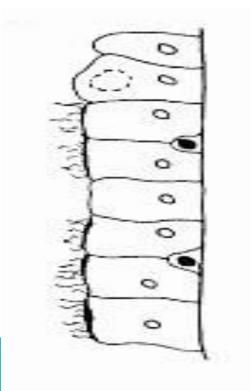






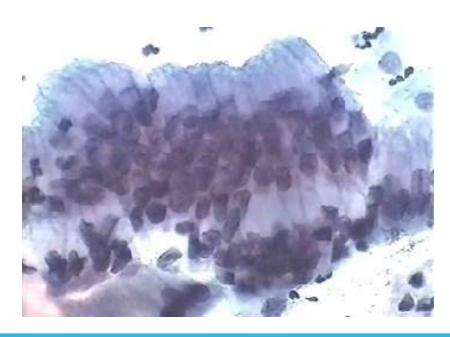
ENDOCERVIX





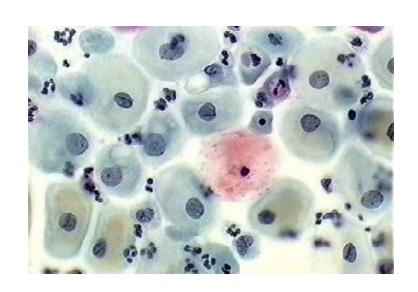
Endocervical CELLS

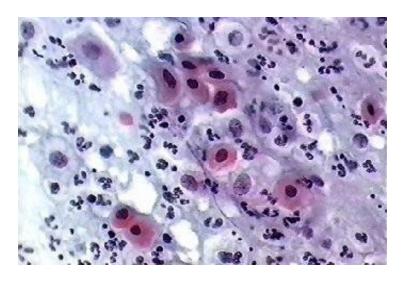


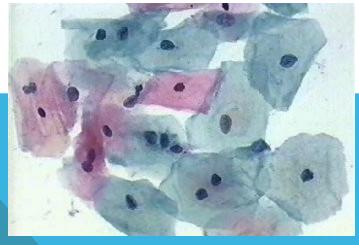


characteristics	Normal Squamous cells	Metaplatic Squamous cells	Malignant Squamous cells
Cell arrangement	Exfoliated singly	Cohesive sheets or groups	Singly or dyscohesive sheets. Variable size & shape of cells.
Cytoplasmic characteristics	Abundant • cytoplasm Well defined cell • borders	Either pale to dense cytoplasm. Cytoplasmic processes. Poor defined cell borders.	Scant cytoplasm
Nuclear characteristics	Centrally located nucleus. Absent nucleoli. Low N/C ratio.	Larger nuclei than normal. With or without nucleoli.	Variable size & shape. High N/c ratio. Prominent nucleoli.

NORMAL SMEAR

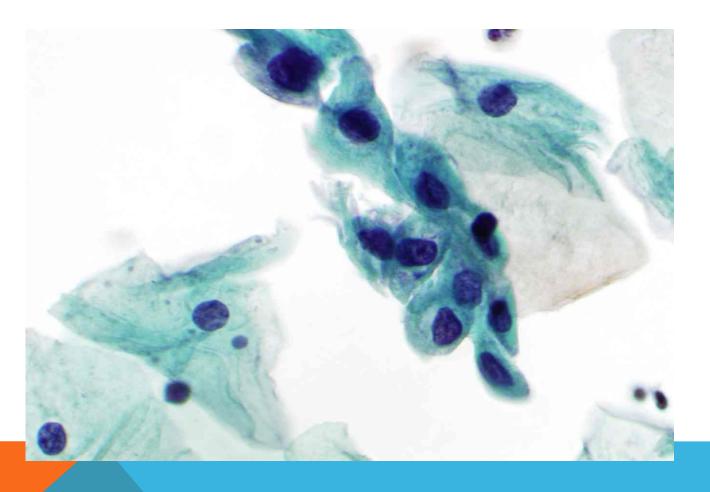




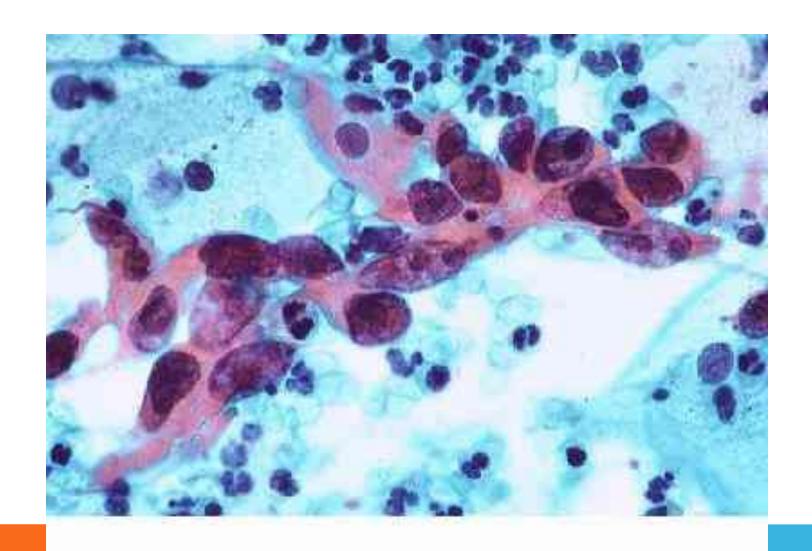




Normal squamous cells



Squamous metaplastic cells



Squamous cell carcinoma

Non-Gynecological cytopathology:

Include cytological examination of all other organs (e.g. respiratory system cytopathology, urine cytopathology.....)

1. Serous effusions:

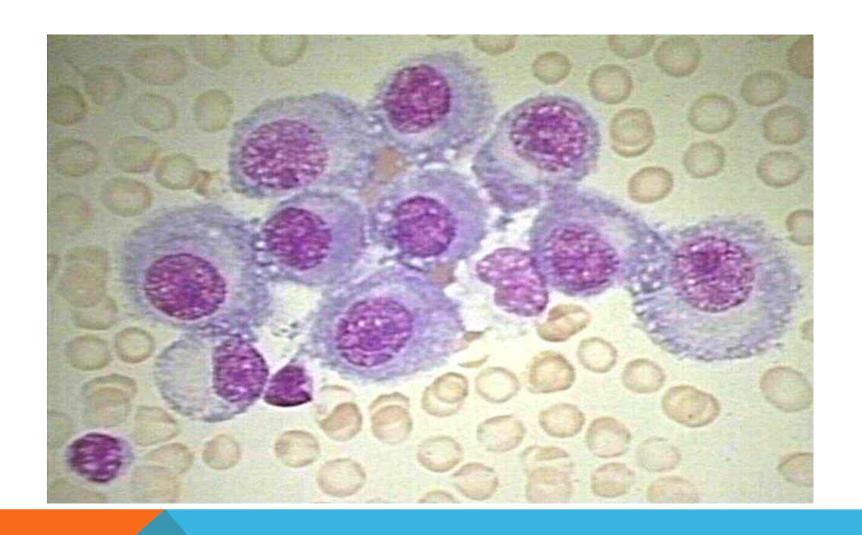
Accumulation of fluid in serous cavities (e.g. pleura, peritoneum)is abnormal & can results from many causes (inflammation, cancers....).

Cytological examination of serous effusions is performed mainly to establish the presence or absence of malignancy (either primary mesothelioma or metastatic carcinoma to these sites).

Metastatic carcinomas to the serous surfaces are from Lung, & ovary breast, colon, stomach,

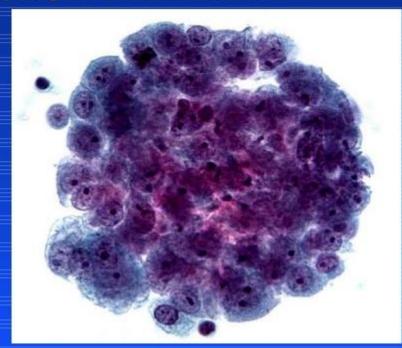
Better to differentiate between mesothelioma & metastatic carcinoma by using special stains & immunohistochemistry.

<u>Characteristic</u>	<u>reactive mesothelial cells</u>	Malignant mesothelioma	<u>Adenocarcinomas</u> (metastatic)
Fluid characteristics	Clear, turbid	Always hemorrhagic	Hemorrhagic
Cells arrangement	Singly mainly, less as tissue fragments. Ill defined cell borders.	Small to large complex sheets of cells. Well defined cell borders.	Acinar (gland like formation). Well defined cell borders.
Nuclear characteristics	Variable in number & shape. Normal N/C ratio Multiple micronucleoli.	Same Increase N/C ratio	Single, macronucleoli, coarse chromatin. Increase N/C ratio
Cytoplasmic characteristics	Moderate to abundant cytoplasm.	Scant to moderate amount.	Scant to moderate amount. Presence of cytoplasmic vacuoles & signet ring cells.



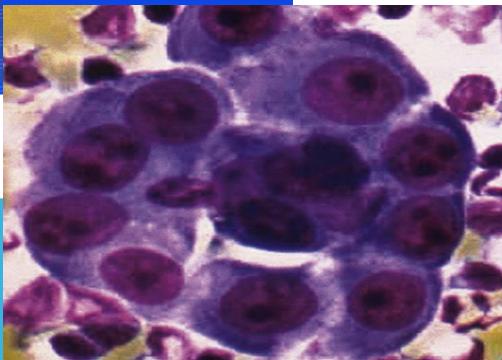
Reactive mesothelial cells

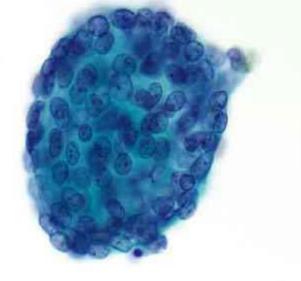
Cytological features of mesothelioma

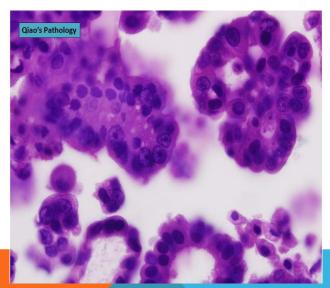


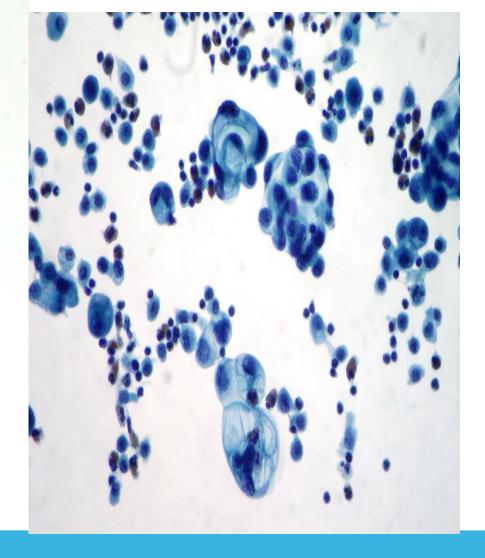
Loosely cohesive threedimensional groups of mesothelioma cells with knobby outline. A multinucleate atypical mesothelial cell is seen at the periphery of the group (arrow).

Thinprep. Pap stain 40x.







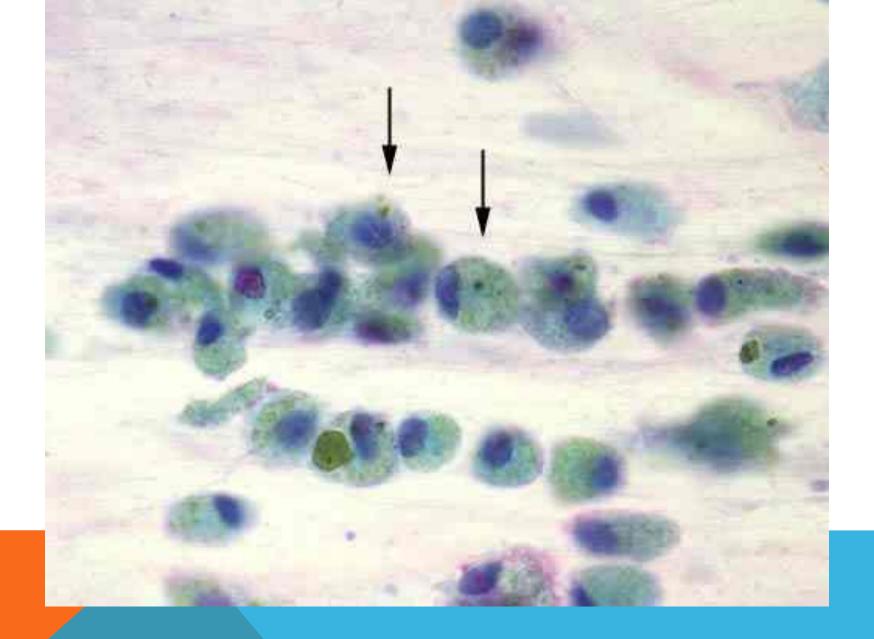


Metastatic malignant cells

2. Sputum Cytology:

- Offers highly diagnostic values for <u>diagnosis of centrally</u> <u>located lung cancer</u> (Squamous, small cell cancers of <u>lung</u>).
- 3 to 5 consecutive daily sputum examinations is advisable to increase the rate of detection of cancers.
- Sputum may spontaneously coughed or induced.
- Sputum sample is <u>fixed with Saccomanno fixative(2% polyethylene glycol With 50% ethyl alcohol).</u>
- Sputum sample is considered satisfactory when it

contains alveolar macrophages.



Alveolar macrophages

FNA Of Breast:

FNA is by far the most popular, simple, cost effective, reliable, & diagnostic procedure in cytological diagnosis of breast lesions.

FNA is important part in <u>triple screening test</u> for early <u>detection of beast carcinoma</u>:

1- (physical examination a thorough physical examination of the whole breast area, including both breasts, nipples, armpits.

2-Imaging tests

Imaging tests for breast cancer may involve:

mammogram: a way of examining the breasts using low-dose X-rays •

ultrasound:

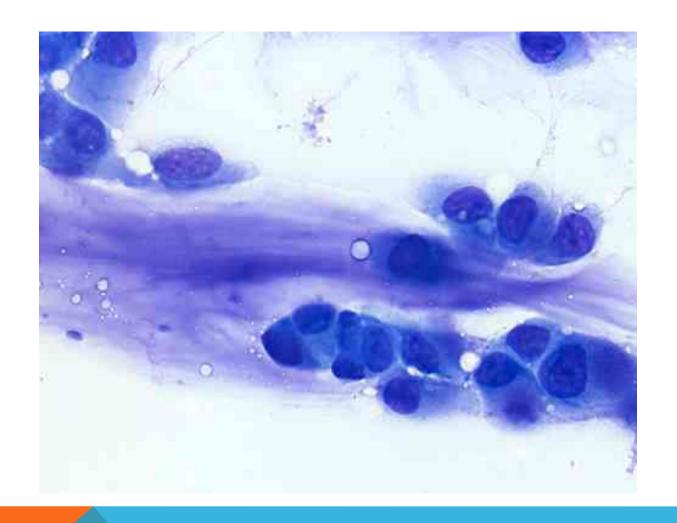
- The tests used will depend on the woman's age. For women aged 35 years or older, both a mammogram and an ultrasound will be used.
- For women younger than 35 years, an ultrasound is used first and a mammogram may also be used if more information is needed.

<u>3- FNA).</u>

<u>Characteristics</u>	Benign pattern	Malignant pattern
Cellularity	Almost low cellularity	High cellularity
Cells arrangements	Sheets of uniform ductal cells (cohesive)	Single, variable size malignant cells. (poorly cohesive)
Single bare nuclei	FREQUENT	ABSENT
Nuclear characteristics	Uniform size	Enlarged, atypical nuclei.



Fibroadenoma of breast



Malignant cells of breast aspirate

