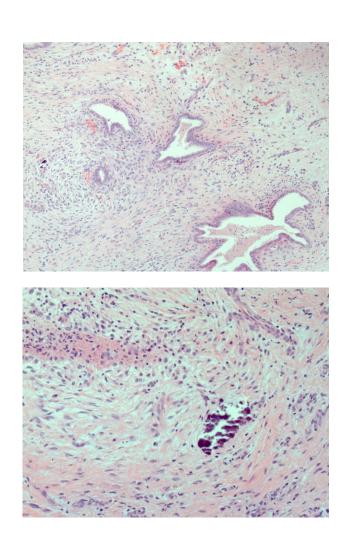
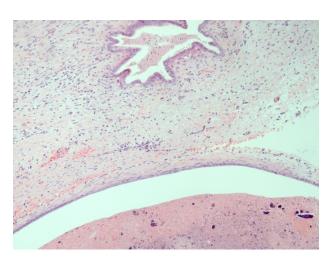
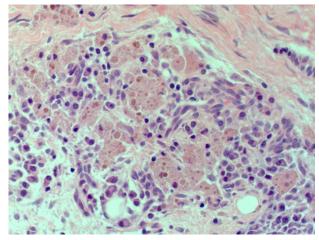
FRCPath Revision Course Gynaecological Pathology

Prof Naveena Singh Consultant Pathologist Barts Health NHS Trust

- 09S36776
- 29/F Excision of fibrous nodule at end of lower abdomen incision.







Endometriosis

- Typically glands+stroma+haemosiderin; 2 out of 3 for diagnosis
- Glands:
 - Inactive or functional
 - Metaplastic changes: ciliated, hobnail, squamous, mucinous
- Stroma:
 - Usually obvious
 - May be inconspicuous cuff
 - Spiral arterioles, haemosiderin, CD10 IHC
 - Decidual change
 - Myxoid change
 - Smooth muscle metaplasia/ elastosis
- Haemosiderin:
 - Pigmented histiocytes
 - Pseudoxanthomatous

Endometriosis – Differential diagnosis

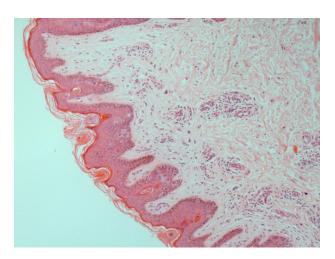
- Endosalpingiosis
- Endometrial stromal sarcoma with glandular differentiation ('aggressive endometriosis')
- Adenosarcoma
- Other causes of pseudoxanthomatous reaction

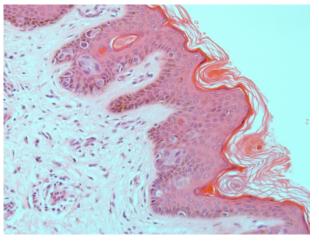
Malignancy in endometriosis

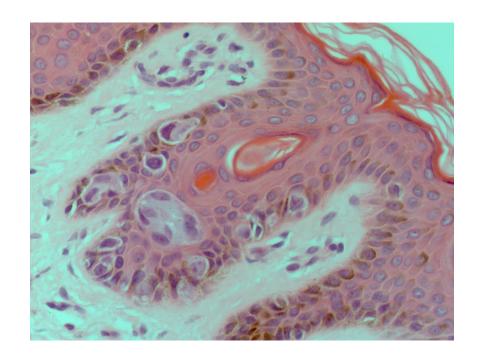
- Up to 10% can have malignancy
- Carcinoma: clear cell or endometrioid
- Other epithelial: Seromucinous borderline tumours; endometrioid adenofibroma
- Mesenchymal: ESS, MMMT, adenosarcoma

Revision: Polypoid tumour-like endometriosis

- 09S14138
- 66/F Previous vulvectomy for extramammary Paget disease







Paget disease of Vulva

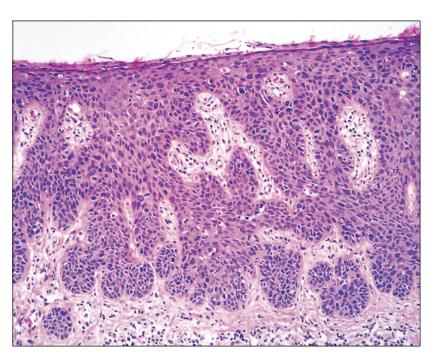
- Malignant cells in epidermis and adnexal structures
- Vesicular nucleus, prominent nucleolus, pale amphophilic cytoplasm which is mucin positive
- IHC: CK7, CEA, Cam 5.2
- 30% have unrelated synchronous/metachronous
 Ca (breast, cervix, UB)
- <5% are secondary to local carcinoma (genital, anal, urinary tract)
- 30% have dermal invasion facilitated by CK7

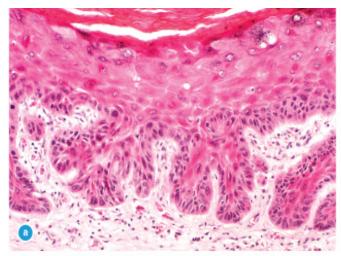
Paget disease of Vulva – Differential diagnosis

- Pagetoid VIN
- Malignant melanoma superficial spreading

Revision: Pathways of VIN and VSCC

VIN





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Classical VIN 3

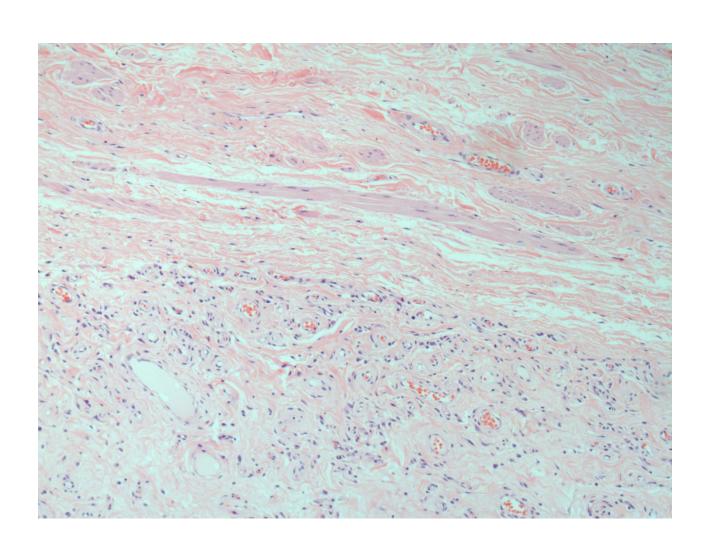
VIN simplex

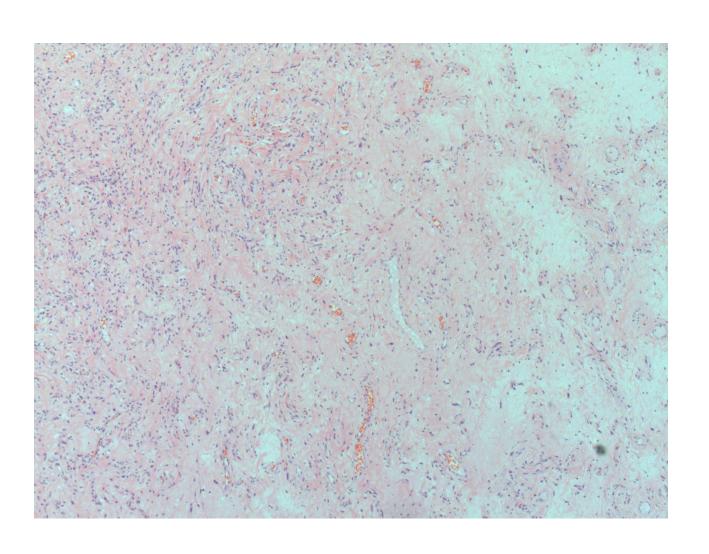
	HPV	Non-HPV	
Age	4 th – 6 th decade	6 th – 9th decade	
Etiology	Oncogenic HPV infection	Chronic inflammation (lichen sclerosus)	
Precursor	High grade VIN, usual type	Differentiated VIN	
Biomarker expression	p16 overexpression	Abnormal p53 expression; p16 negative	
Outcome	Favorable	Less favorable (more likely to have nodal mets, recur	

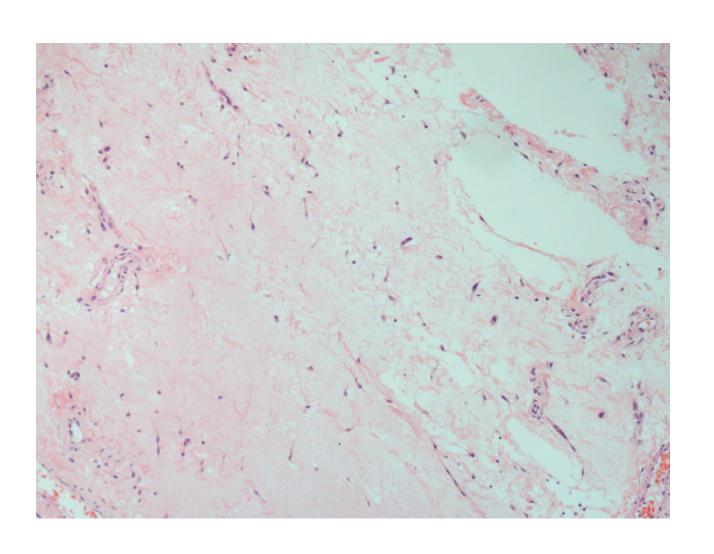
Clinical Significance of HPV Status in VSCC

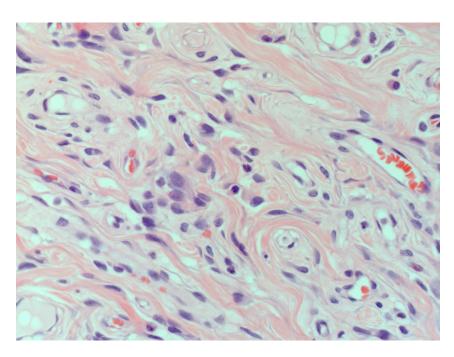
- HPV-related VSCC has better outcomes
 - Younger
 - Less advanced
 - Lower incidence of LN mets
 - Better outcomes with surgery, even when conservative
 - Better outcomes with RT/chemoRT
- Essential to classify VIN/VSCC as HPVmediated or HPV-independent

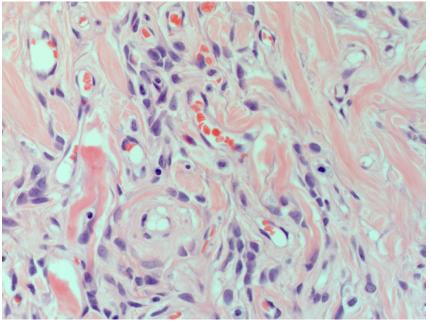
- 07S8184
- 30/F Excision of vulval cyst











Angiomyofibroblastoma

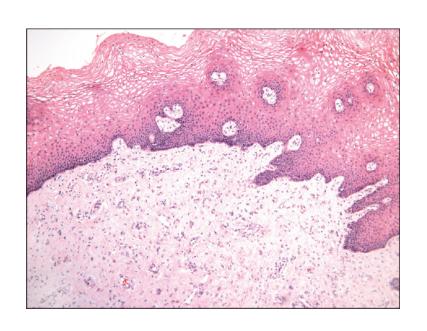
- Well circumscribed
- Alternating hyper- and hypocellular zones
- Numerous small to medium-sized blood vessels
- Perivascular fibrosis
- Spindled, oval, epithelioid or plasmacytoid cells
- Collagen strands or bundles
- IHC: Vimentin, desmin+, occasionally actin, CD34

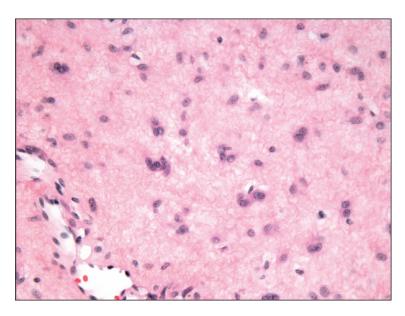
Angiomyofibroblastoma – Differential diagnosis

- Aggressive angiomyxoma
- Cellular angiofibroma

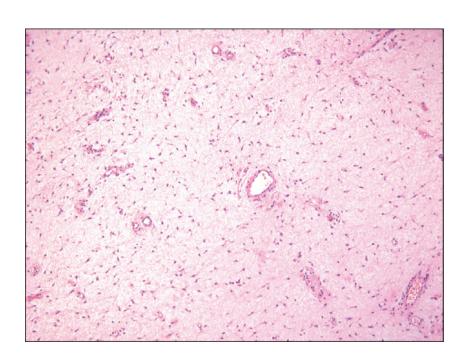
Revision: Site-specific vulvar mesenchymal lesions and their differentiating features

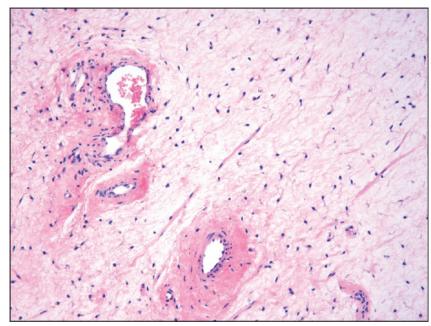
Fibroepithelial stromal polyp





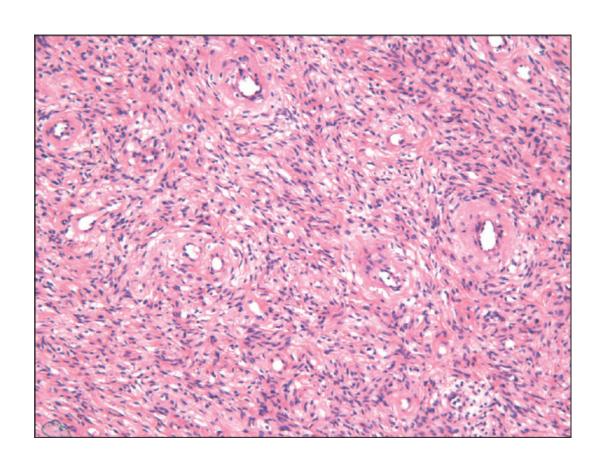
Aggressive/deep angiomyxoma



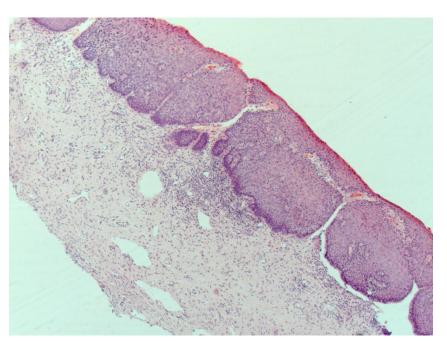


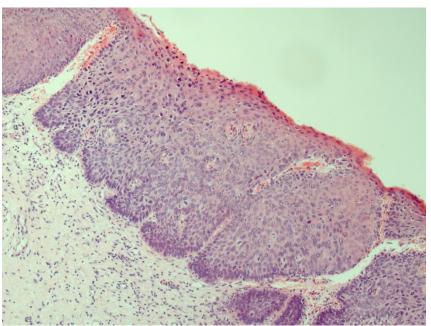
New IHC marker: HMGA2

Cellular angiofibroma



- 09S28705
- 34/F Vaginal biopsy, retrovirus positive





VAIN 3

 Only 1% as common as CIN 3, may be associated with invasive SCC

Revision: Vulvovaginal cysts

Also: Prostatic type lesions in vagina

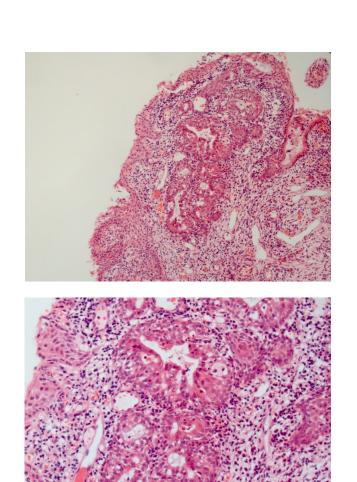
Remember: Prolapsed FT; tubulosquamous polyp

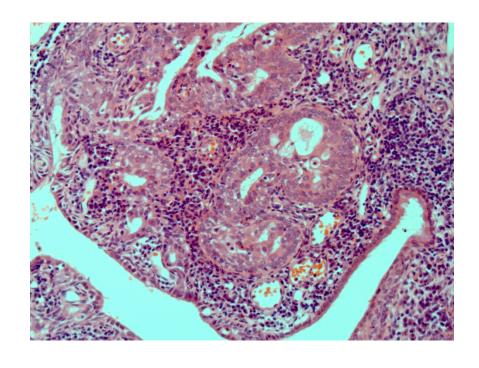
Look up: LAST project: unified 2-tier terminology for anogenital HPV related squamous lesions (with -IN1, 2 and 3)

Vulvovaginal cysts

Lining		Site	Origin
Mucinous	Mullerian cyst	Upper vagina, usually anterolateral wall	Mullerian duct
	Mucous vestibular	Usu vestibule, other	Minor vestibular glands
	Bartholin cyst	Posterolat vagina	Bartholin gland
Non-mucinous, non-ciliated cuboidal	Gartner's cyst	Lateral wall of vagina	Wolffian remnants
Columnar endometrial type/ciliated	Mullerian cyst	Upper vagina, usually anterolateral wall	Mullerian duct
	Endometriotic cyst	Vulvovaginal endometriosis	Site of trauma/ vault or posterior fornix
Transitional	Bartholin's cyst		Posterolat vagina
	Skene's cyst	Close to urethral meatus	Skene's duct

- 10S326
- 35/F High grade dyskaryosis; recent pregnancy. Cervix looks suspicious with contact bleeding





Microglandular hyperplasia with cytological atypia

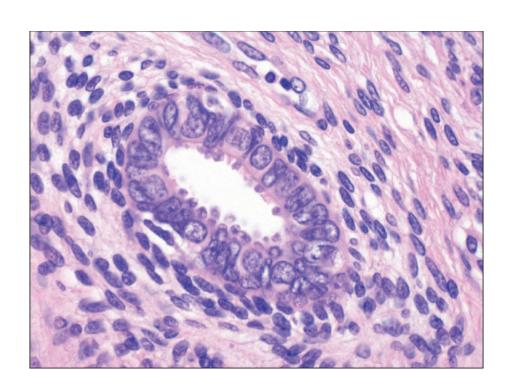
- Closely packed glands with variable shape, cribriform architecture
- Lumen contains inspissated mucin and inflammatory cells
- Stromal inflammatory infiltrate
- Glandular cells low columnar or cuboidal, subnuclear vacuolation present
- Small regular nuclei, inconspicuous nucleoli
- Focal squamous metaplasia
- Mild to moderate atypia may occur but is unusual

Microglandular hyperplasia with cytological atypia – Differential diagnosis

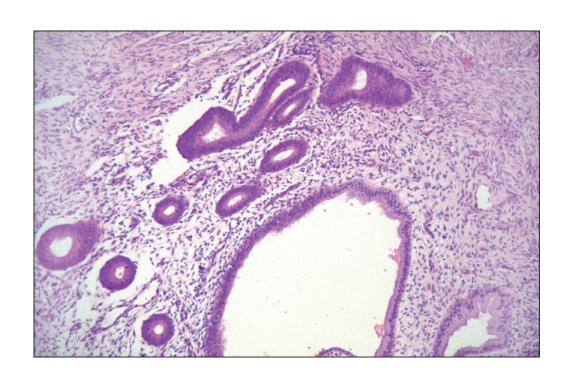
- Endometrium with secretory change
- Endometrial carcinoma: endometrioid or clear cell
- Cervical adenocarcinoma or adenosquamous carcinoma

Revision: benign mimics of cervical adenocarcinoma; role of p16 in diagnosis of cervical neoplasia

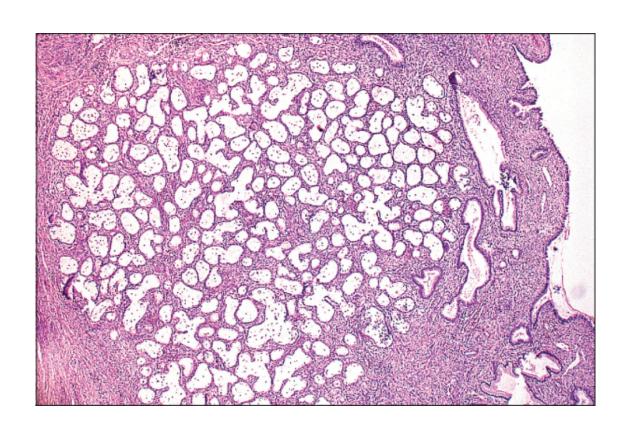
Tuboendometrioid metaplasia



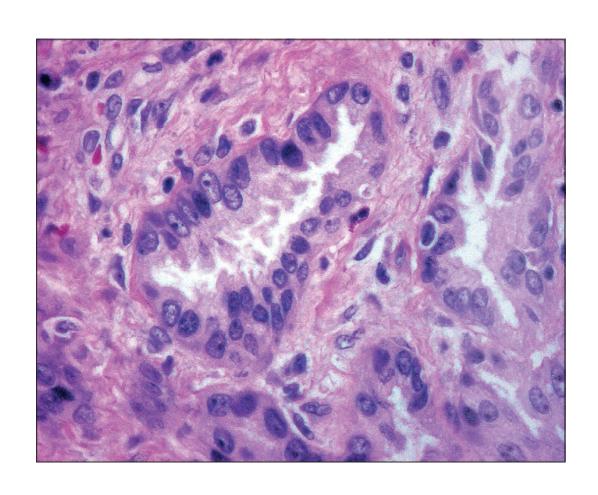
Endometriosis



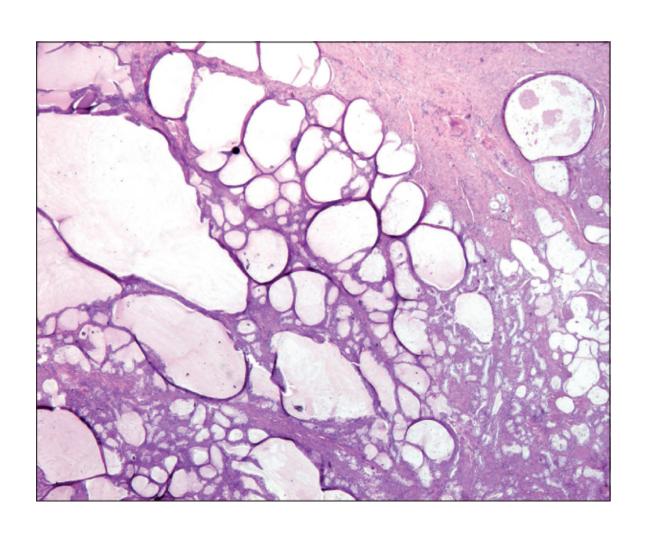
Tunnel clusters type B



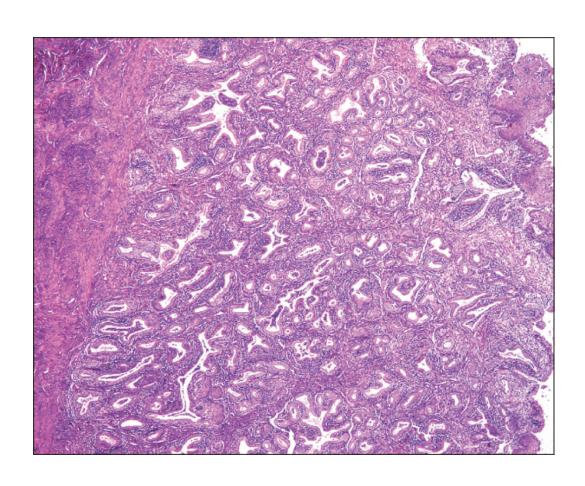
Tunnel clusters type A



Lobular endocervical glandular hyperplasia

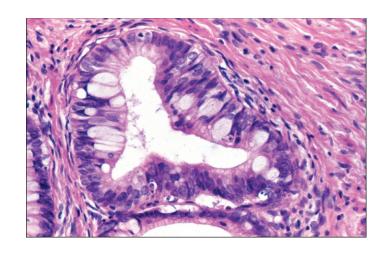


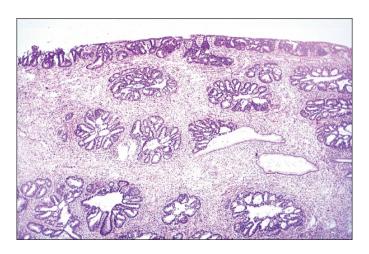
Diffuse laminar endocervical glandular hyperplasia

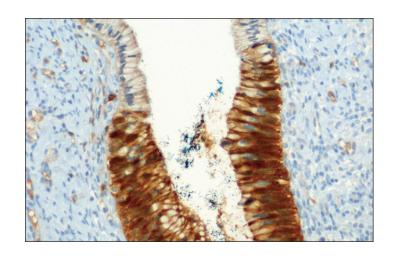


CGIN

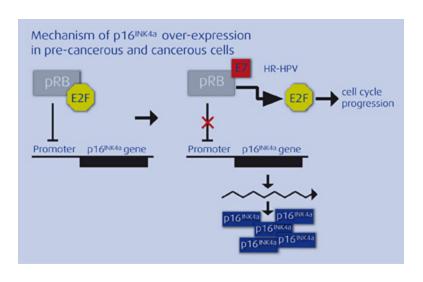








p16ink4a



- hrHPV E7 inactivation of pRB results in p16 overexpression by disturbing E2F dependent feedback loop
- p16 is not expressed in ALL hrHPV infected cells
- Some normal cells also express p16
- Diffuse and strong p16
 overexpression is a sensitive
 and specific marker for a
 transforming hrHPV infection
- p16 is not directly related to HPV and has diagnostic value in other settings

What does p16 expression signify in HPV neoplasia?

 Diagnostic marker of transforming HRHPV infection (necessary but not sufficient for progression to cancer)

What does p16 **NOT** signify?

Not present in ALL HRHPV infection

- SPECIFIC for hrHPV but not seen in ALL cases
- HRHPV spectrum: silent-productive-transforming/abortive
- Not seen in IrHPV

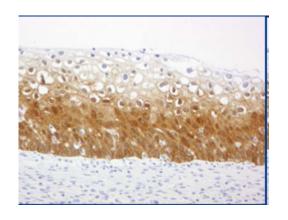
Not diagnostic of high grade CIN

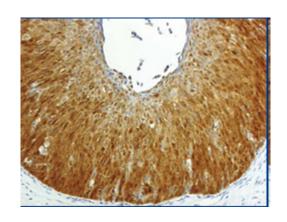
About 50% of CIN1 is also p16 positive

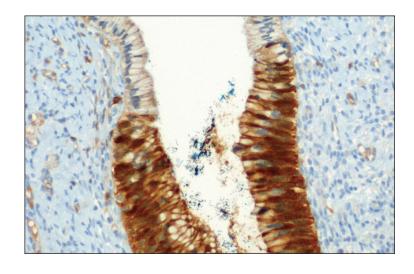
Does NOT indicate likelihood of progression

- Many conflicting papers but overall p16 (in CIN1) does NOT have predictive value for progression
- >90% CIN1 and up to 2/3rds CIN2 in young women will spontaneously regress after biopsy
- (CIN1 overcalled; CIN2 poorly reproducible)

p16





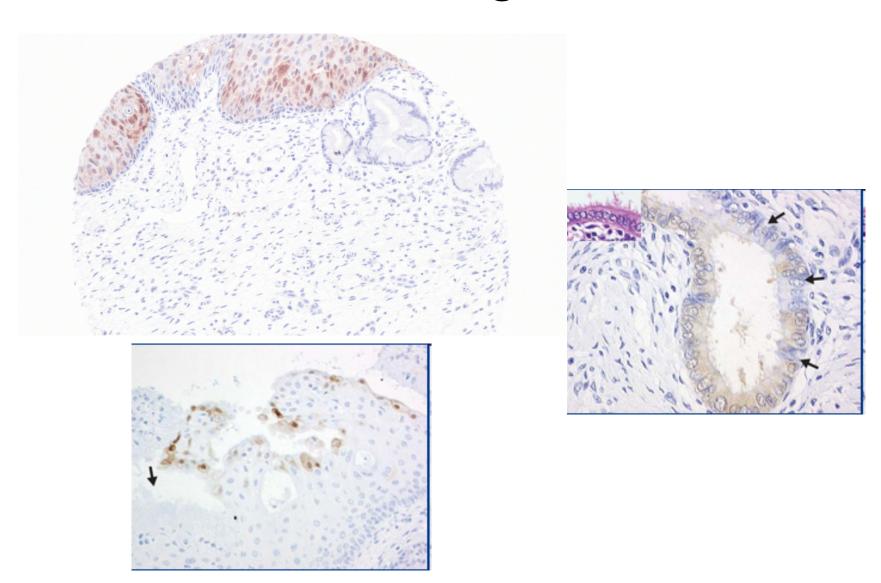


It is the pattern not the presence of positive staining that is specific:

Squamous: Diffuse basal nuclear or nuc+cytoplasmic for at least lower onethird epithelium ('block-positive')

Glandular: Diffuse, strong nuclear or nuc+cytoplasmic

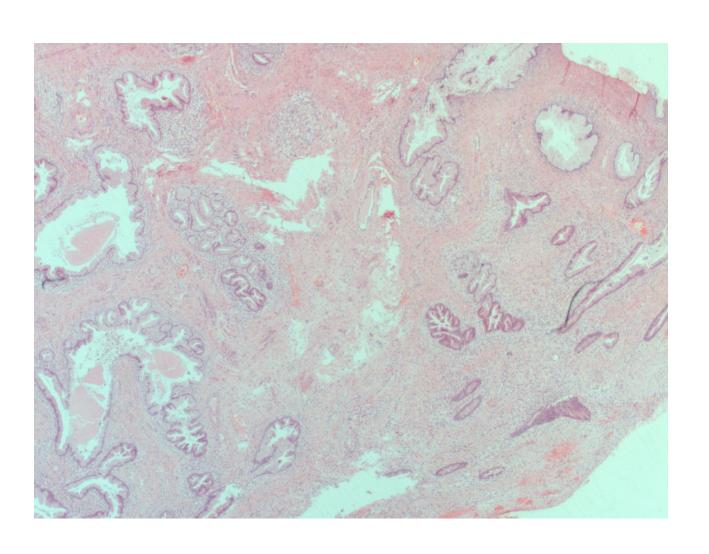
P16 in benign mimics

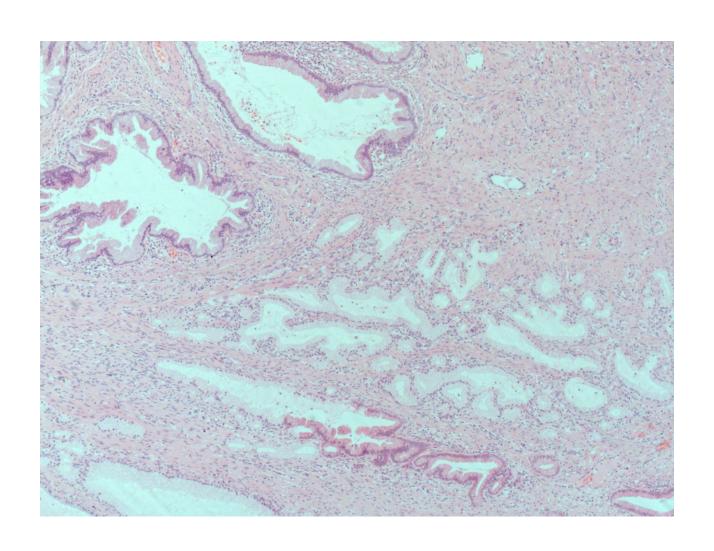


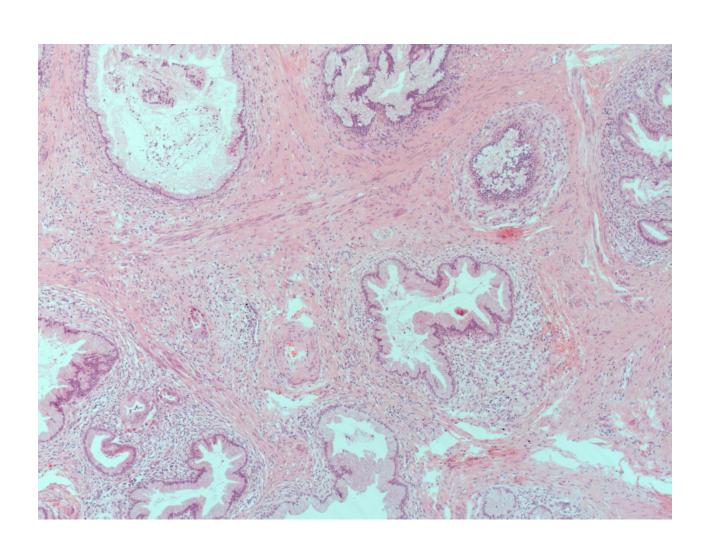
Recommendations for use of p16 (LAST project)

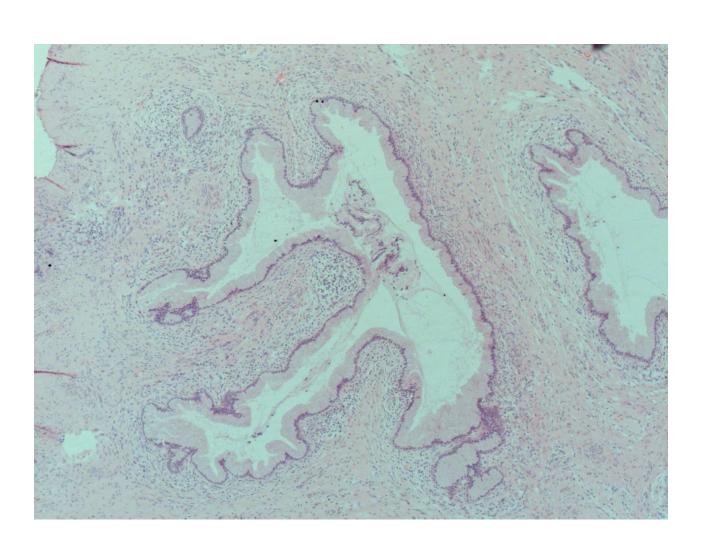
- DD between HG CIN and a mimic: strong and diffuse block-positive p16 → HGCIN
- To help clarify a diagnosis of CIN 2: negative or non-block-positive staining → LGCIN or non-HPV associated pathology
- p16 should <u>not</u> be used if H&E DD is between LGCIN and negative, as CIN 1 can be either p16 negative or positive: p16 IHC is not recommended if diagnosis is "obvious" CIN 1.

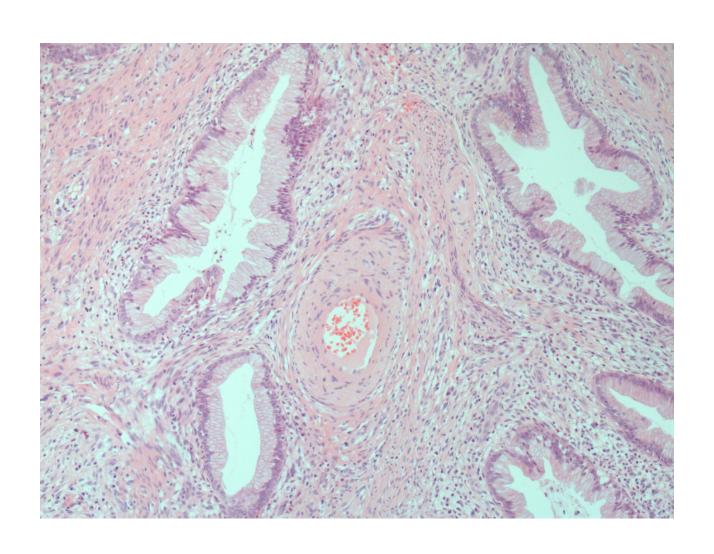
- 09S39415
- 42/F Glandular abnormality. On colposcopy - Unusual appearance with fibroid on posterior lip

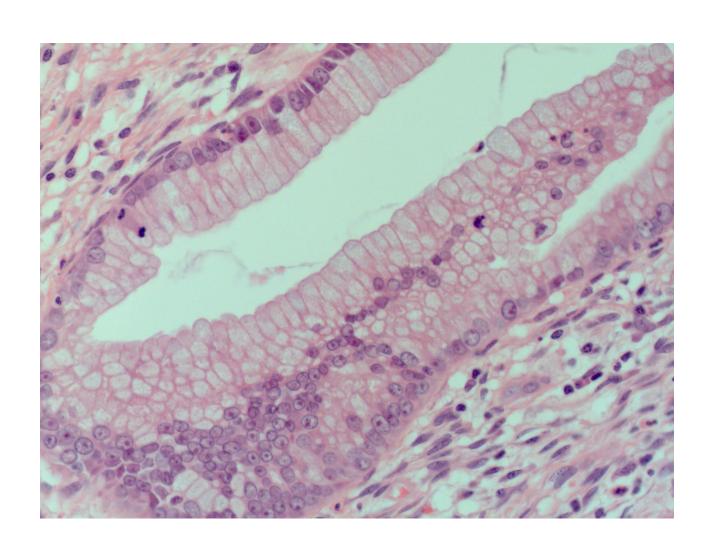


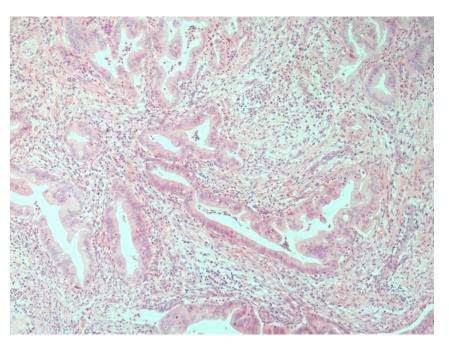


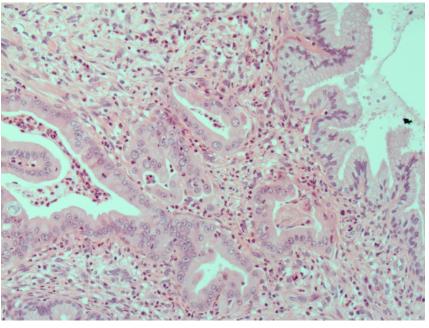


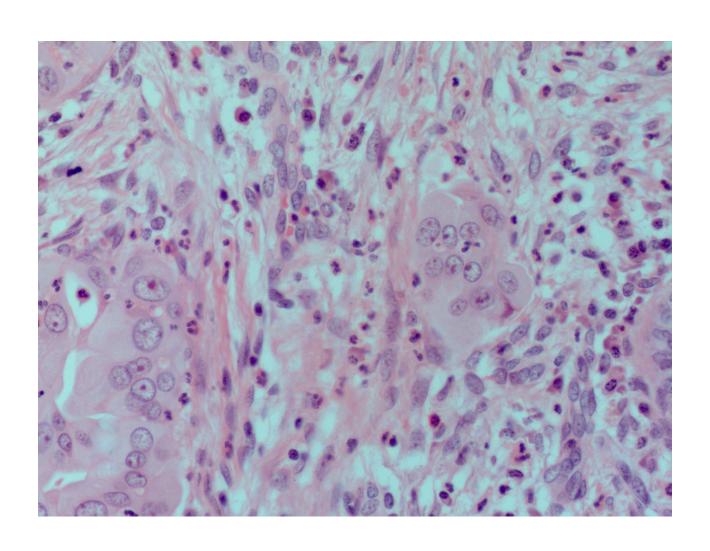


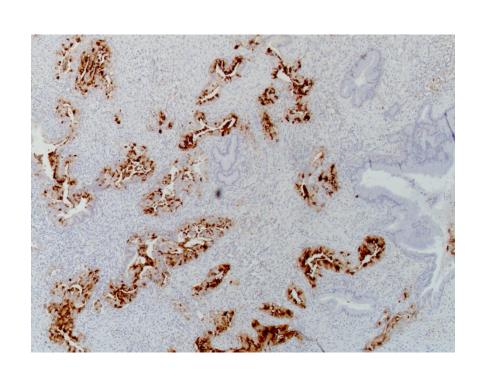


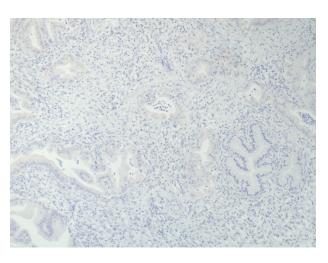


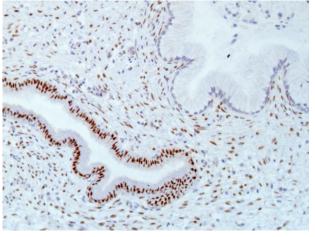












Minimal deviation adenocarcinoma, mucinous type

- 1% of cervical adenoca; HPV negative
- Neoplastic glands lined by deceptively benign appearing epithelium
- Mucinous epithelium, basal nuclei, occasional mitotic figures
- Irregular outlines, variable, claw-like contours
- Desmoplastic stroma seen focally
- Glands adjacent to thick-walled vessels
- At least focal dysplasia present or frank invasion
- CEA positive, ER/PR/CA125 negative

Minimal deviation adenocarcinoma, mucinous type— Differential diagnosis

 Deep glands, cysts, tunnel clusters, diffuse laminar hyperplasia, adenomyoma of endocervical type

Revision: rare subtypes of cervical cancer, including gastric type carcinomas; hereditary gyn neoplasms

Endocervical Glandular Lesions Showing Gastric Differentiation

Category	Diagnostic term		
Benign	Lobular endocervical glandular hyperplasia (LEGH) / pyloric gland metaplasia (PGM)		
	Simple gastric / pyloric gland metaplasia		
	Tunnel cluster (type A)		
Premalignant	Atypical lobular endocervical glandular hyperplasia (atypical LEGH)		
	Adenocarcinoma in situ (gastric type)		
Malignant	Gastric type adenocarcinoma (GAS)		
	Minimal deviation adenocarcinoma (MDA) (AKA: adenoma malignum)		
Specific clinical or clinicopathologic conditions	Synchronous mucinous metaplasia and neoplasia of the female genital tract (SMMN-FGT)		
	Peutz-Jeghers syndrome		

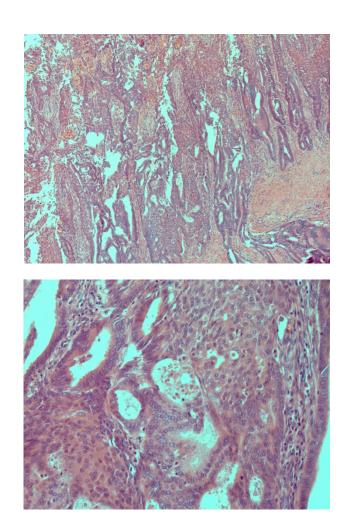
		RCA and Lync	` h)
Syndrome	Gene	Gynaecological tumours	Associated tumours
Peutz-Jeghers Syndrome	STK11/LKB 1	Ovary: SCST (5-15% risk: SCTAT, Sertoli cell tumours) Cervix: Adenoma malignum (10% risk)	Hamartomatous GI polyps Breast, GI, lung, pancrease, testis cancers
Hereditary leiomyomatosis and renal cell carcinoma (HLRCC)	Fumarase hydratase	Uterus: Leiomyomas (prominent nucleoli, perinuclear halos, young patients (most patients)	Renal cell carcinoma (15% risk Cutaneous leiomyomas
Gorlin Syndrome	PTCH	Ovary: Fibromas, bilateral	Basal cell carcinomas

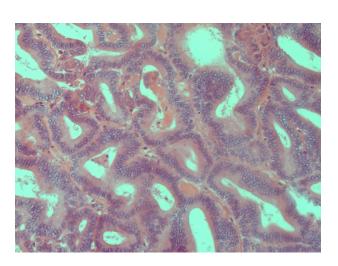
■ (nevoid basal cell and calcified (2-25% risk) Odontogenic keratocysts Syndrome) Medulloblastomas

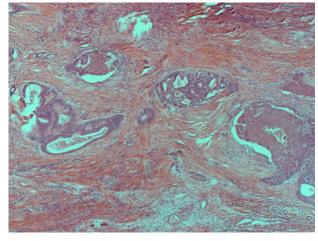
Syndrome)

Uterus: Leiomyomas Cowden PTEN Hamartomas of GI tract, and Syndrome (PTEN multiple other sites Endometrial carcinoma Breast (25-50% risk) and hamartoma (5-19% risk) Thyroid(3-10% risk) carcinomas tumour

- 10S1197
- 62/F Total hysterectomy for PMB







Grade 1 endometrioid adenocarcinoma, infiltrating inner myometrium

- Confluent proliferating glands with mild to moderate nuclear atypia
- Glandular architecture
- Squamous metaplasia

Grade 1 endometrioid adenocarcinoma

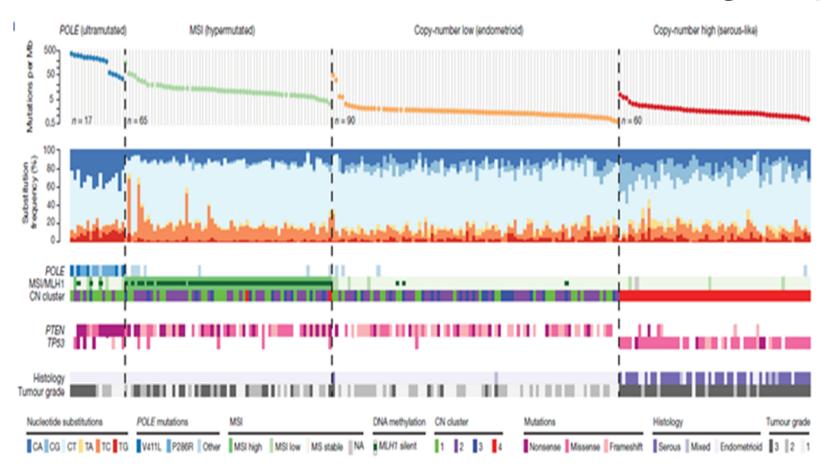
Revision: Molecular classification of endometrial carcinoma

Uterine neoplasia: Endometrial carcinoma

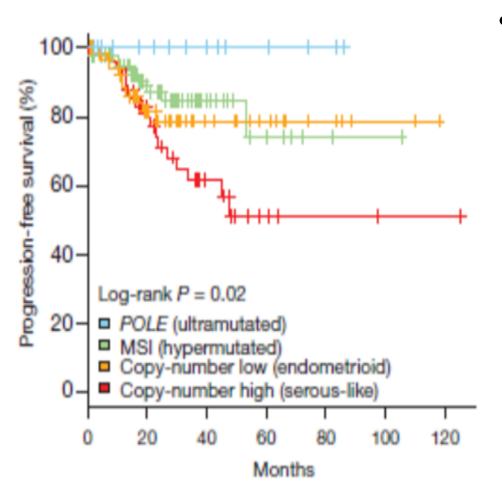
 Type 1: Endometrioid, oestrogen driven, favourable prognosis

 Type 2: Serous, ?clear cell, carcinosarcoma, oestrogen-independent, aggressive clinical course

Endometrial cancer: Molecular groups

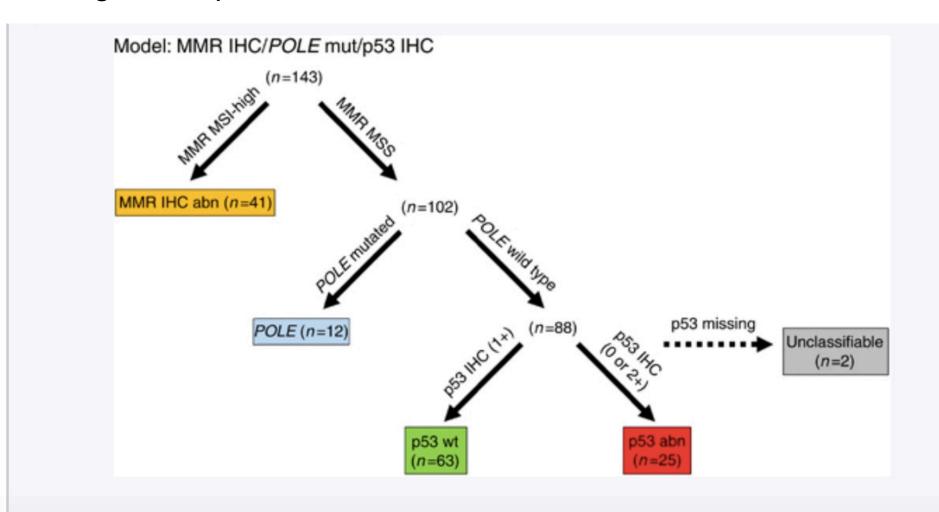


Endometrial cancer: Molecular groups



- 4 distinct groups:
 - POLE
 - MSI
 - Copy-number low (endometrioid)
 - Copy-number high (serous-like)

Endometrial carcinoma: Molecular classification strategies being developed



LS and endometrial cancer

- >95% LS undiagnosed
- Only 2-5% EC is caused by LS; but this gives opportunity to screen for LS
- Women with LS have comparable risk of EC and CRC
- LS EC patients are at substantial risk for second of more malignancies
- Screening/preventative strategies reduce incidence and mortality

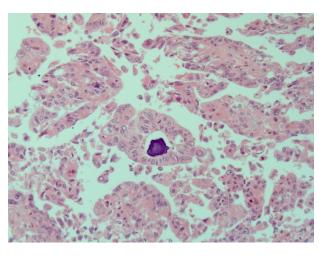
LS and endometrial cancer

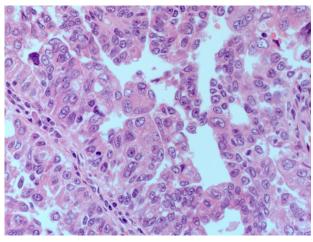
Look up:

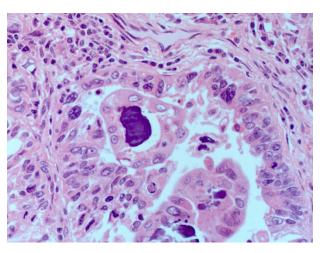
LS IHC

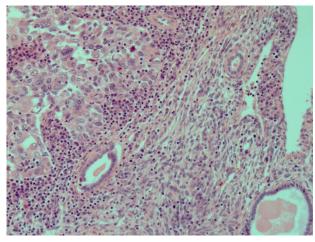
LS related morphology in EC

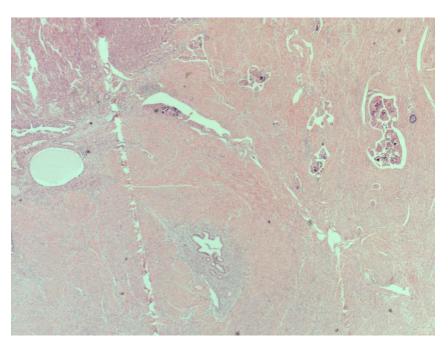
- 10S177
- 73/F Total hysterectomy for PMB; biopsy showed high grade carcinoma

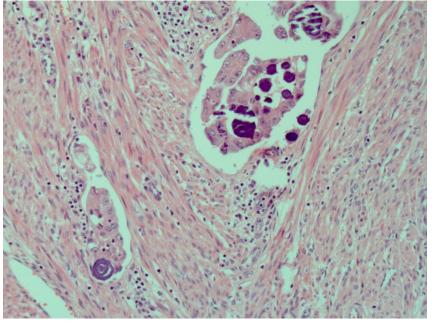












Uterine serous carcinoma

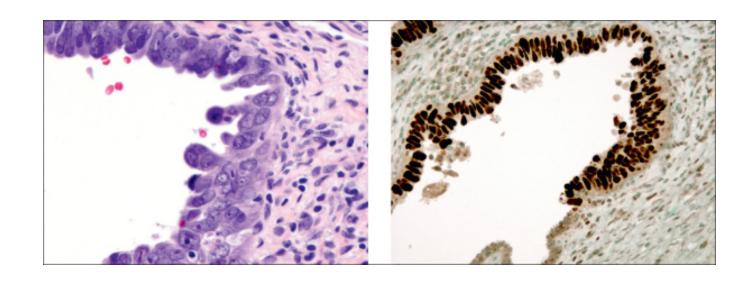
- Marked nuclear atypia out of proportion to architecture
- High mitotic activity
- Bizarre nuclei, vesicular nuclei, prominent nucleoli
- Papillary architecture +/-; tubuloglandular or slitlike common, 'hobnail' luminal outline
- LVSI common
- Extrauterine spread even if minimally invasive
- IHC: ER/PR negative; p53 abnormal/p16 positive

Uterine serous carcinoma – Differential diagnosis

- Endometrioid carcinoma with papillary architecture
- Clear cell carcinoma
- MMMT
- Metastatic serous ca from FT or ovary (WT1)
- Papillary syncytial metaplasia (p53 and Ki67)

Revision: as for endometrioid carcinoma

Serous endometrial intraepithelial carcinoma (EIC)



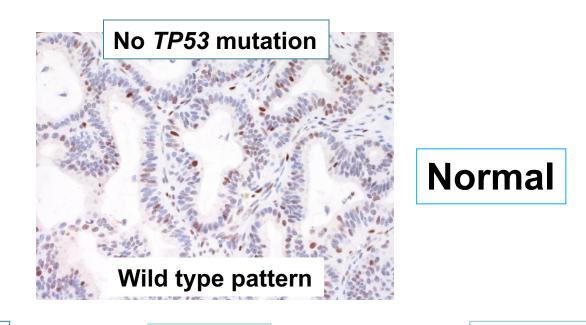
p53 IHC

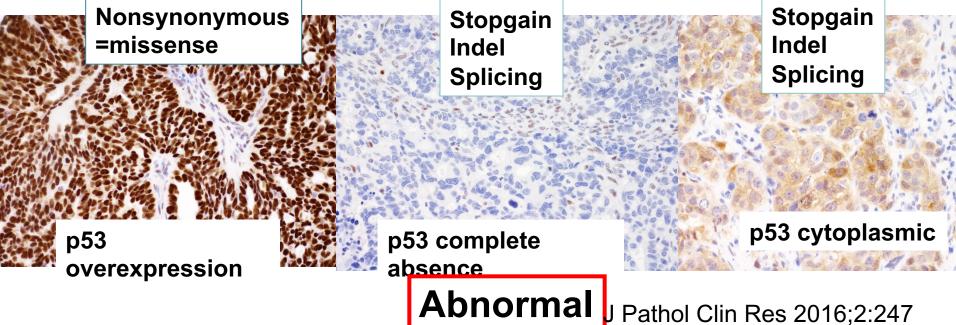
ABERRANT/ABNORMAL

- •POSITIVE: diffuse strong nuclear immunoreactivity (75% cells suggested) (missense mutation)
- NULL (nonsense mutation or deletion)

NORMAL = focal, weak, heterogenous staining ("wild-type" staining)

Interpretation of p53 immunohistochemistry





p53 immunohistochemistry pattern and interpretation

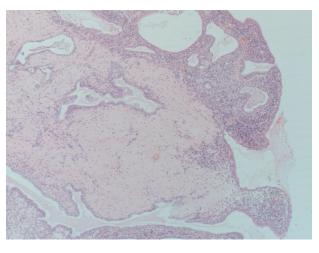
Pattern	p53 IHC Interpretat	TP53 mutation type	% in HGSC
TP53 MUTATION ABSENT			
Wild type	Normal	No mutation	0
TP53 MUTATION PRESENT			
Overexpression	Abnormal	Non-synonymous (missense); also inframe deletion, splicing	66%
Complete absence/null	Abnormal	Indels, stopgains, splicing mutations	25%
Cytoplasmic	Abnormal	Indels and stopgains with disruption of the nuclear localization domain	4%
Wild type	Normal*	Truncating mutation	5%

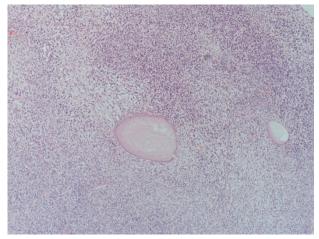
HGSC- high-grade serous carcinoma

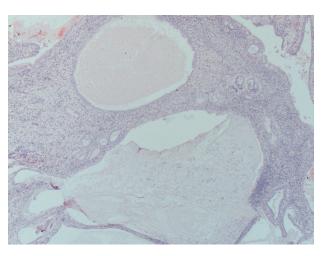
EIN

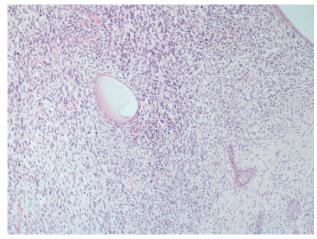
- Not used in the UK
- Refers to endometrioid carcinogenesis
- Identification of endometrioid precancerous change with higher prediction than hyperplasia classification
- Cytoarchitectural demarcation and/or morphometric definition, rather than nuclear atypia
- WHO (2014) equated EIN to atypical hyperplasia ('endometrioid intraepithelial neoplasia')

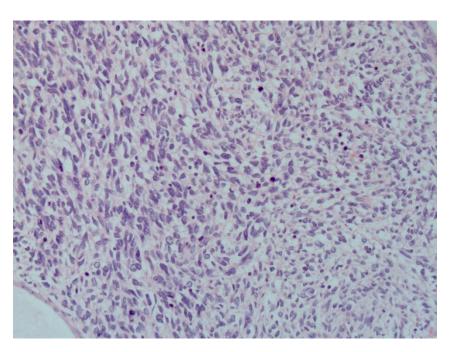
- 09S21401
- 63/F Uterine mass

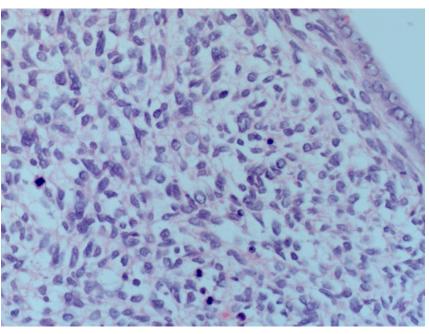


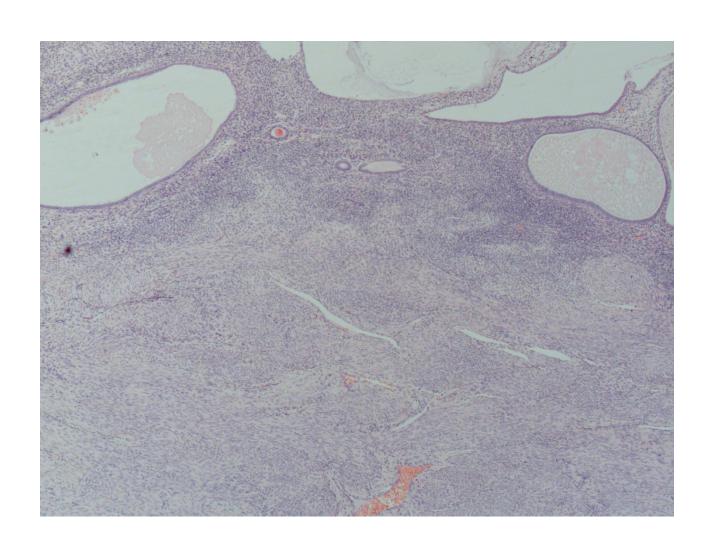












Mullerian Adenosarcoma

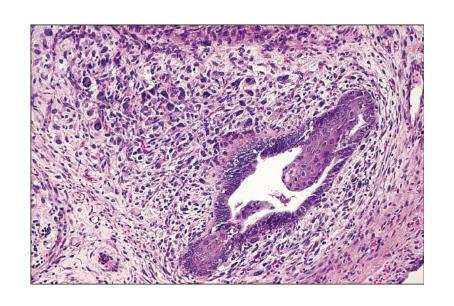
- Biphasic
- Glands should be scattered throughout the mesenchymal component
- Papillary (phyllodes-like) architecture
- Gland lining variable with at most focal and mils cytoarchitectural atypia
- Mesenchymal component consists of low grade sarcoma resembling ESS or fibrosarcoma
- Periglandular cuffing
- Mitosis >4/10 hpf (CAREFUL)
- Heterologous elements +/-
- Low malignant potential

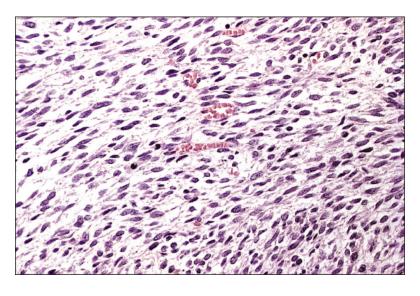
Mullerian Adenosarcoma – Differential diagnosis

- Mullerian adenosarcoma with high grade (heterologous) stroma
- Mullerian adenosarcoma with sarcomatous overgrowth (10% of all)
- Adenofibroma
- Benign endometrial (or endocervical) polyp
- Atypical polypoid adenomyoma
- MMMT
- ESS

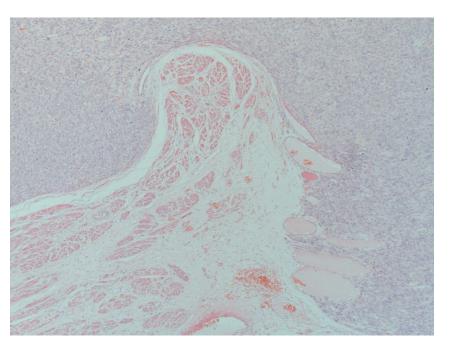
Revision: FIGO staging of uterine sarcoma (2009)

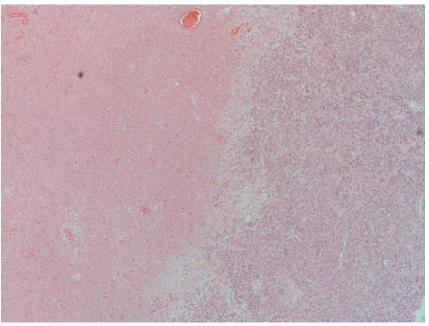
Adenosarcoma with sarcomatous overgrowth

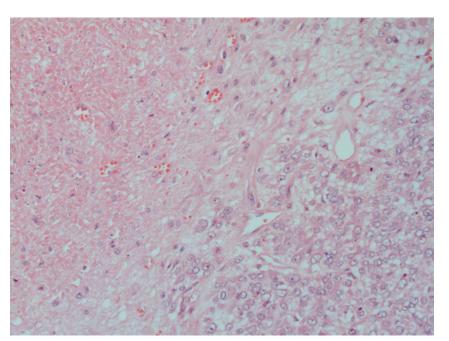


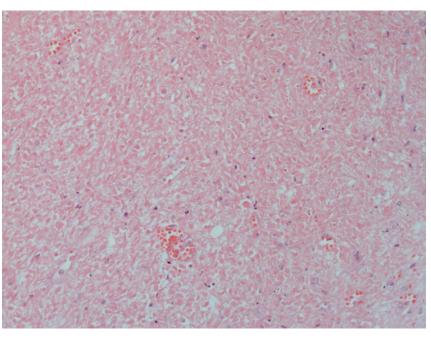


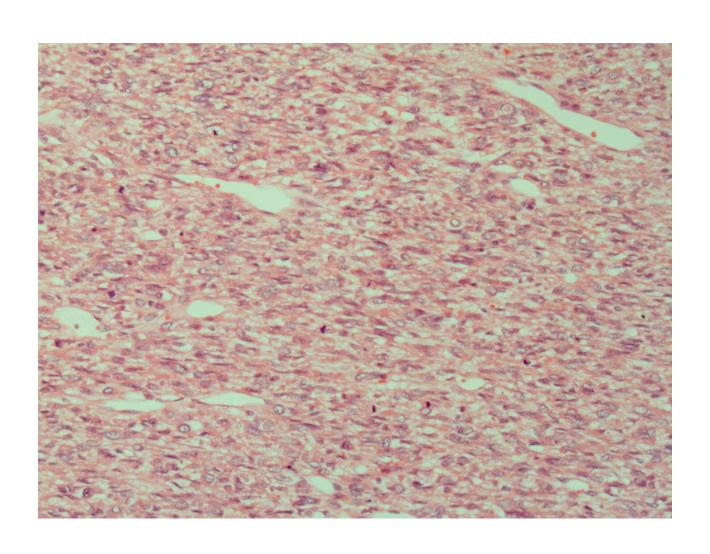
- 09S26072
- 63/F Uterine mass

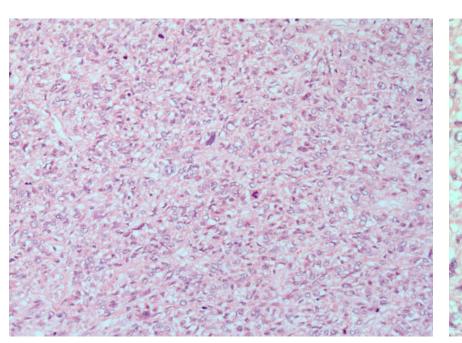


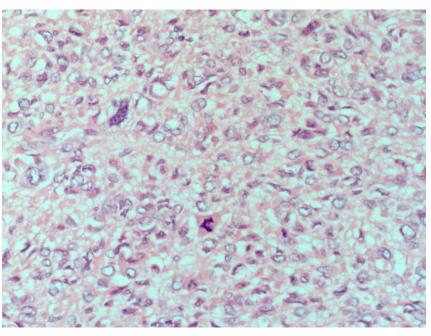












Uterine leiomyosarcoma

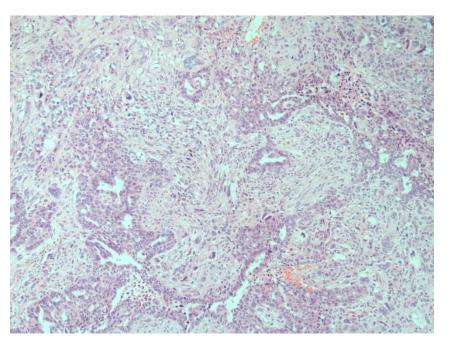
- Hypercellular with 2 of 3 criteria present:
- Diffuse moderate to marked atypia
- High mitotic rate >/= 10/10 hpf
- Tumour cell necrosis (abrupt transition, ghost-like outlines of atypical cells, hyperchromatic tumour nuclei, perivascular survival pattern)
- Other features: atypical mitotic figures, infiltrative edge, vascular invasion

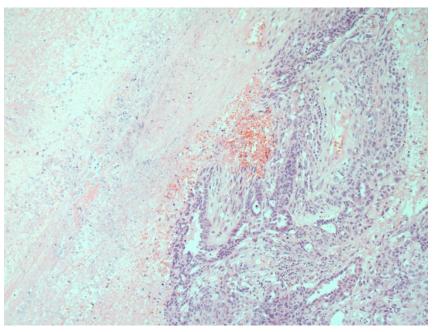
Uterine leiomyosarcoma – Differential diagnosis

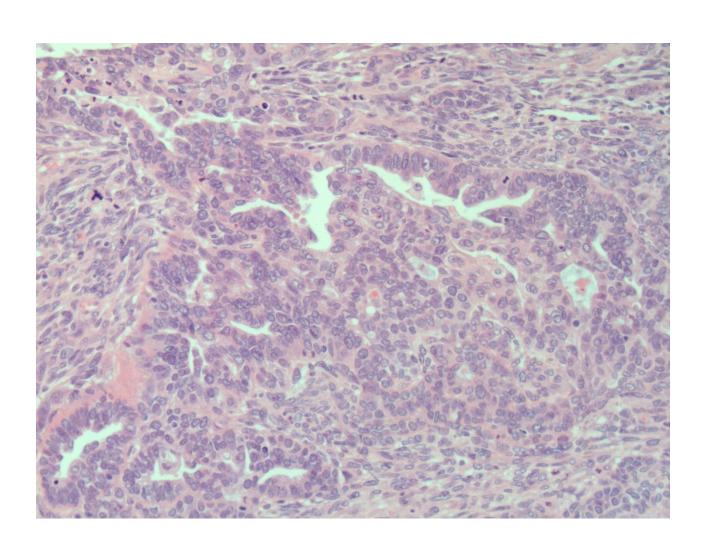
- Leiomyoma variants: cellular, atypical/symplastic, mitotically active, myxoid, intravascular
- ESS
- PEComa

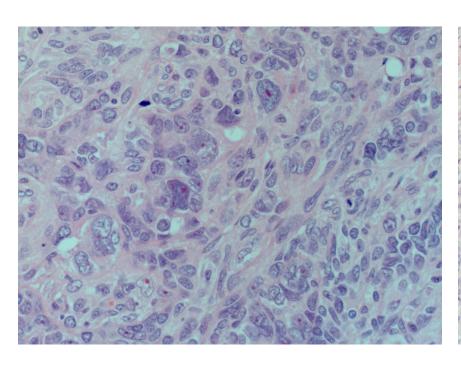
Revision: leiomyoma variants

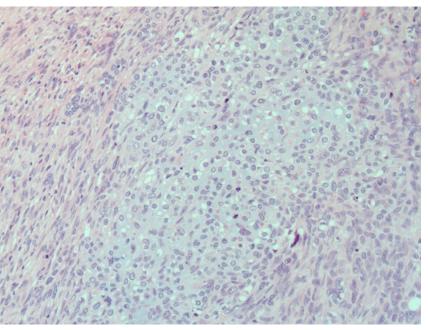
- 10S246
- 59/F Ovarian mass











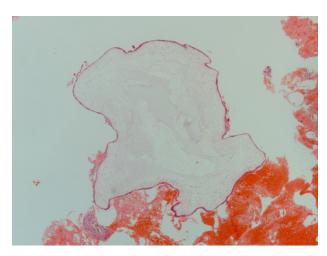
High grade carcinosarcoma (Malignant Mixed Mullerian Tumour)

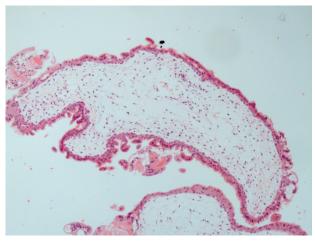
- Most represent carcinoma with sarcomatous transformation
- Biphasic with intermingled carcinomatous and sarcomatous components
- One component may overwhelmingly predominate
- Carcinoma: serous/ high grade endometrioid/ mucinous/ clear cell/ squamous/ mixed
- Sarcoma: homologous (LMS/ fibrosarcoma) or heterologous (cartilage/ skeletal muscle)
- Metastases often purely carcinomatous, rarely purely sarcomatous

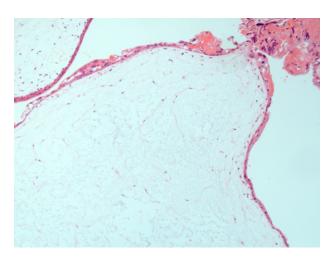
High grade carcinosarcoma (Malignant Mixed Mullerian Tumour) – Differential diagnosis (in endometrial tumours)

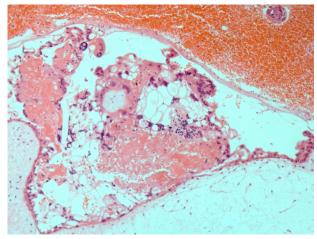
- Mullerian adenosarcoma
- Dedifferentiated endometrial carcinoma
- Sarcomatoid carcinoma = endometrioid carcinoma with spindle cells
- Pure uterine sarcomas
- Endometrioid carcinoma with heterologous elements (cartilage/osteoid/fat, NOT sarcomatous)
- Uterine (or ovarian) serous carcinoma with prominent desmoplastic stroma

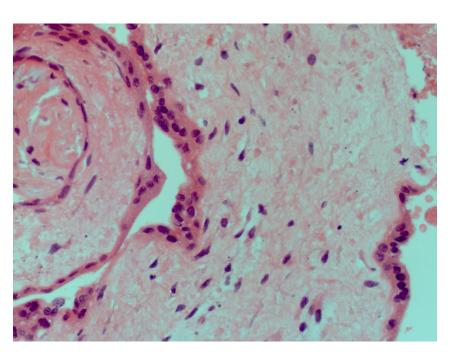
- 09S32117
- 28/F POC

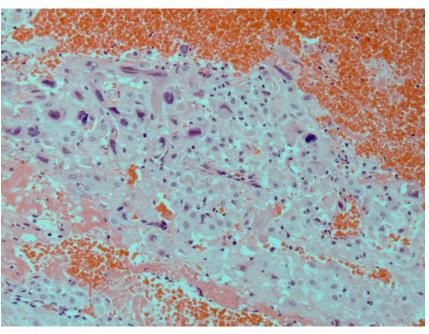






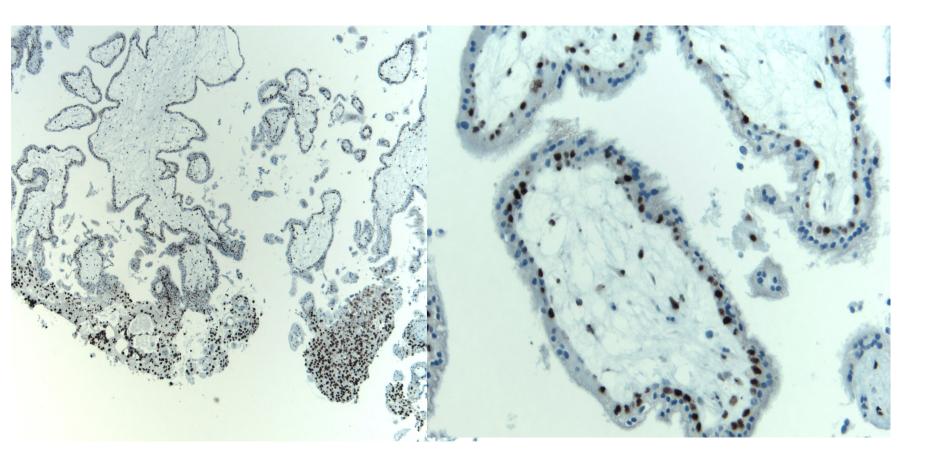






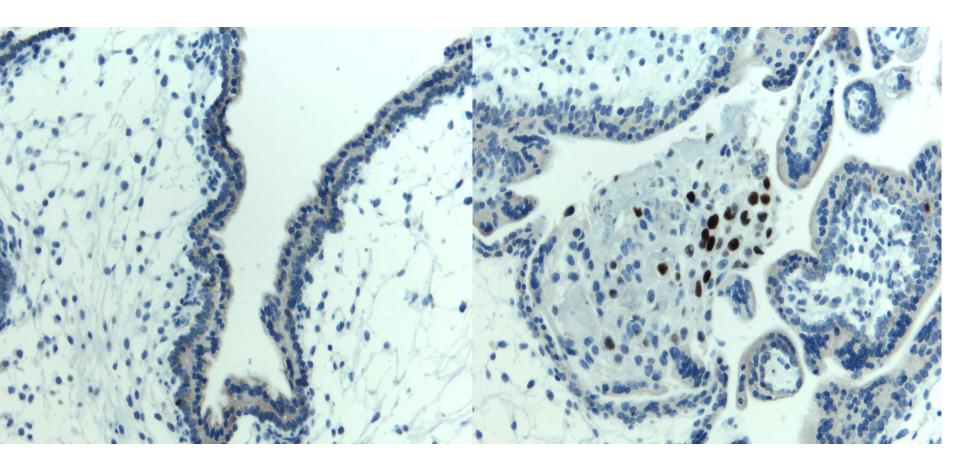
Hydatidiform mole - complete

- Diffuse villous hydropic change, cisterns
- Diffuse trophoblastic hyperplasia, circumferential
- Karyorrhectic <u>stromal</u> debris (NOT vascular)
- Cytotrophoblast atypical
- Syncytiotrophoblast lacelike
- Inclusions not as frequent as in PM
- Intervillous trophoblast is hyperplastic and atypical
- Fetal tissues + in <2% of CM
- Early CM: Subtle club like protrusions, myxoid stroma with karyorrhexis, at least focal circumferential trophoblast hyperplasia
- P57 negative



Intervillous trophoblast is p57 POSITIVE in **all** POCs

Villous cytotrophoblast POSITIVE in non-molar and PHM



CHM: Villous cytotrophoblast and villoous stromal cells NEGATIVE

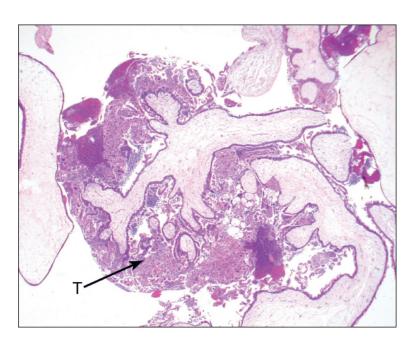
!Intervillous trophoblast will be POSITIVE!

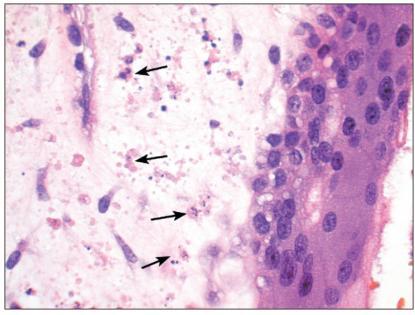
Complete mole – Differential diagnosis

- Partial mole
- Non-molar hydropic change

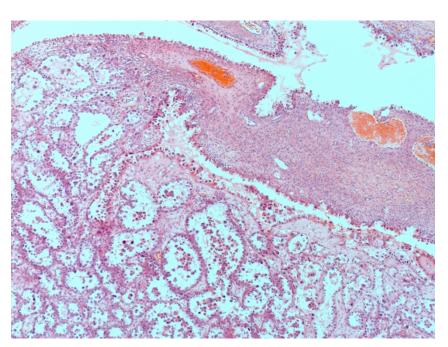
Revision: Pathology of molar pregnancy; pathology of miscarriage (including early and recurrent - has no role in establishing cause of fetal demise – except molar pregnancy)

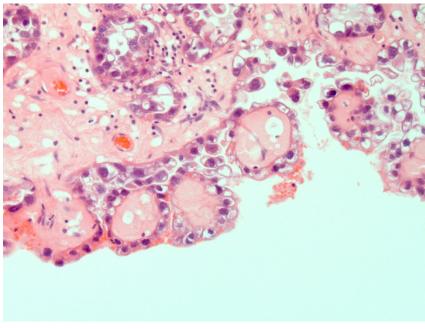
Early complete mole

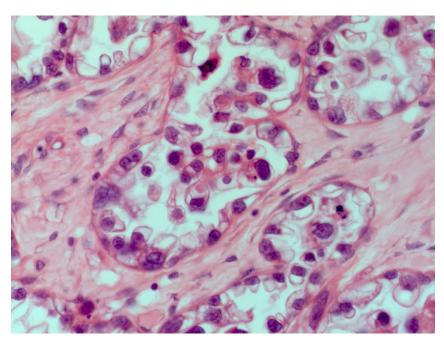


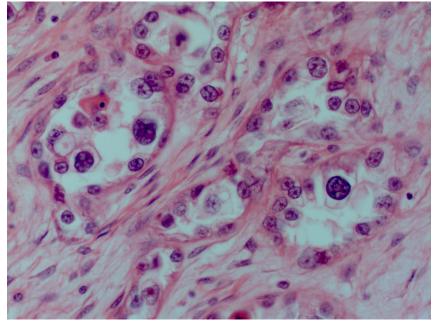


- 09S36075
- 56/F Ovarian mass









Clear cell carcinoma

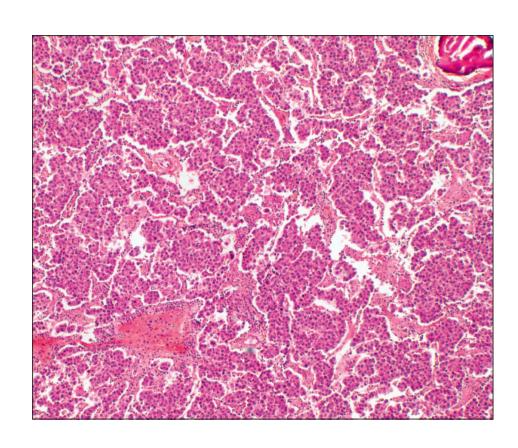
- Variety of architectural patterns: commonly tubulopapillary with hyalinised cores
- Clear cells: polyhedral, distinct cell outlines, eccentric nuclei, glycogen-rich
- Hobnail: bulbous dark nuclei protruding into lumina
- Oxyphilic: eosinophilic cytoplasm
- Others: Cells may be flat and deceptively bland, signet ring, undifferentiated
- Luminal contents: mucin or colloid-like
- Background endometriosis
- New IHC marker: HNF1β

Clear cell carcinoma— Differential diagnosis

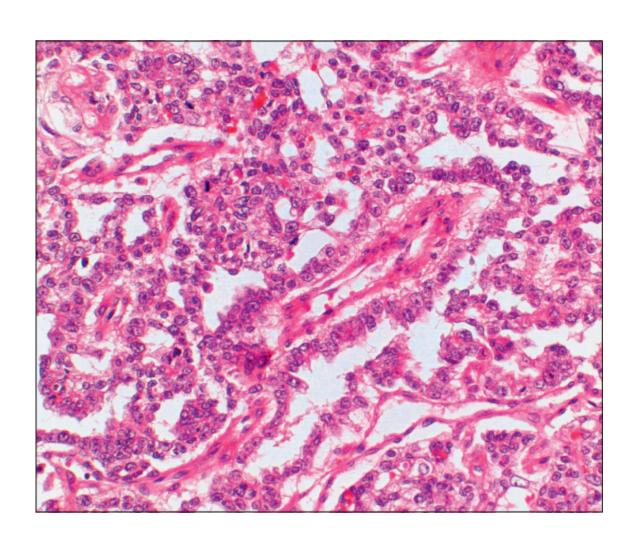
- Other ovarian primaries: endometrioid; YST; dysgerminoma; Brenner; struma ovarii; Sertoli; steroid cell; epithelioid smooth muscle tumours
- Metastatic tumours: other FGT CCC;
 Melanoma; renal cell ca
- Arias-Stella reaction in endometriosis
- Benign/BL clear cell neoplasms

Revision: Synchronous tumours of FGT

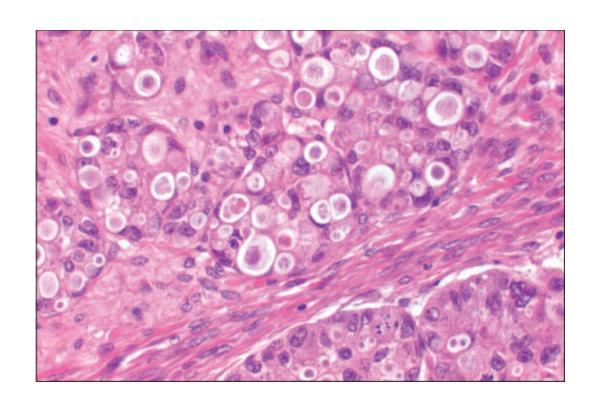
Dysgerminoma



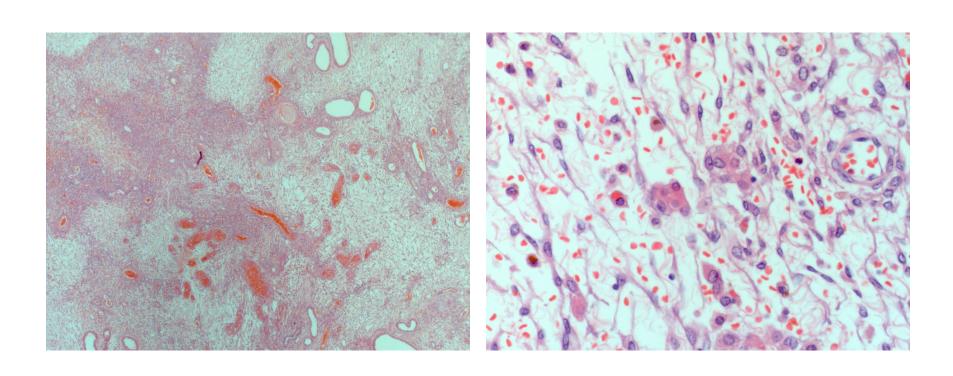
Yolk sac tumour

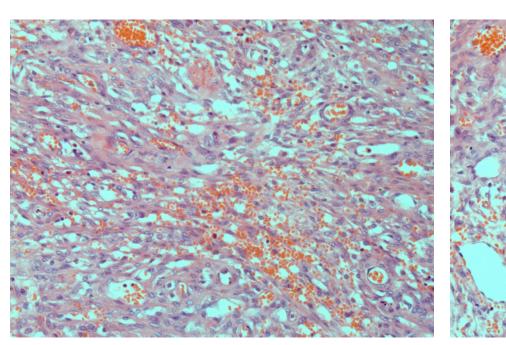


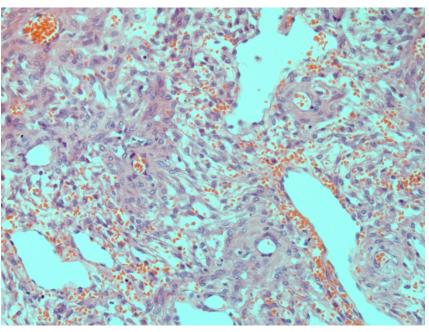
CCC with signet ring morphology

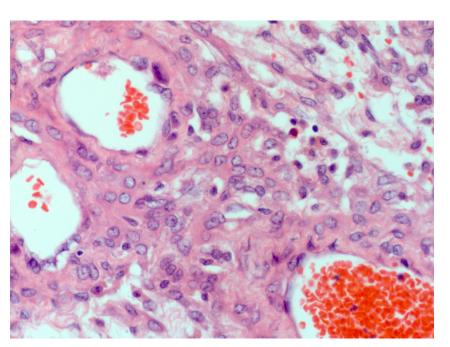


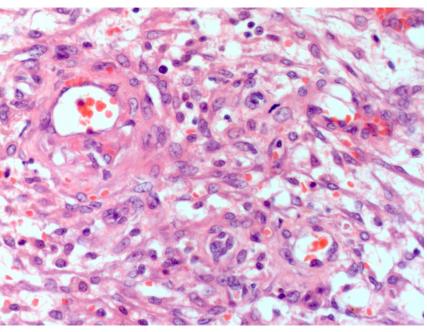
- 09S35285
- 36/F Torsion of right ovary











Sclerosing Stromal Tumour

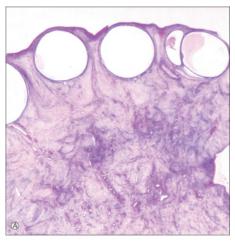
- Unilateral, discrete, sharply demarcated
- Pseudolobular pattern
- Cellular areas alternating with paucicellular densely collagenous or oedematous areas
- Sclerosis variable
- Prominent thin-walled vessels, hemangiopericytomatous pattern
- Cellular areas have disorganised mixture of stromal cells, vacuolated cells, luteinised cells

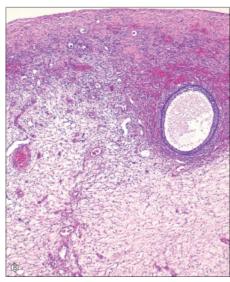
Sclerosing Stromal Tumour – Differential diagnosis

- Fibroma, thecoma
- Krukenberg tumour
- Hemangiopericytoma

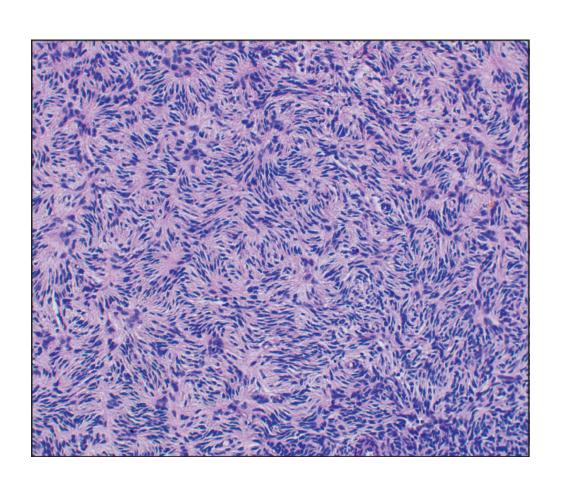
Revision: Ovarian sex-cord stromal tumours

Massive ovarian oedema

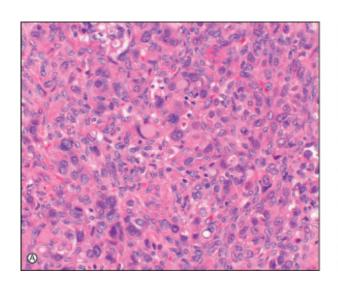


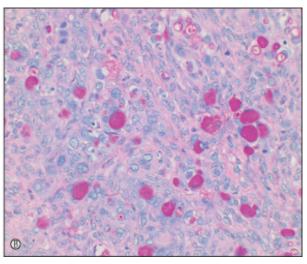


Ovarian fibroma

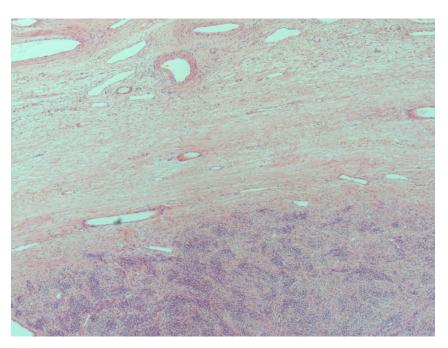


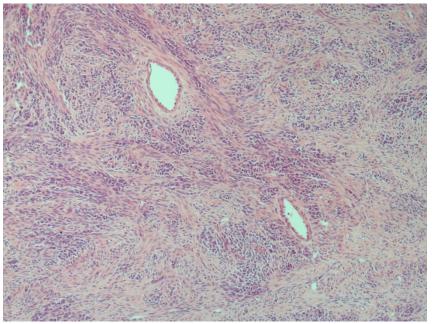
Krukenberg tumour

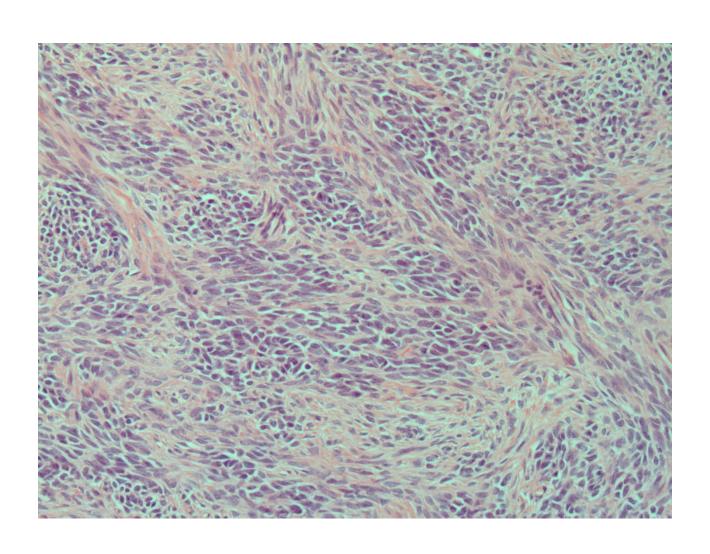


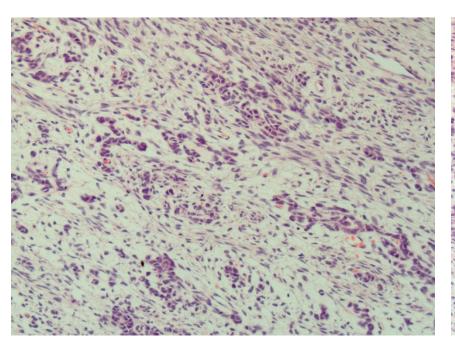


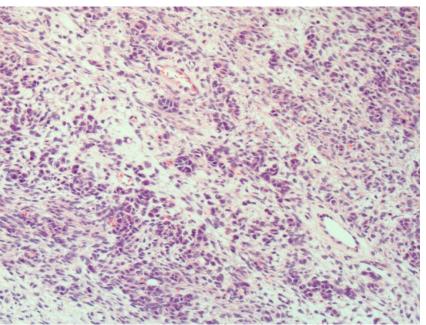
- 09S31420
- 33/F Complex right ovarian mass with raised CA125

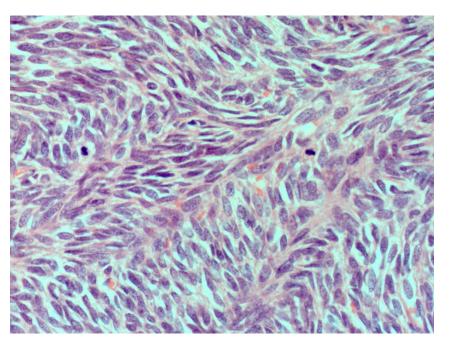


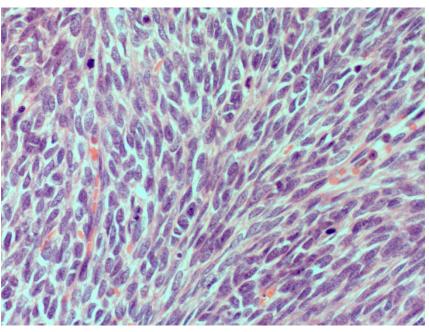












Mitotically active cellular fibroma

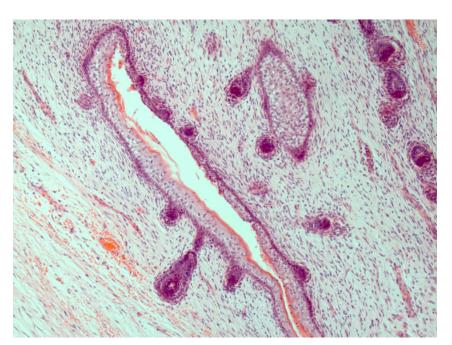
- Cellular proliferation of spindle cells
- Variable lipidisation
- >4 mf/10 hpf
- Minor component o sex-cord elements
- Pseudocysts
- Necrosis
- NO/MILD/FOCAL MODERATE CYTOLOGICAL ATYPIA

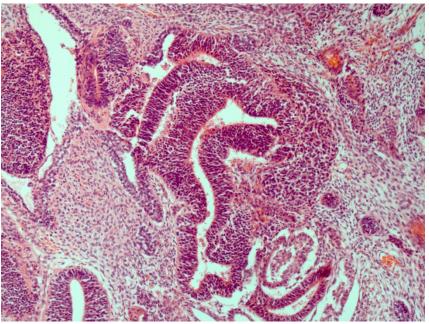
Mitotically active cellular fibroma – Differential diagnosis

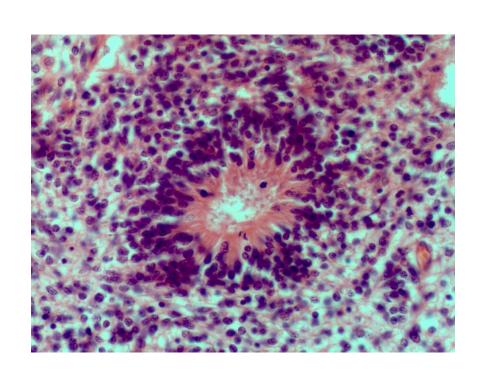
Fibrosarcoma

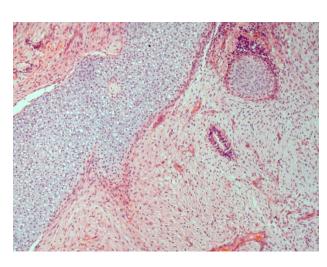
Revision: Diagnostic features of MACF

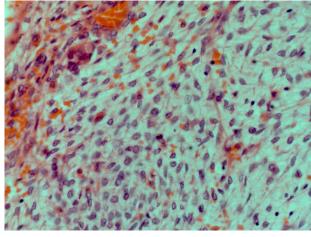
- 09S30787
- 31/F Complex right ovarian mass with raised CA125 and AFP











Immature teratoma

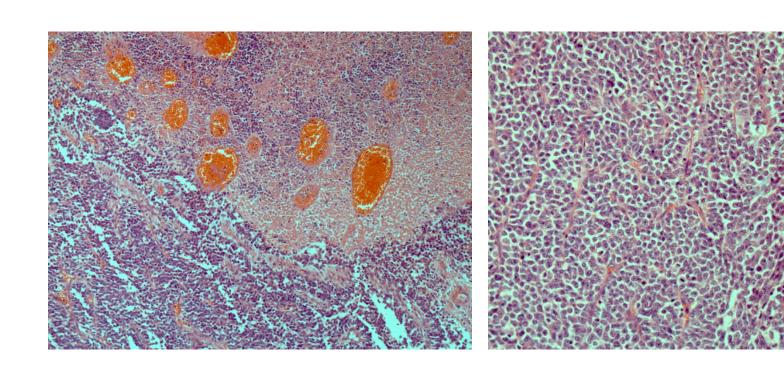
- Diagnostic feature: Presence of embryonic-appearing tissue - usually neuroectodermal as tubules, rosettes; cellular and mitotically active glial tissue
- Other immature tissues: cartilage, skeletal muscle, epithelial, hepatic

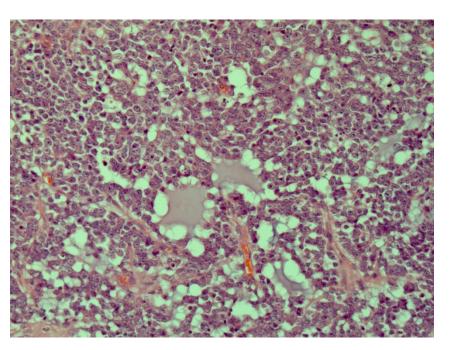
Grading of Immature Teratoma

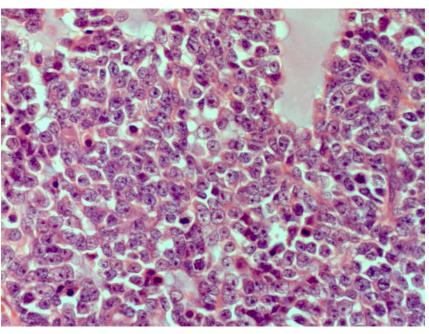
- Grade 1: rare foci <1 per lpf in any one slide
- Grade 2: 2 or 3 foci per lpf in any one slide
- Grade 3: >4 foci per lpf in one slide

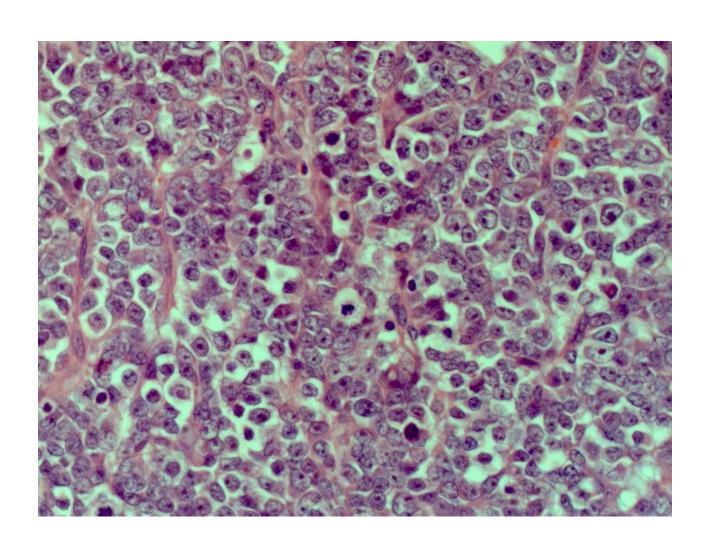
Revision: Peritoneal gliomatosis and growing teratoma syndrome

- 06S14449
- 36/F 6 months post TAH + BSO for granulosa cell tumour ovary Stage 1A. Now pelvic recurrence









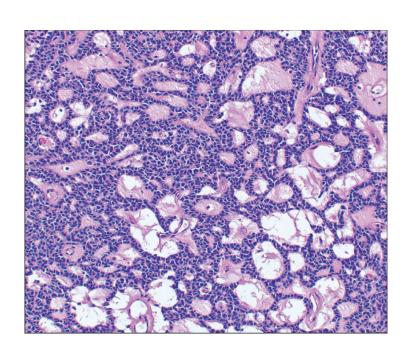
Small cell carcinoma of ovary (hypercalcemic type)

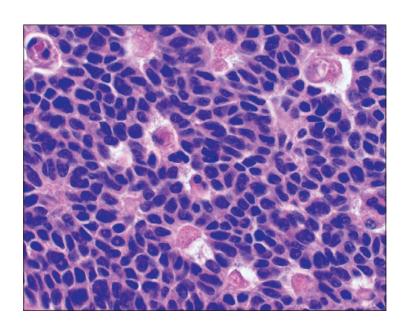
- Solid architecture, sheets of malignant cells
- Variably sized follicle like spaces typical
- Tumour cells small, round, scanty cytoplasm, hyperchromatic nuclei, single small nucleoli
- Frequent mitotic figures
- Large cells seen focally but may predominate
- Minor foci of the following may occur: spindle cells, mucinous epithelium, clear cells
- Vascular invasion ++
- IHC: CK, EMA, WT1, CD10, p53 positive
- 2/3rd of cases have hypercalcaemia
- SMARCA4 mutation (2014; SCCHT may be hereditary)

Small cell carcinoma of hypercalcemic type— Differential diagnosis

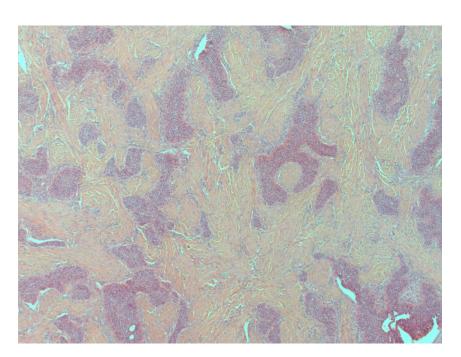
- Granulosa cell tumour
- Lymphoma
- Small cell ca of pulmonary type
- Dysgerminoma
- Melanoma
- Primitive round cell tumours eg PNET
- Undifferentiated/ poorly differentiated ca

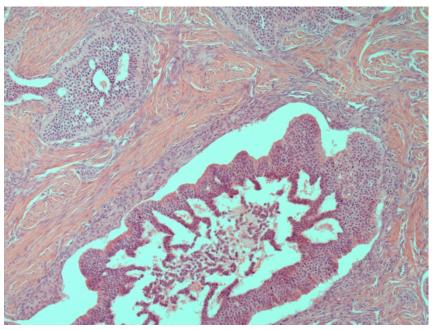
Granulosa cell tumour

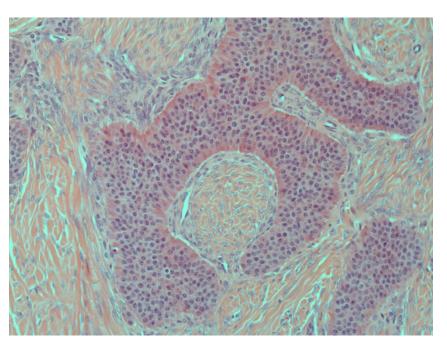


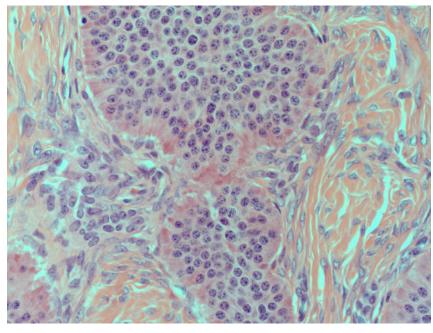


- 08S35916
- 50/F Ovarian mass. Normal CA125. Low RMI









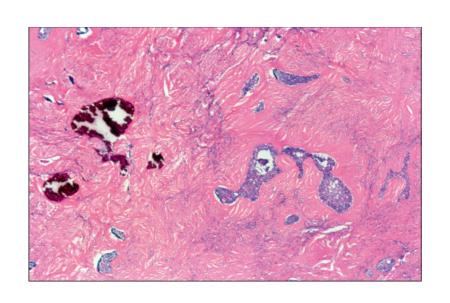
Insular carcinoid

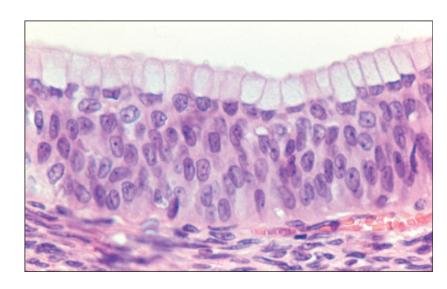
- Discrete cellular nests with variable numbers of acini
- Separated by fibrous stroma
- Tumours cells have moderate amount of eosinophilic cytoplasm, red-brown argentaffin granules, sometimes oxyphilic
- Nuclei round and uniform, finely stippled chromatin
- Mitosis rare or absent
- Positive for neuroendocrine markers

Insular carcinoid— Differential diagnosis

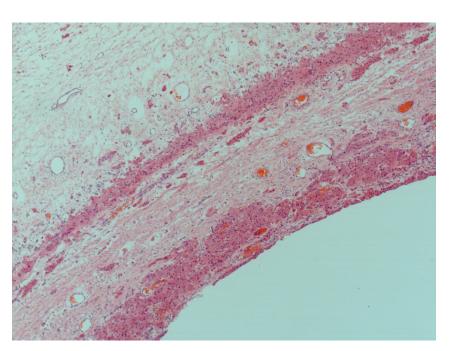
- Metastatic carcinoid tumour
- Granulosa cell tumour with microfollicular pattern
- Brenner tumour

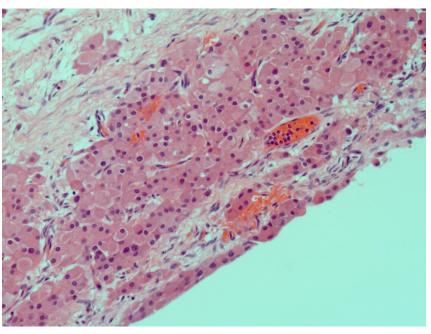
Brenner tumour

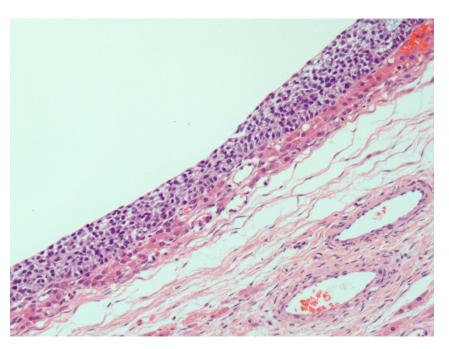


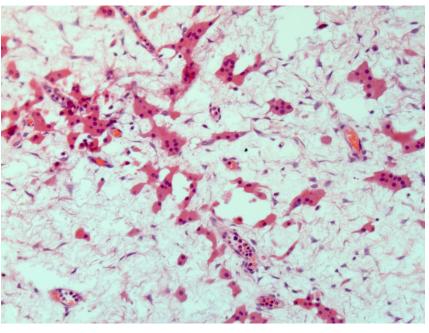


- 07S27230
- 27/F Right ovarian cyst found at LSCS,
 12cm diameter





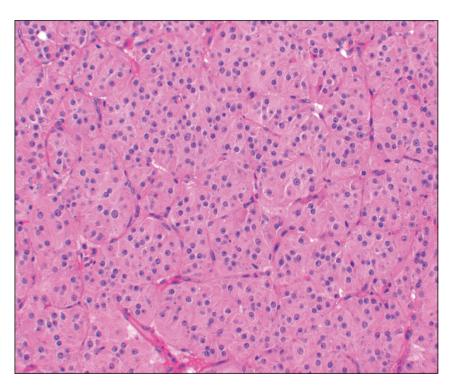




Hyperreactio luteinalis

- Associated with increased HCG levels: GTD, multiple gestation, normal singleton pregnancy, ovarian hyperstimulation (after ovulation in women who conceive)
- Removal necessitated by cyst complication or virilisation; or because of clinical suspicion
- Bilateral, multiple thin-walled cysts causing ovarian enlargement
- Multiple large luteinised follicles; oedema of stroma; groups of luteinised stromal cells

Tumour like conditions of pregnancy

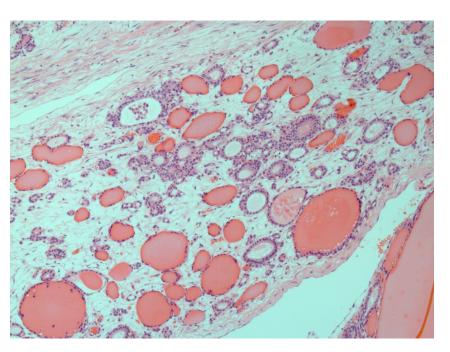


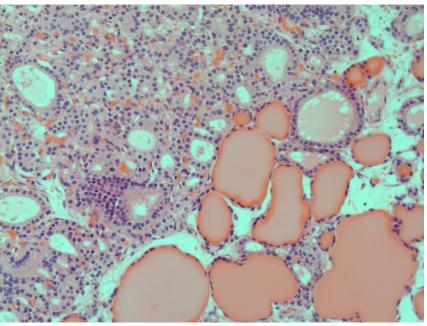
Pregnancy luteoma



Giant solitary luteinised follicular cyst of pregnancy and puerperium

- 10S10059
- 55/F Ovarian mass. CA-125 44. Normal germ cell markers



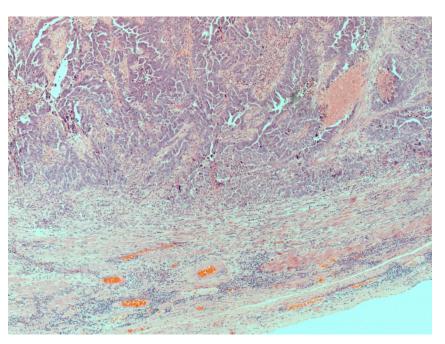


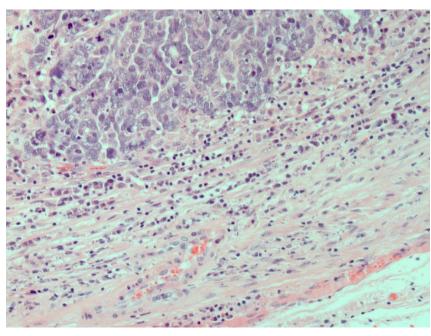
Struma ovarii

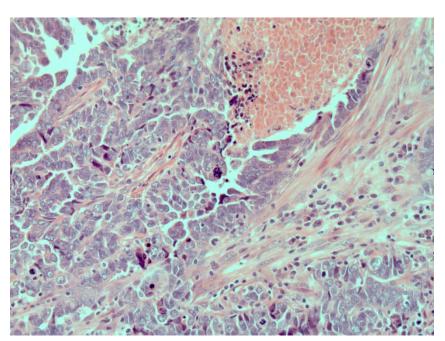
- Mature thyroid tissue, sometimes resembles follicular adenoma
- No or very mild atypia
- Malignancy:
 - Papillary carcinoma
 - Follicular carcinoma
 - Non-specific adenocarcinoma resembling endometrioid

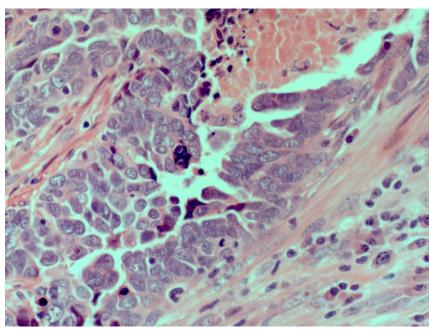
Struma ovarii – Differential diagnosis

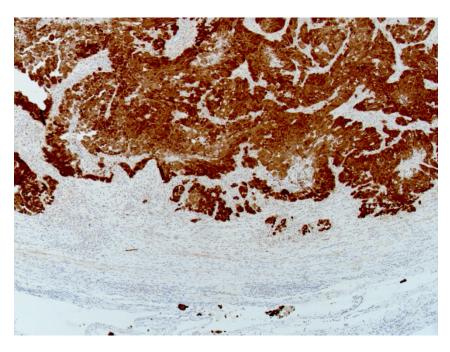
- Cystadenoma
- Clear cell carcinoma
- Endometrioid carcinoma
- Other tumours can be simulated in presence of clear cell or oxyphilic change

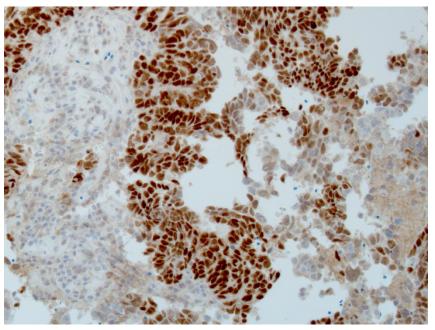












Primary high grade serous carcinoma of Fallopian tube

- Papillary architecture
- High grade adenocarcinoma cells of serous type
- Psammoma bodies
- Invasion

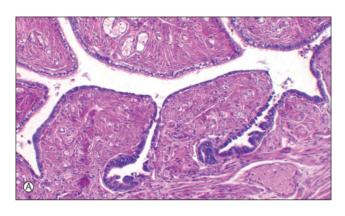
Primary high grade serous carcinoma of Fallopian tube

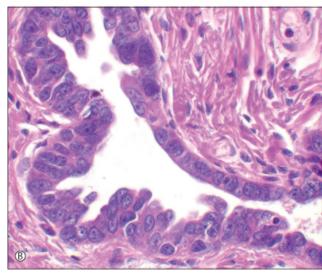
DD: Secondary involvement from ovarian or endometrial primary

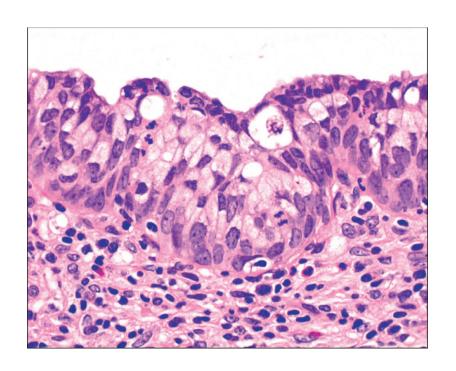
Revision: IMPORTANT

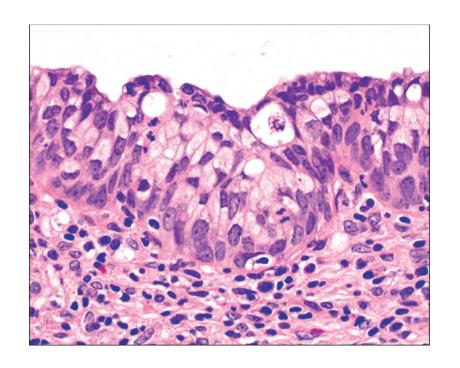
- Classification of serous carcinoma (LG vs HG, dual origin)
- Concept of tubal origin of serous neoplasia (Crum)

Intraepithelial carcinoma of FT









SMILE....

And good luck!