Ovarian Mucinous Tumours

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Epithelial tumours

- 65-70% serous
- 3% MUCINOUS
- 9-11% endometrioid
- 12-13% clear cell
- 1% transitional
- 6% mixed ??

 Mucinous carcinoma – 3% of all epithelial tumours

- Older literature 12-%
- Reason for decline-

Better awareness and recognition of secondaries Better defined criteria for BMT

What's new

- Until 2014 classification two types intestinal type and endocervical type/Mullerian-type
- Now only one type gastrointestinal-type
- Endocervical type renamed seromucinoustype

General issues

Benign ---- Borderline ---- Carcinoma

- Benign vs borderline
- Borderline vs malignant
- Other tumours with mucinous differentiation
- Primary vs secondary
- Appendiceal pathology

Presentation

- wide age range but may occur in <u>young</u> females)
- large and unilateral
- usually confined to ovary
- multiloculated with or without solid areas
- Ca 125 normal or just mildly elevated
- serum CA19.9 and CEA

Mucinous tumours

- Benign mucinous cystadenoma
- Borderline mucinous tumour (BMT)
- BMT with microinvasion
- Microinvasive carcinoma
- Adenocarcinoma

Molecular framework

- KRAS mutations common (occur early in pathway)
- Tp53 mutation occurs in minority of ovarian mucinous carcinomas
- minority arise in teratomas (may exhibit upper or lower intestinal differentiation)

Macroscopic

- Thorough examination of every locule
- 1 block/cm
- More from solid areas but also cystic areas
- Look for capsular breach and secondary signs of leakage
- Look for adhesions (may upstage)

Stage I (FIGO 2014)

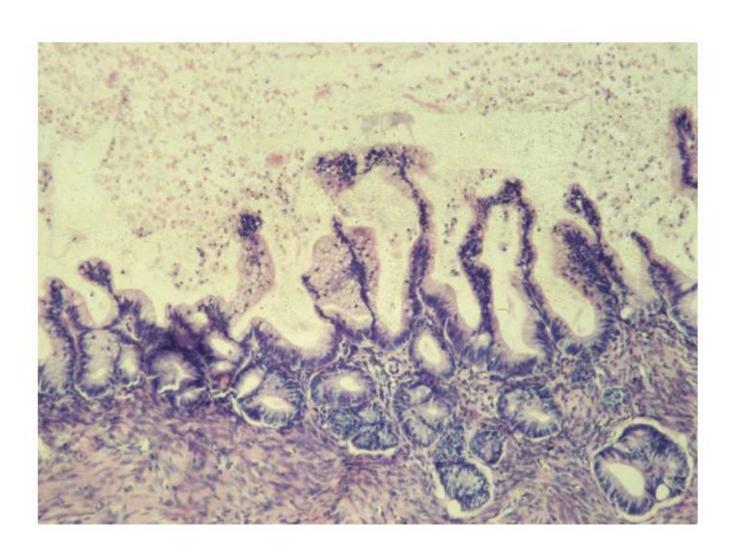
Stage I		Growth limited to ovaries		
IA	T1a N0 M0	Growth limited to one ovary; no tumour on the external surface, capsule intact, no ascites		
IB	T1b N0 M0	Growth limited to both ovaries; no tumour on the external surface, capsule intact, no ascites		
IC	T1c N0 M0	Tumor limited to one or both ovaries		
	IC1	Surgical spill		
	IC2	Capsule rupture before surgery or tumor on ovarian surface		
	IC3	Malignant cells in the ascites or peritoneal washings		

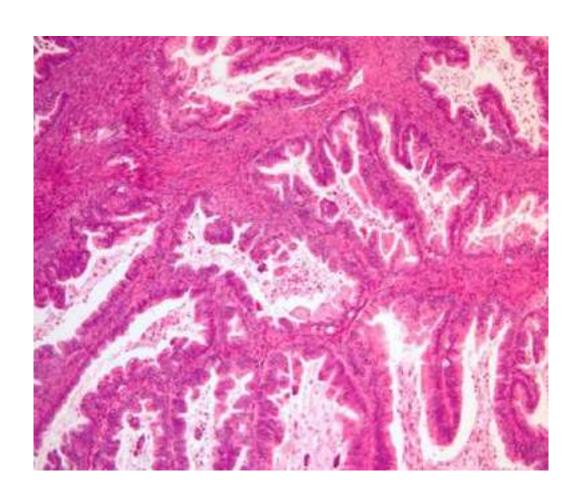
Cystadenoma/adenofibroma

- Gastric-type or intestinal type or both type of epithelia
- <10% epithelial proliferation
- Association with Brenner tumour is well known

Borderline

- Proliferation in >10%
- Nuclear stratification, mild to mod nuclear atypia, occasional mitoses
- Villous architecture, papillation





Microinvasion vs microinvasive

- Microinvasion- occasional scattered cells
 - no stromal reaction
 - <5mm
 - no clinical implication

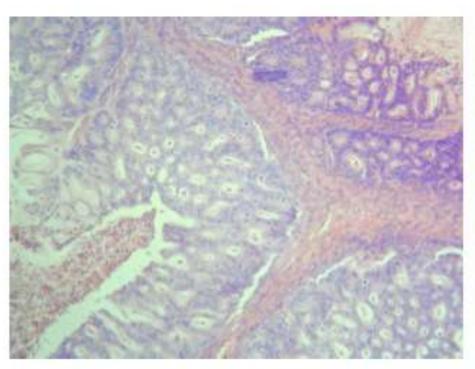
Microinvasive – small focus of high grade carcinoma

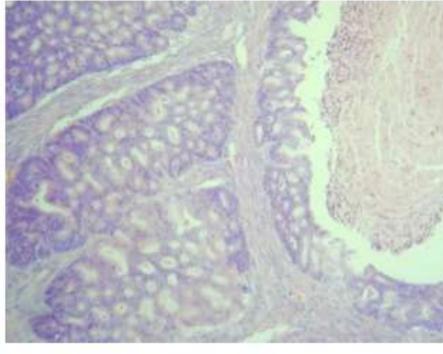
poor prognosis

Invasion

- Expansile vs infiltrative
- DD of muc ca with expansile invasion from BMT
- Pathologist should state what type of invasion
- Grading is like in endometrioid carcinoma

CRIBRIFORM





Expansile invasion (slide)

- back to back small to medium sized glands with stromal exclusion
- cribriform growth pattern
- nuclear atypia, although may be low grade
- at least 5mm area
- marked inter-observer variability
- LOW POWER DIAGNOSIS

Destructive invasion

- obvious invasion- easy to diagnose but not very common
- always think of secondary

PROBLEMS WITH OVARIAN MUCINOUS NEOPLASMS

- BL vs carcinoma
 - -significant inter-observer variation

- Primary vs secondary
 - -variable threshold to suggest further investigations to rule out metastasis
 - cause of unrest/debate/arguments in MDM

Prognosis

- Borderline -TUMOUR OF NO MALIGNANT POTENTIALmay recur following cystectomy and possibly spillage
- borderline with intraepithelial carcinoma (extremely good prognosis)
- borderline with microinvasion (extremely good prognosis provided adequate sampling)
- stage I carcinoma with expansile invasion (good prognosis)
- stage I carcinoma with destructive invasion (guarded prognosis)
- advanced stage mucinous carcinoma (poor prognosis; exclude secondary)

Immunohistochemistry

CK 7, CK20, CDX 2, CEA, p16

Ovary – CK7 > CK20 (unless..), PAX8, (CDX2,ca19.9)

Colon - CK20 > CK7, CDX2

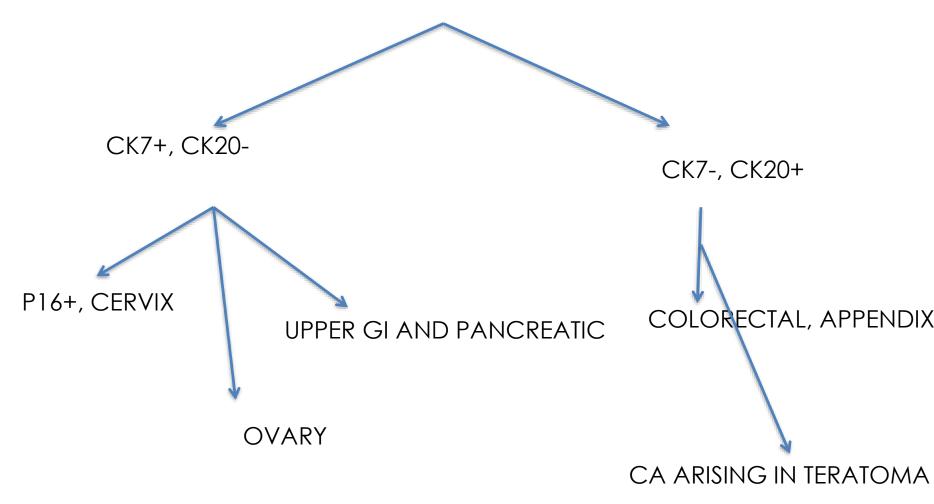
Appendix CK20 +, CK7 -

Pancreatobiliary – CK7 > CK20

Stomach - CK7 > CK20

Cervix – CK7 > CK20, p16 +

CK7, CK20



Seromucinous tumours

- New category in WHO 2014 classification
- Previously classified as endocervical type mucinous tumours
- Benign, borderline and low grade carcinoma spectrum
- Arise in background of endometriosis
- Can be predominantly serous or mucinous type posing diagnostic challenge

Seromucinous tumours- morphology

Mixed epithelia
Associated with endometriosis
Inflammatory infiltrate is a constant feature

Differential diagnosis

- If predominantly mucinous ER and vimentin positive, CDX2 negative
- If predominantly serous papillary architecture
 - WT1 negative

- Mixed mucinous, serous, clear cell, endometrioid areas
- Association with endometriosis
- Presence of polymorphs

How do we grade?

 Mucinous carcinoma is graded like endometrioid carcinoma

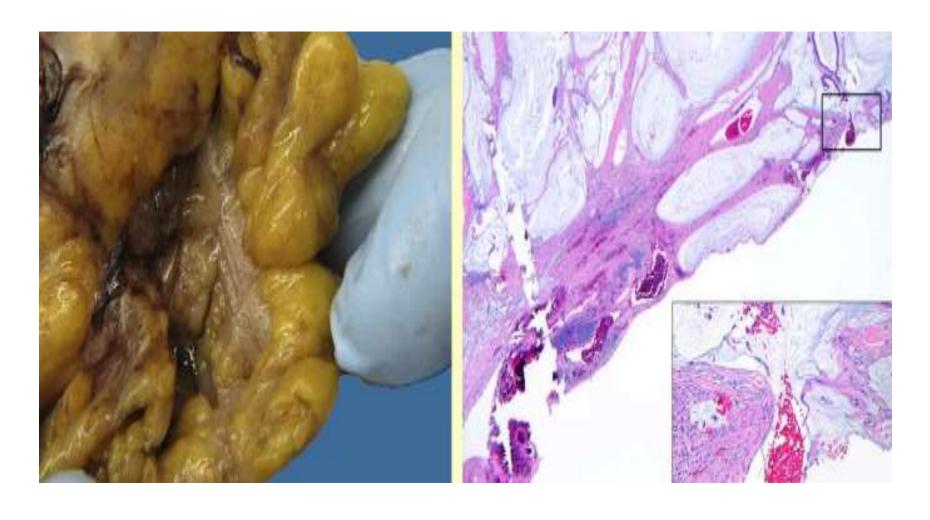
Appendix

- Hyperplastic polyp
- SSP
- LAMN
- AC
- GCC

• Biopsy of the omental nodule

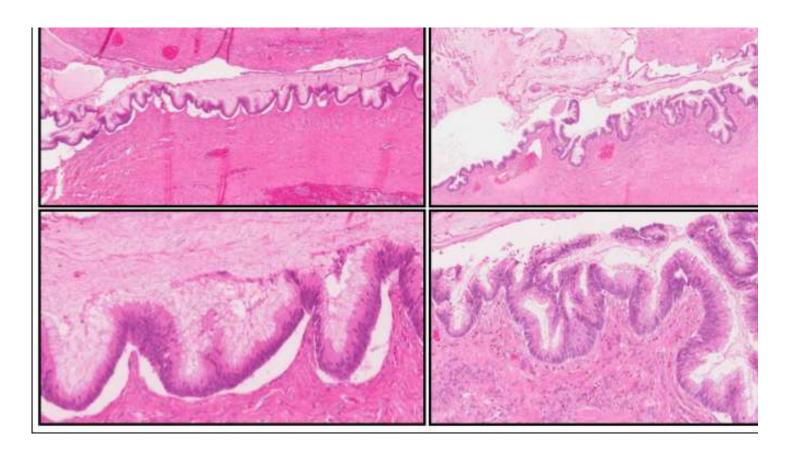
Debulking surgery

PMP

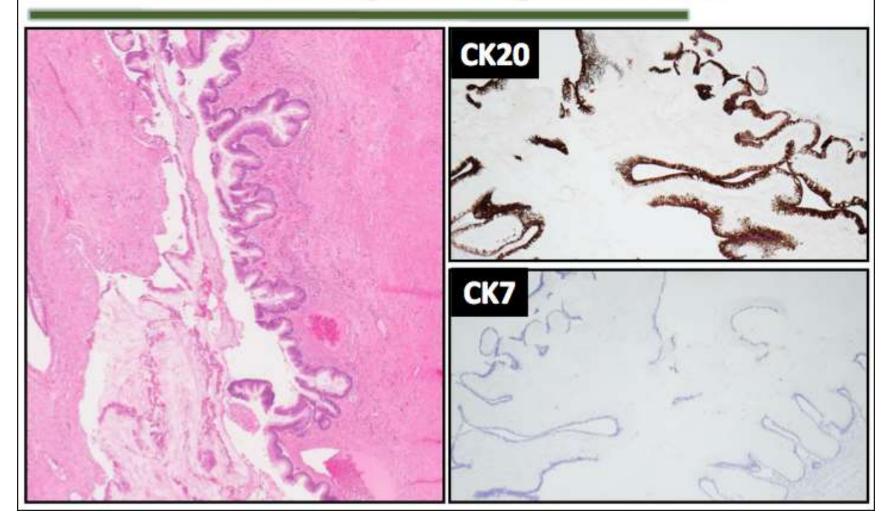


Case of the month (BAGP)

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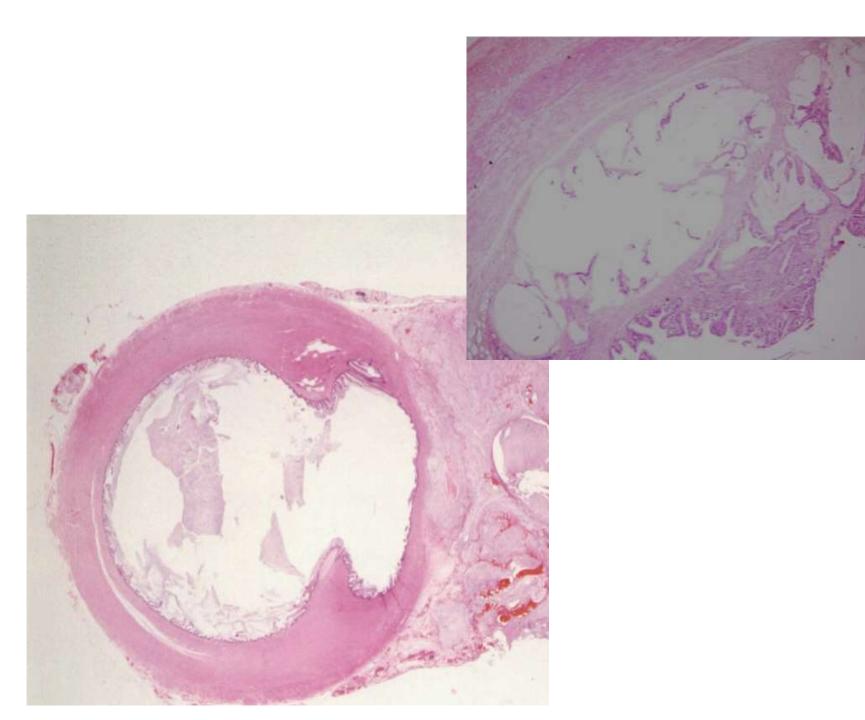


Ovarian tumour + mucinous ascites Pseudomyxoma peritoneii



Mucocele





LAMN(Low garde Appendiceal Mucinous Neoplasm)

- Villous or cystic epithelial neoplasm
- Low grade dysplasia
- May be associated with ovarian involvement
- Risk of PMP

Reporting

- Examination of entire appendix (esp if macroscopically normal)
- Degree of dysplasia
- Perforation present/absent
- Mucin on the serosal surface present /absent
- Acellular vs epithelium outside appendix
- Comment on risk of PMP

Management

- Simple appendicectomy even when ruptured
- Close follow up required because of the possibility of pelvic recurrence

- Presence of any epithelium worsens prognosis
- Must examine multiple sections if no epithelium found in mucin

PMP

- Clinical diagnosis
- Right ovary more commonly affected
- Usually due to spread from appendix
- Appendix may be embedded in mucinous or fibrous mass

Pseudomyxoma peritonei

- Low grade
- High grade

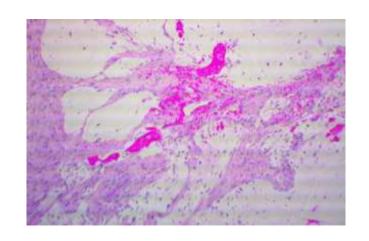
Takes into account

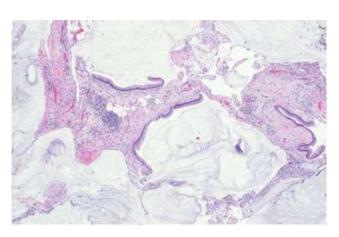
Mucin pools

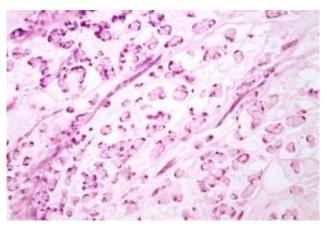
Cytological atypia

Architectural complexity

LG =<10% of epithelium, nonstratified columnar epithelium, bland cytology



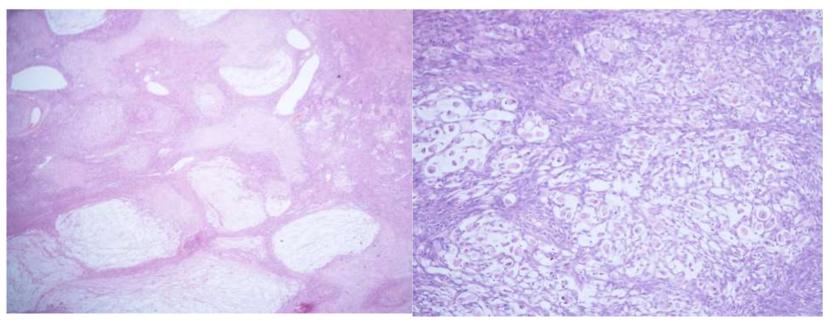




Case discussion

Metastatic carcinoma (CK7 +, CK20-, ER-, ,PAX8 -,

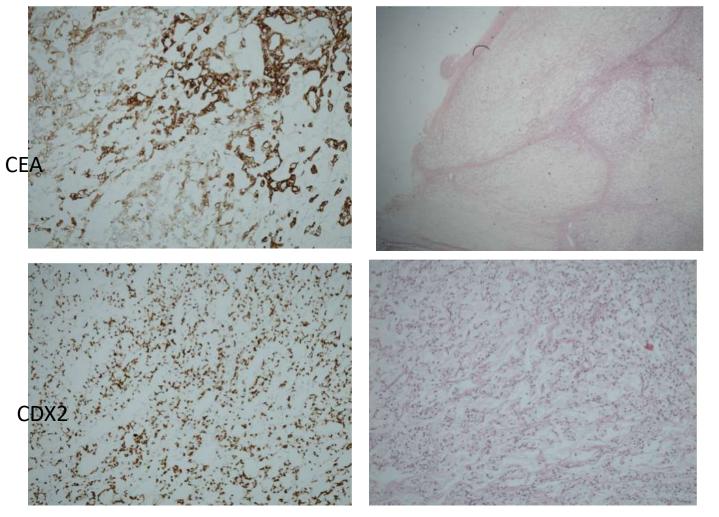
P16-, CDX2 -) upper GI and breast likely primary sites.



Exracellular mucin

Infiltrative invasion Signet ring cells

Case discussion 2 Metastatic mucinous carcinoma



Ck20 + CK7-, CDX2+ (colonic primary)



Secondaries

- Not uncommon
- History of primary may or may not be there
- Unilateral or bilateral
- Solid or cystic
- Often exhibit maturation phenomenon (also consider D/D in BMT)

Features favouring metastasis

- Bilaterality
- <10cm size (small tumours)
- Surface involvement
- Advanced stage

Features favouring metastasis

- Signet ring cells
- Extracellular mucin
- LVI
- Hilar infiltration
- Surface involvement
- Nodular growth pattern

Take home....

- Mucinous tumours tend to be large, multilocular and usually unilateral
- Gross examination and adequate sampling is a key to correct diagnosis
- All ovarian mucinous tumours are now called gastrointestinal type
- Previous endocervical-type is now reclassified as seromucinous tumour

Take home....

- BL vs carcinoma cribriforming, villiformity, severe atypia, necrosis are clue to carcinoma
- In carcinoma grading is like endometrioid carcinoma
- In carcinoma- mention pushing or infiltrative type invasion
- Infiltrative type invasion- think of metastasis
- Clues to metastatic nature bilaterality, <10cm size, multilobulated surface, infiltrative pattern of invasion, LVI

- IHC-
- Metastatic lower GI CK20 +, CK7- (teratoma premise to be excluded)
- Metastatic upper GI CK7+, CK20 -, infiltrative pattern, signet ring morphology
- Seromucinous ER+ PGR+ Pax8 +, CK7+,WT1associated with endometriosis

PMP

Causes – colonic, appendiceal, ovarian

- Appendix examine whole appendix,
- Presence of rupture, acellular/cellular mucin at the surface in the mesoappendix most important prognostic features

That's it....!

Thank you.