

# Cases from My Referral Practice

## *Teaching Files & Dermpathpro Spot Diagnosis*

# Theme : Adnexal Lesions

*Self-Assessment / Spot Diagnosis Challenge*

## Cases 1 to 3

**Benign v's Malignant**

A. Adenoma (**1 point**)

B. Adnexal Ca (**1 point**)

**Subtype (bonus 1 point)**

Papillary adenoma

Hidradenoma

Spiradenoma

Other

Digital papillary AdenoCa

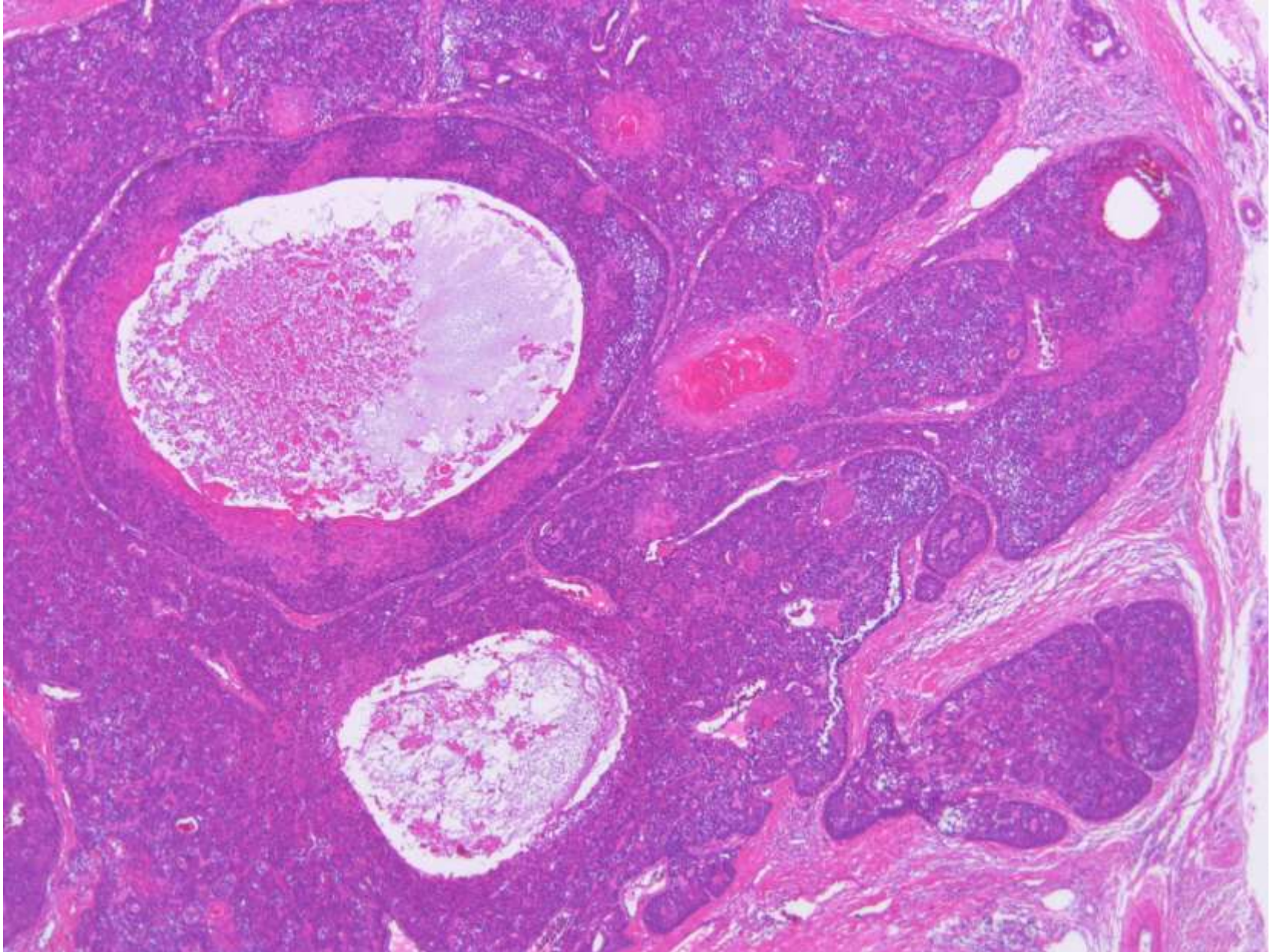
Spiradenocarcinoma

Hidradenocarcinoma

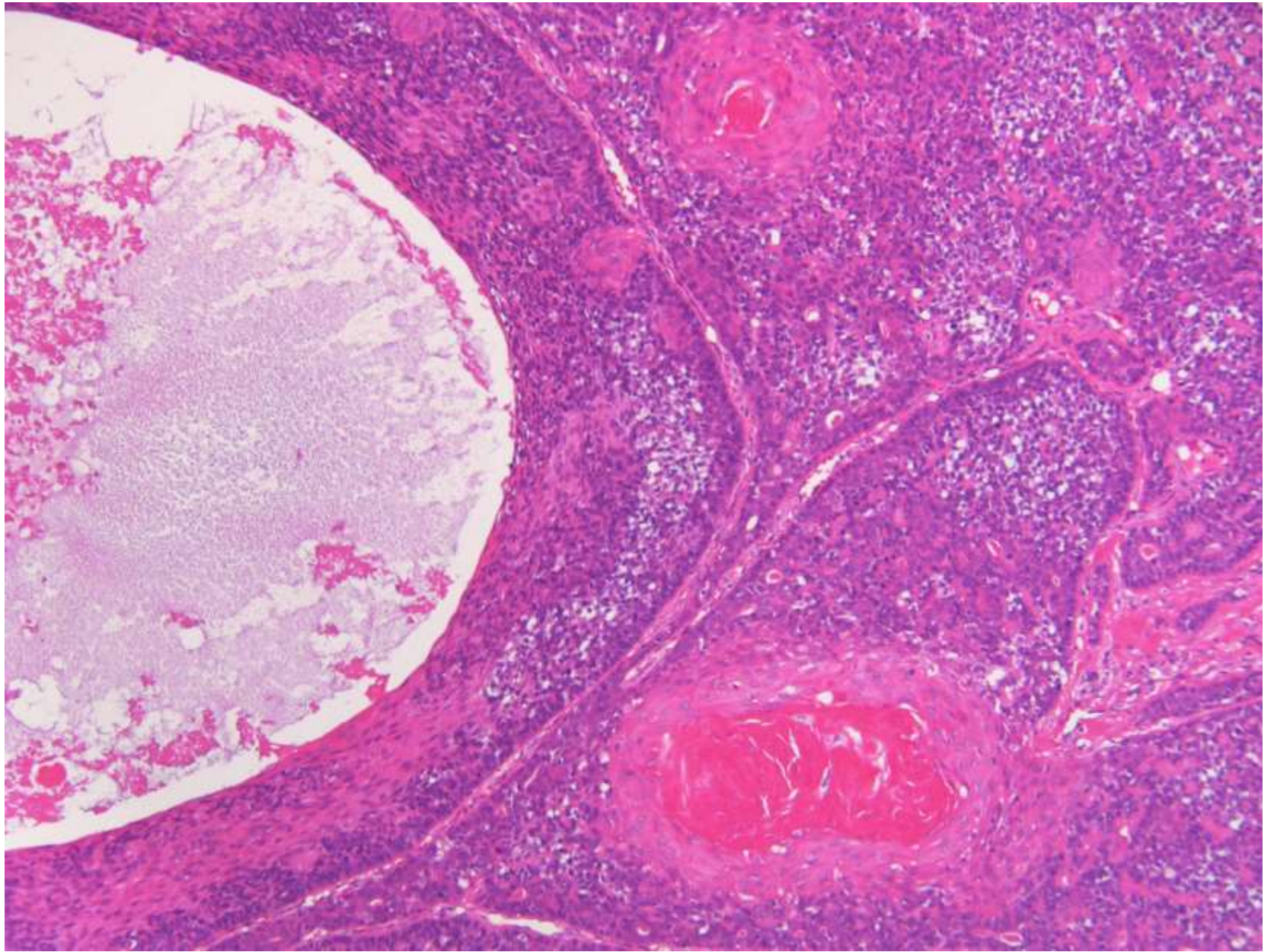
Adenoid cystic carcinoma

Other

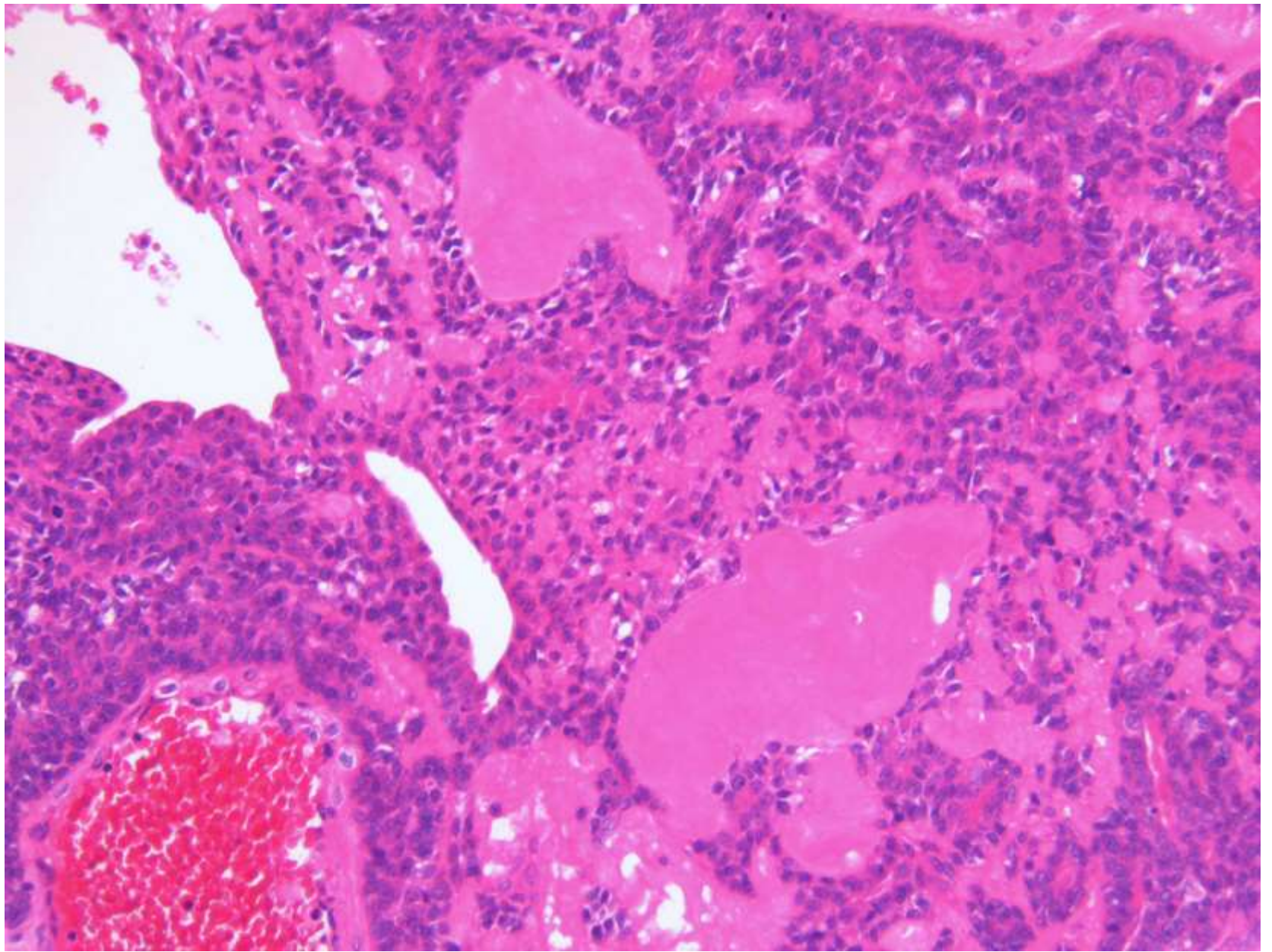
## Spot Diagnosis: Case 1



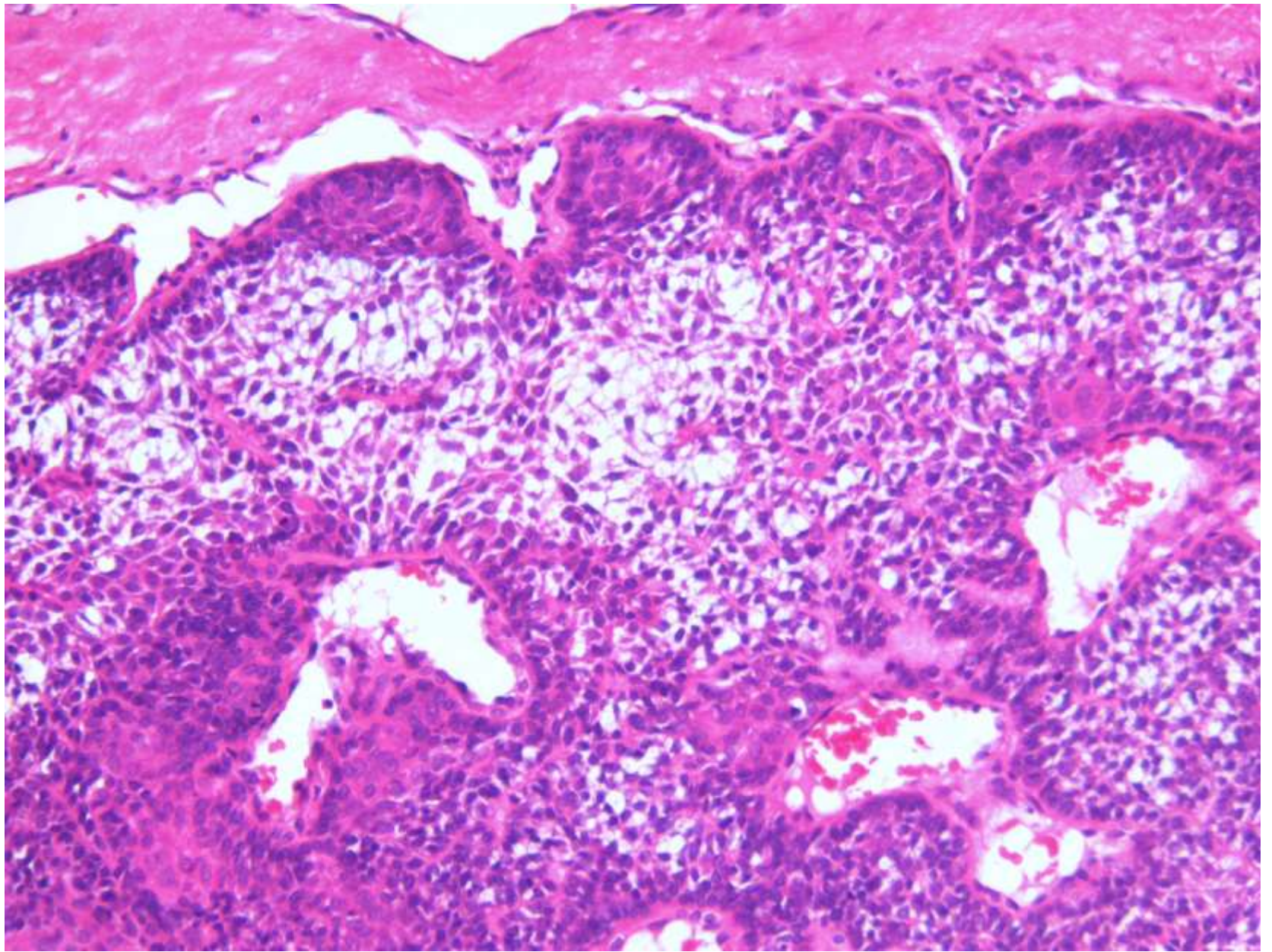














# Case 1

## Benign v's Malignant

A. Adenoma (**1 point**)

Subtype (**bonus 1 point**)

Papillary adenoma

Hidradenoma

Spiradenoma

Other

B. Adnexal Ca (**1 point**)

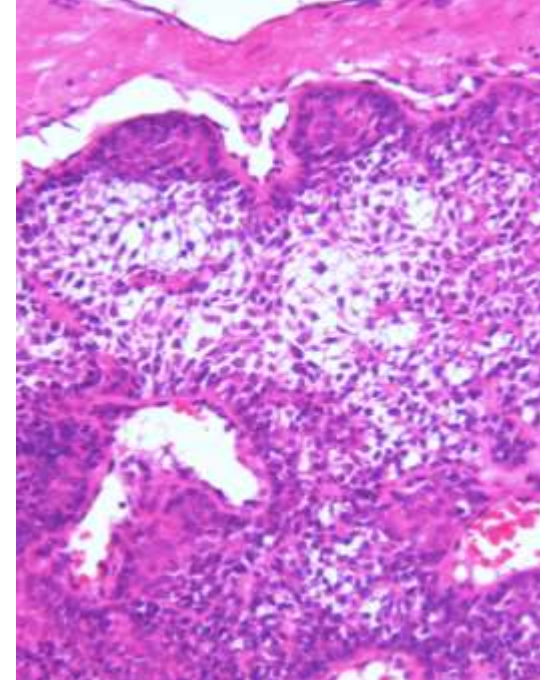
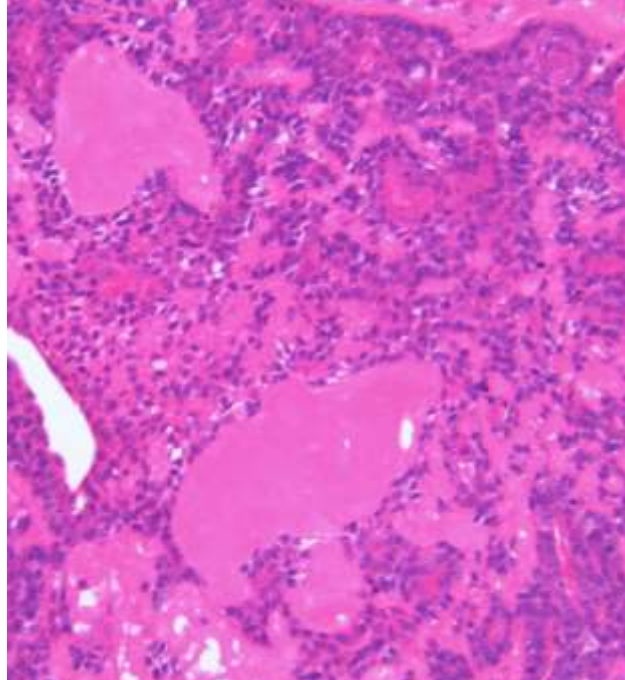
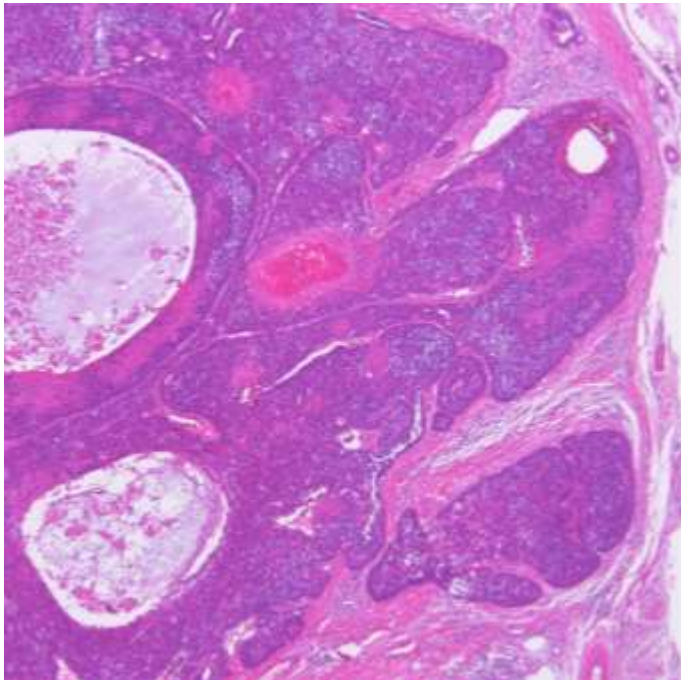
Digital papillary AdenoCa

Spiradenocarcinoma

Hidradenocarcinoma

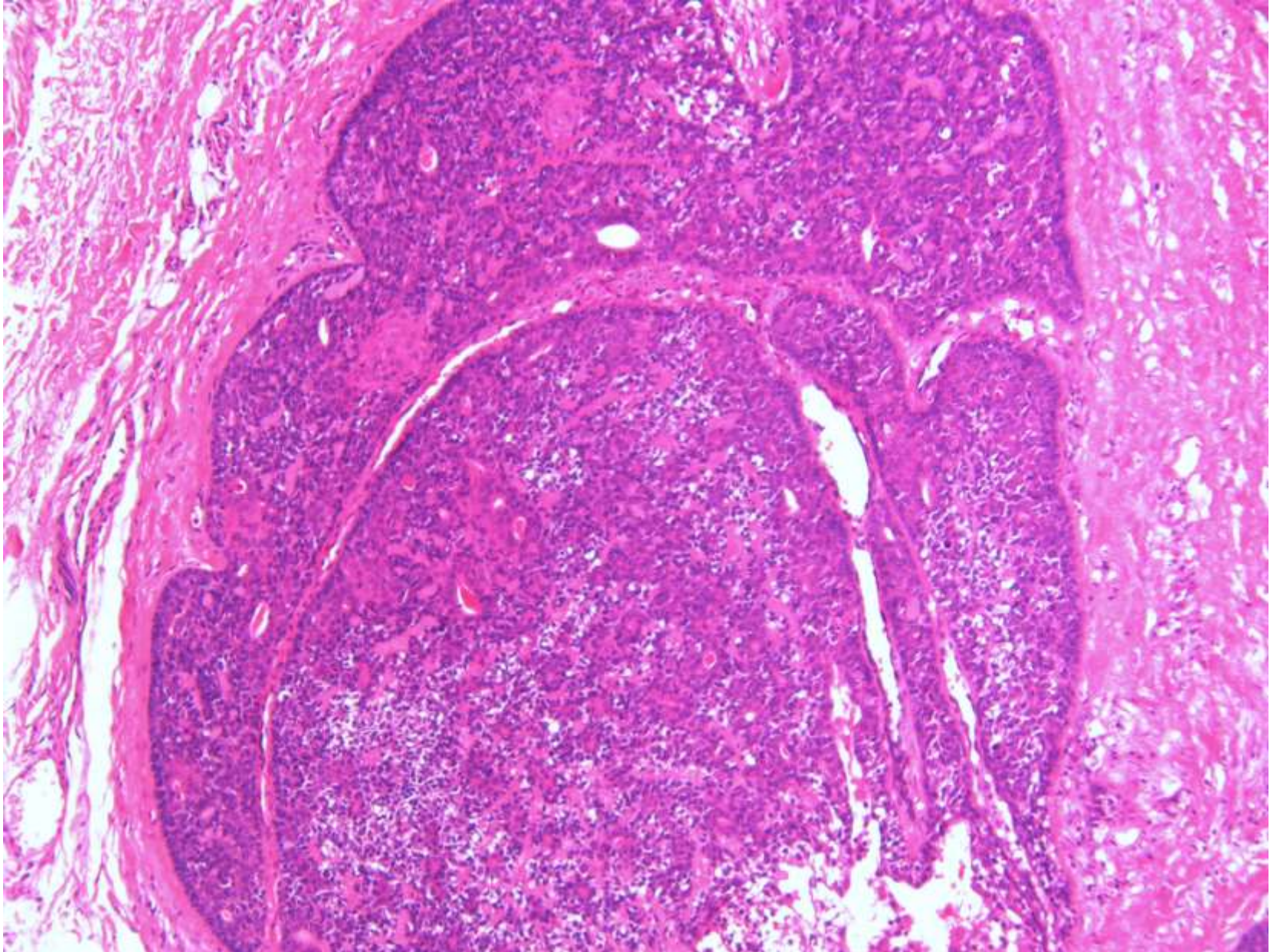
Adenoid cystic carcinoma

Other

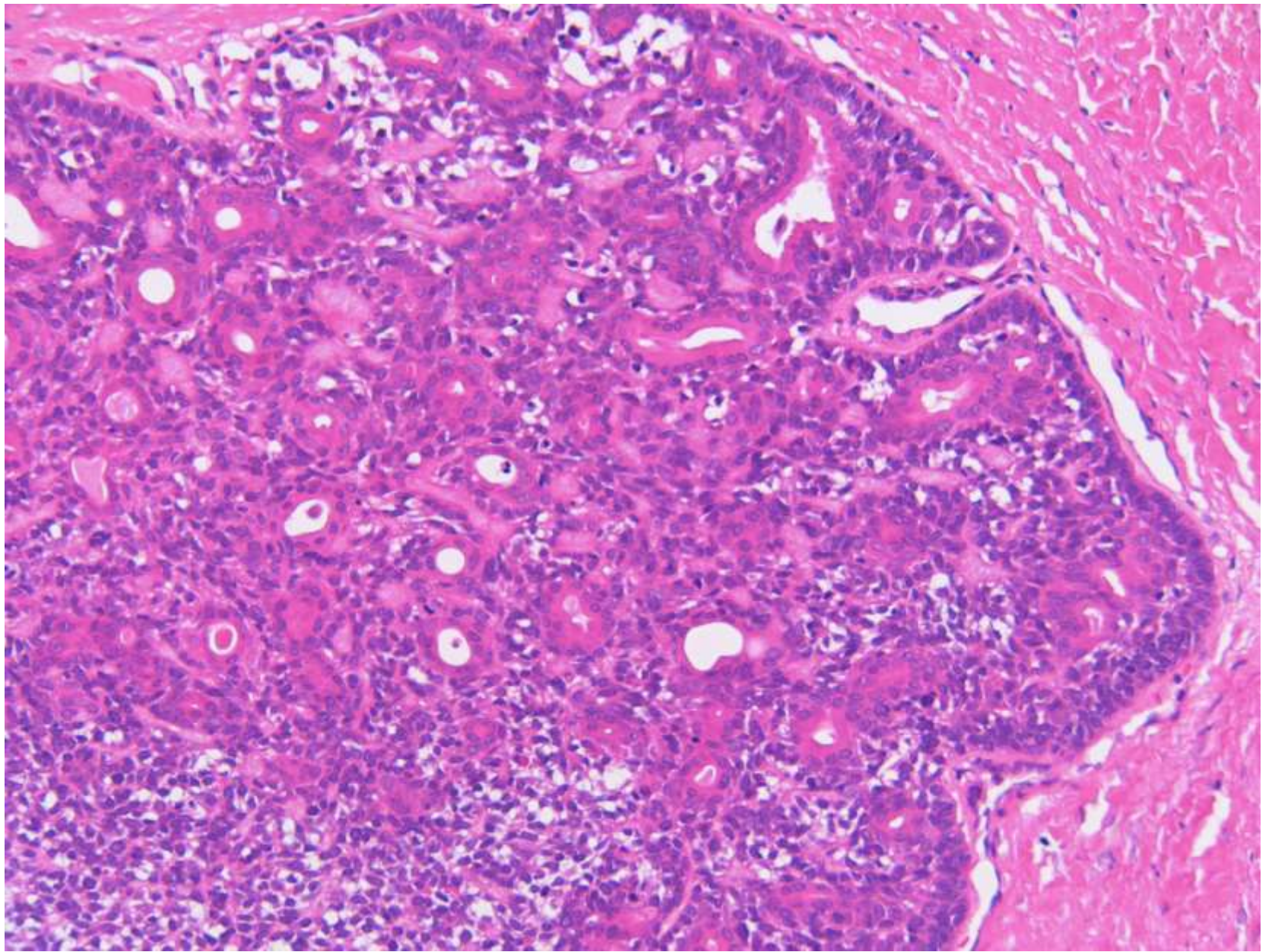




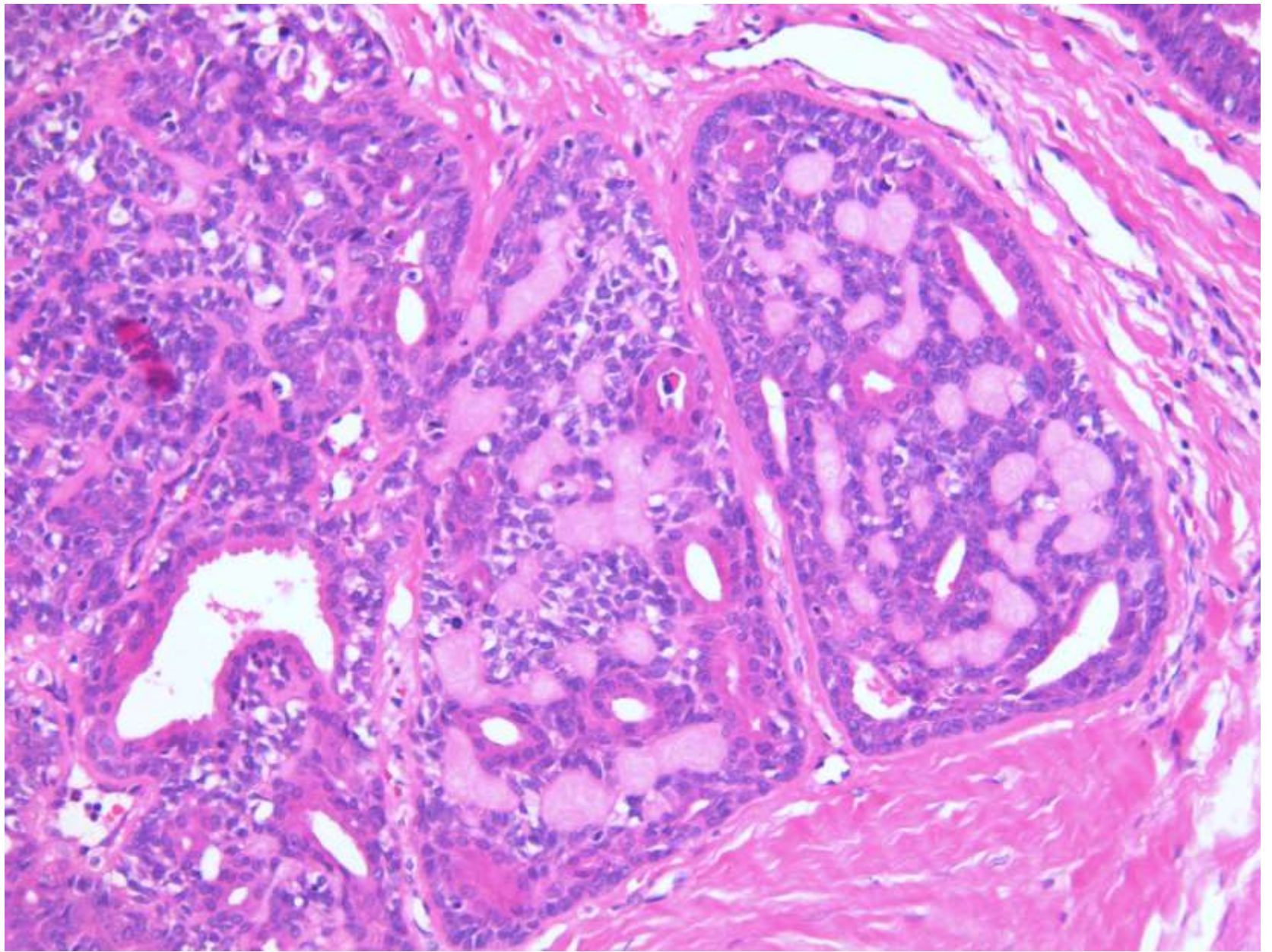
## Spot Diagnosis: Case 2













## Case 2

### Benign v's Malignant

A. Adenoma (**1 point**)

Subtype (**bonus 1 point**)

Papillary adenoma

Hidradenoma

Spiradenoma

Other

B. Adnexal Ca (**1 point**)

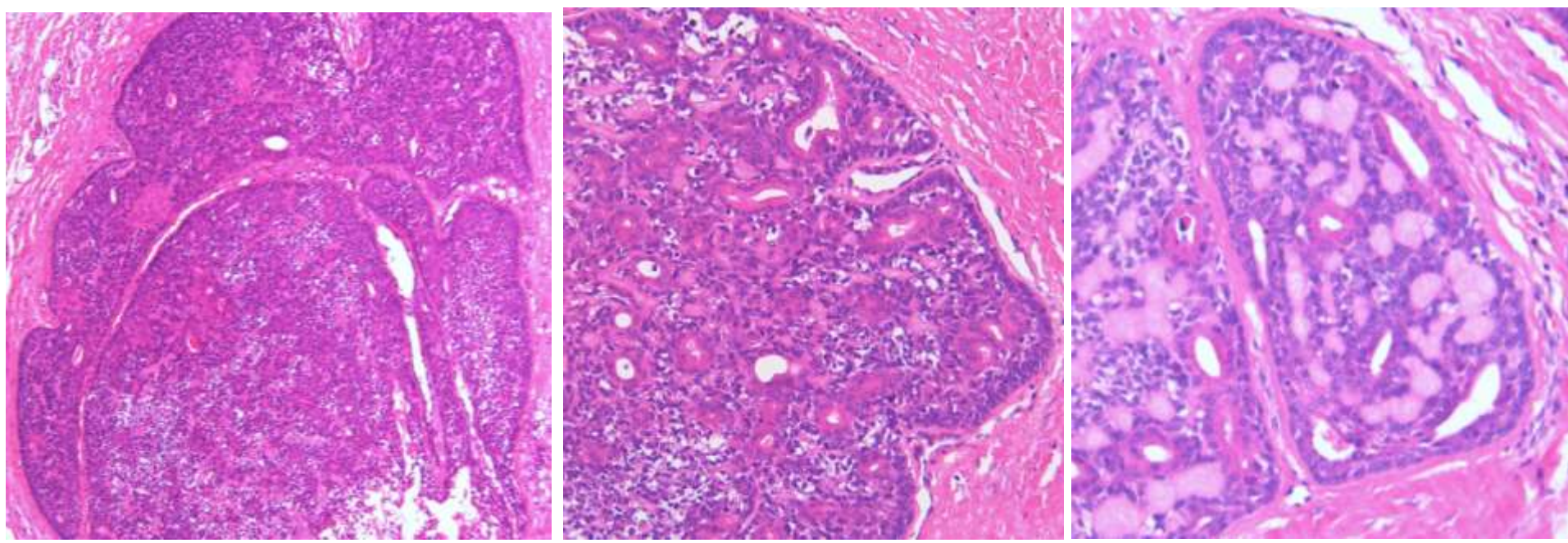
Digital papillary AdenoCa

Spiradenocarcinoma

Hidradenocarcinoma

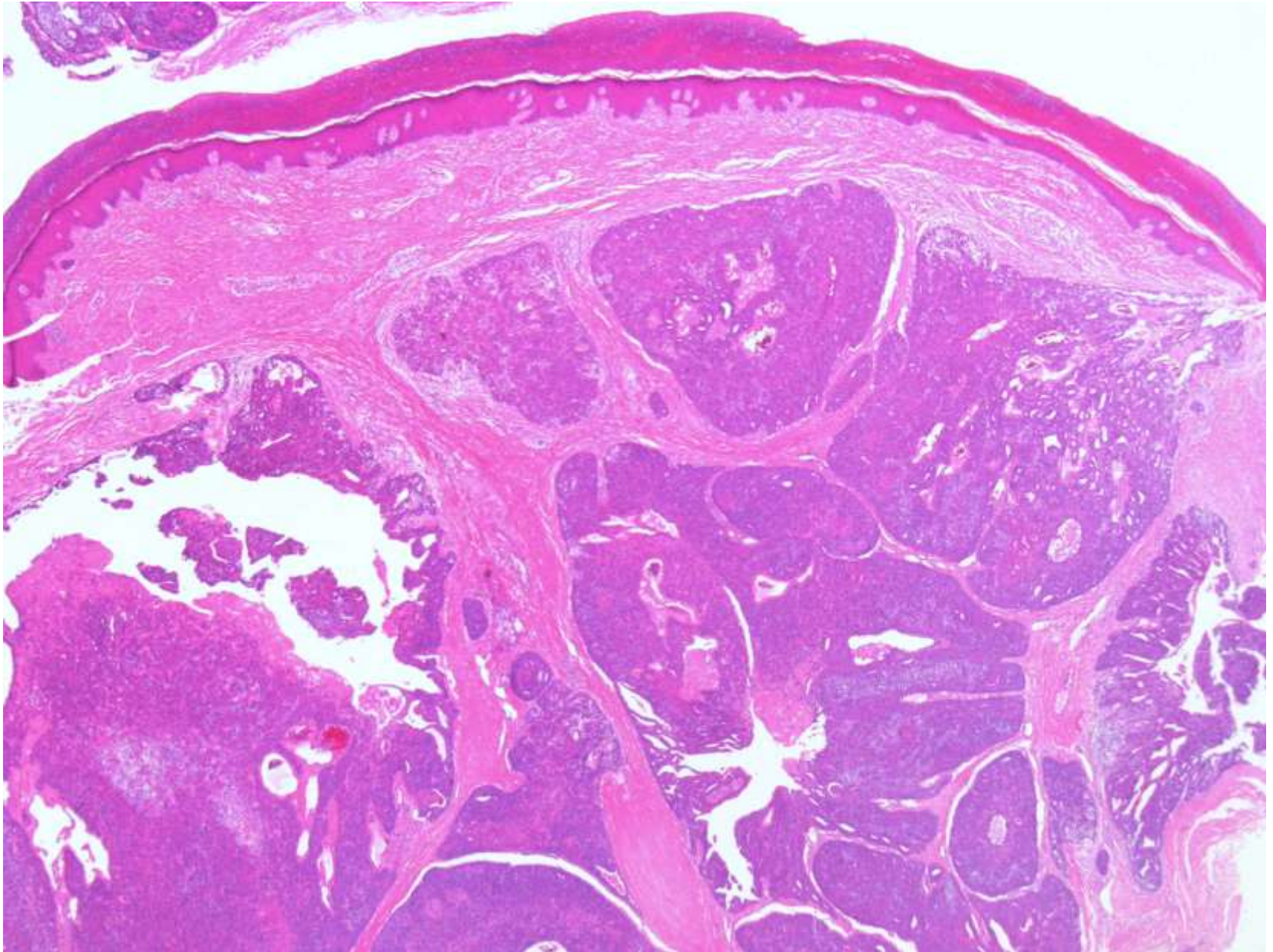
Adenoid cystic carcinoma

Other

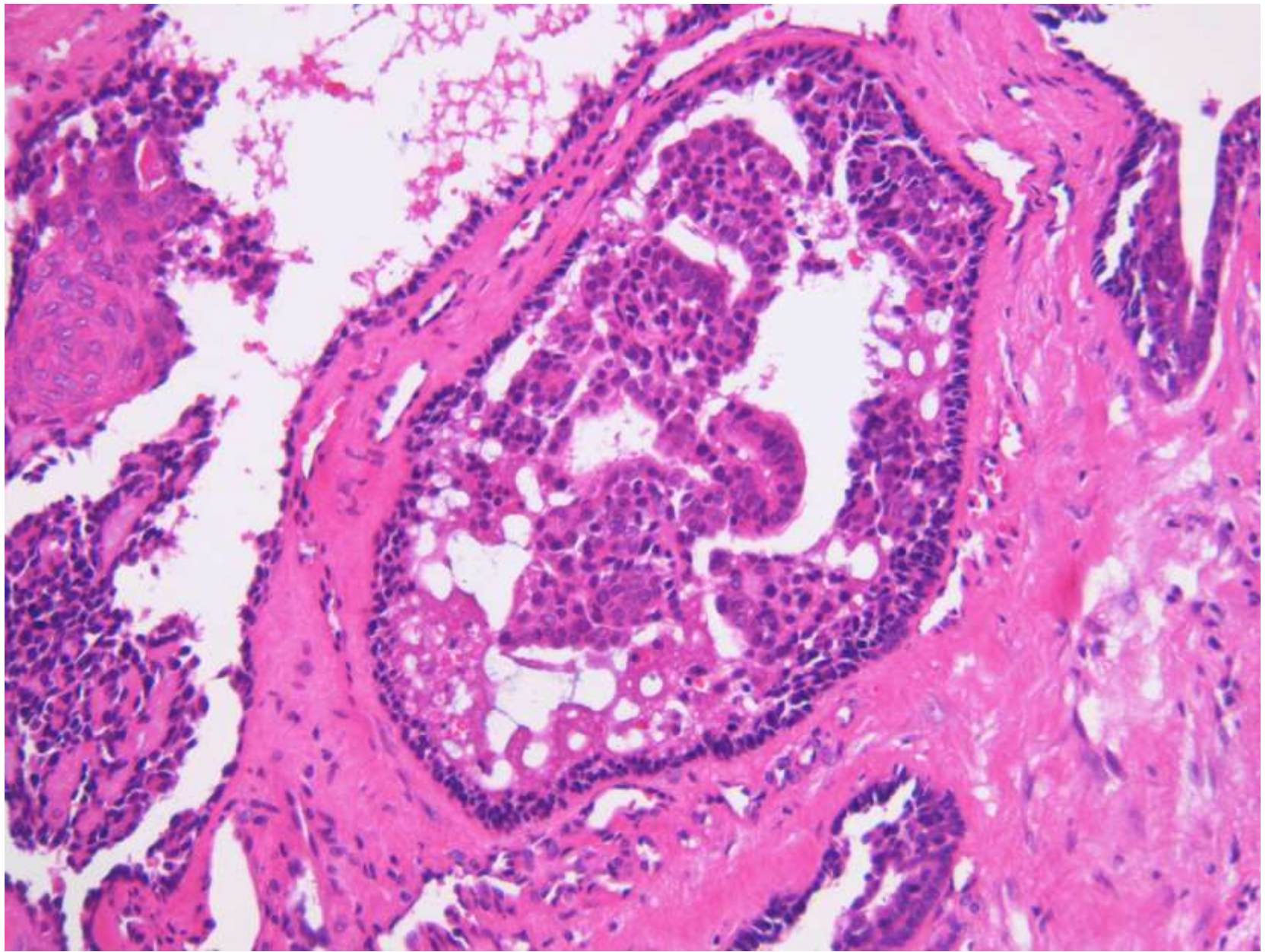




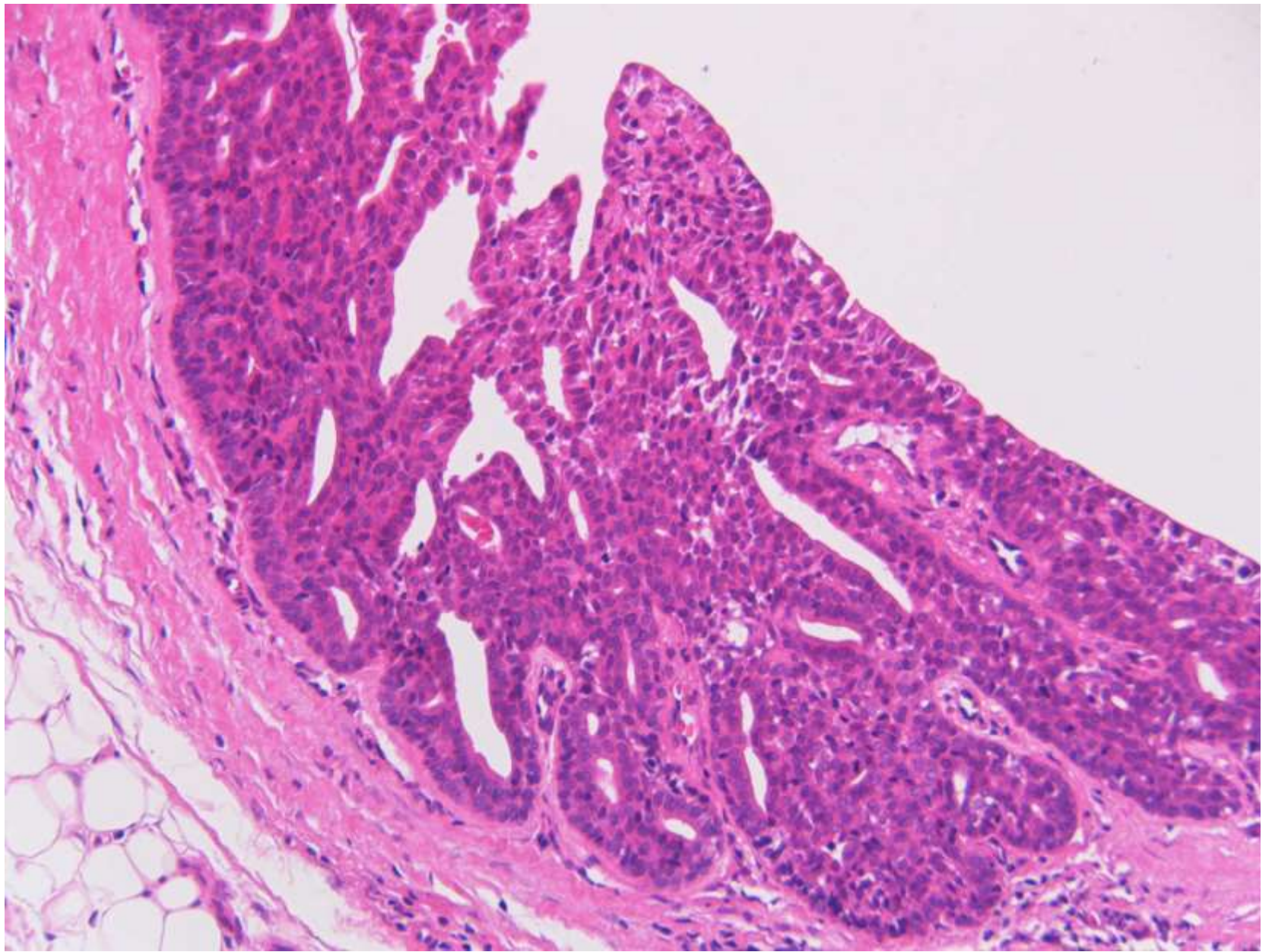
## Spot Diagnosis: Case 3



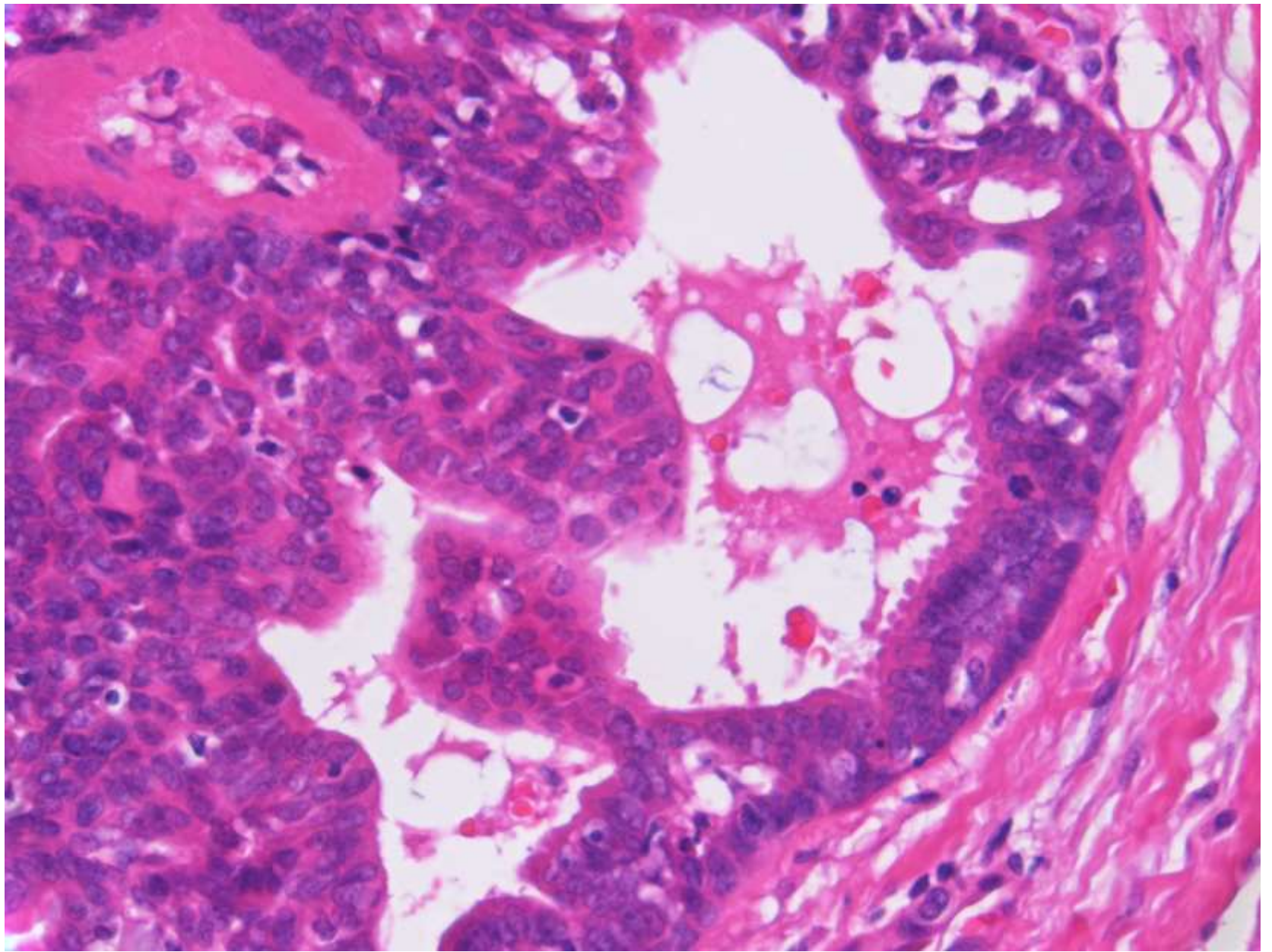












# Case 3

## Benign v's Malignant

A. Adenoma (**1 point**)

Subtype (**bonus 1 point**)

Papillary adenoma

Hidradenoma

Spiradenoma

Other

B. Adnexal Ca (**1 point**)

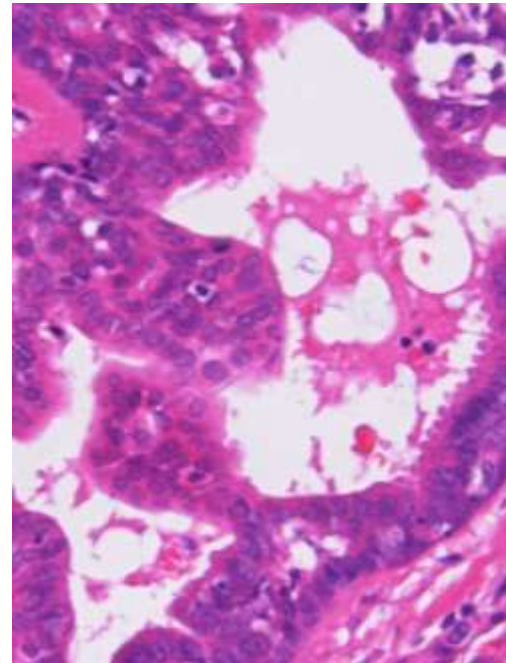
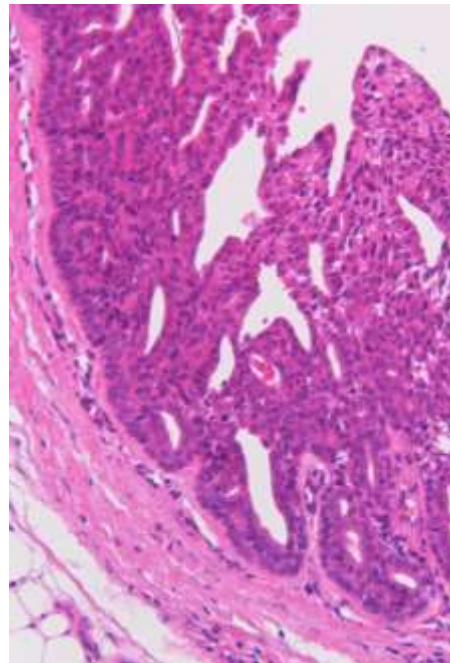
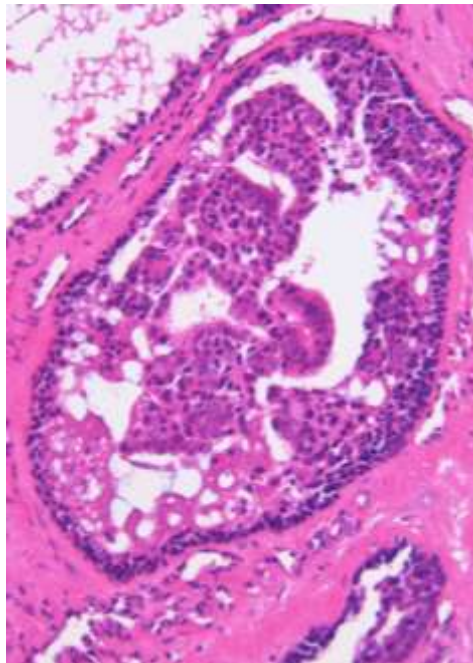
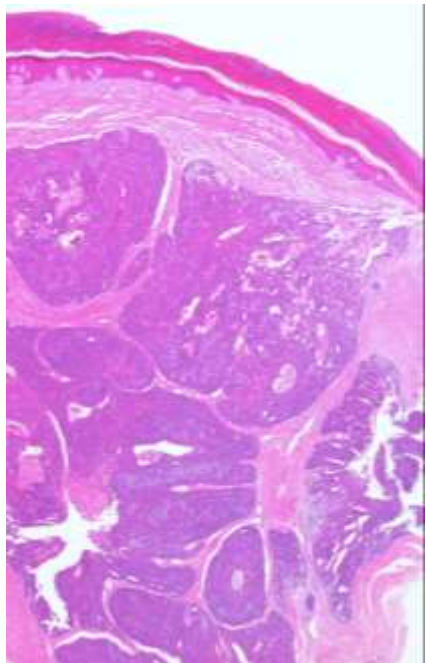
Digital papillary AdenoCa

Spiradenocarcinoma

Hidradenocarcinoma

Adenoid cystic carcinoma

Other





# Theme : Adnexal Lesions

*Self-Assessment / Spot Diagnosis Challenge*

Cases 1 to 3

## Case 1

### Benign v's Malignant

A. Adenoma (**1 point**)

Subtype (**bonus 1 point**)

Papillary adenoma

Hidradenoma

Spiradenoma

Other

B. Adnexal Ca (**1 point**)

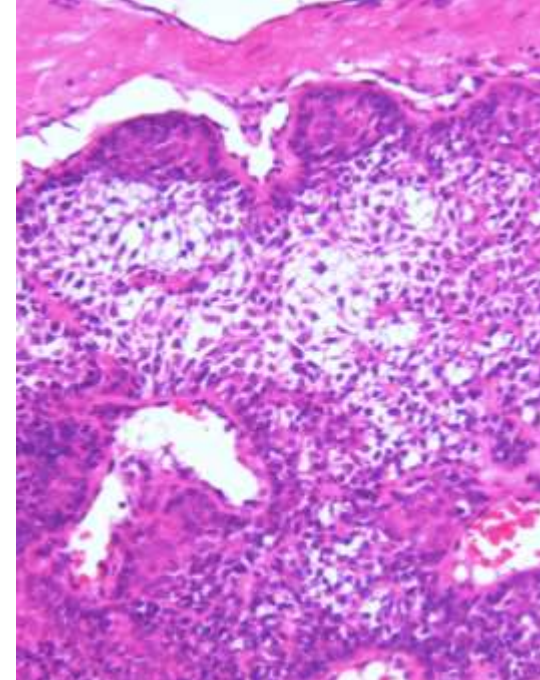
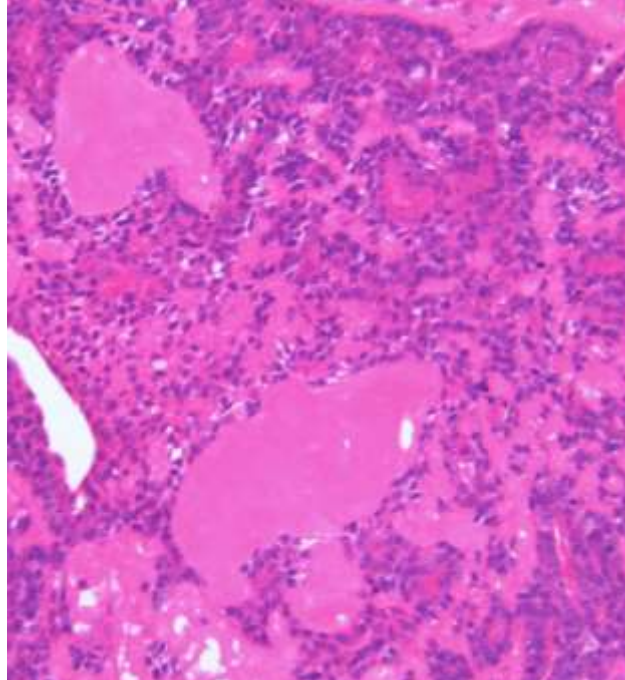
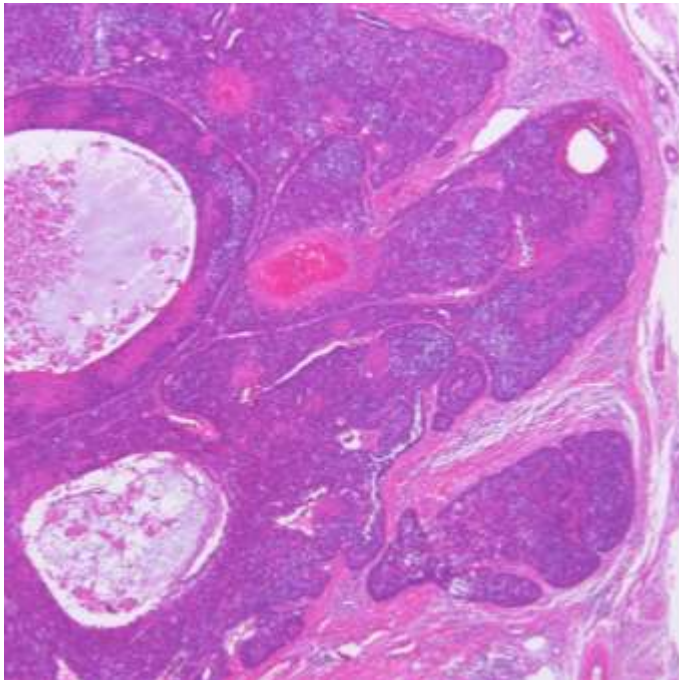
Digital papillary AdenoCa

Spiradenocarcinoma

Hidradenocarcinoma

Adenoid cystic carcinoma

Other





## Case 1

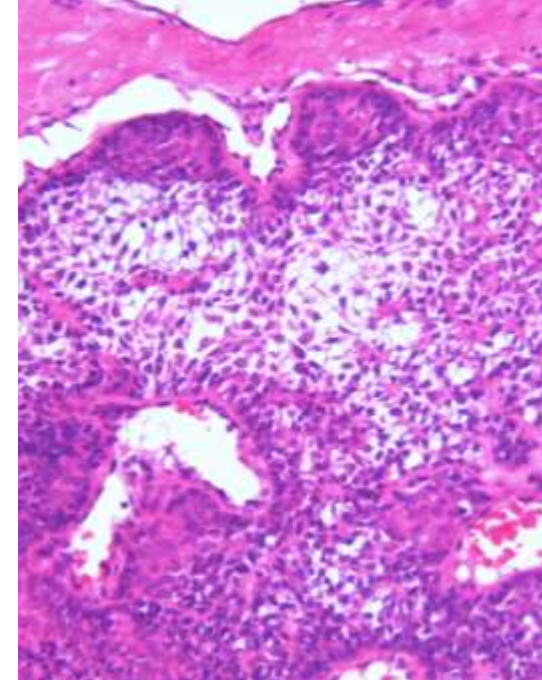
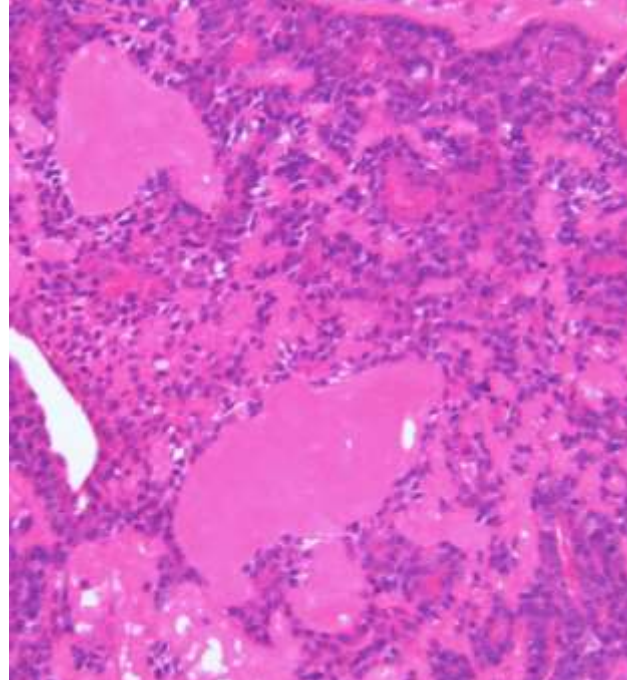
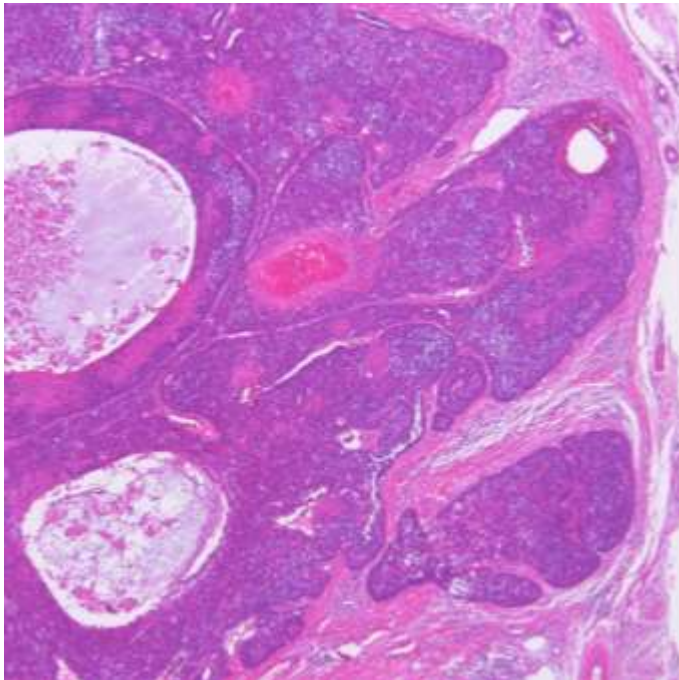
**Benign v's Malignant**  
**Adnexal Carcinoma**

**(1 point)**

**Subtype**

**Digital Papillary**  
**Adenocarcinoma**

**(1 point)**



## Case 2

Benign v's Malignant

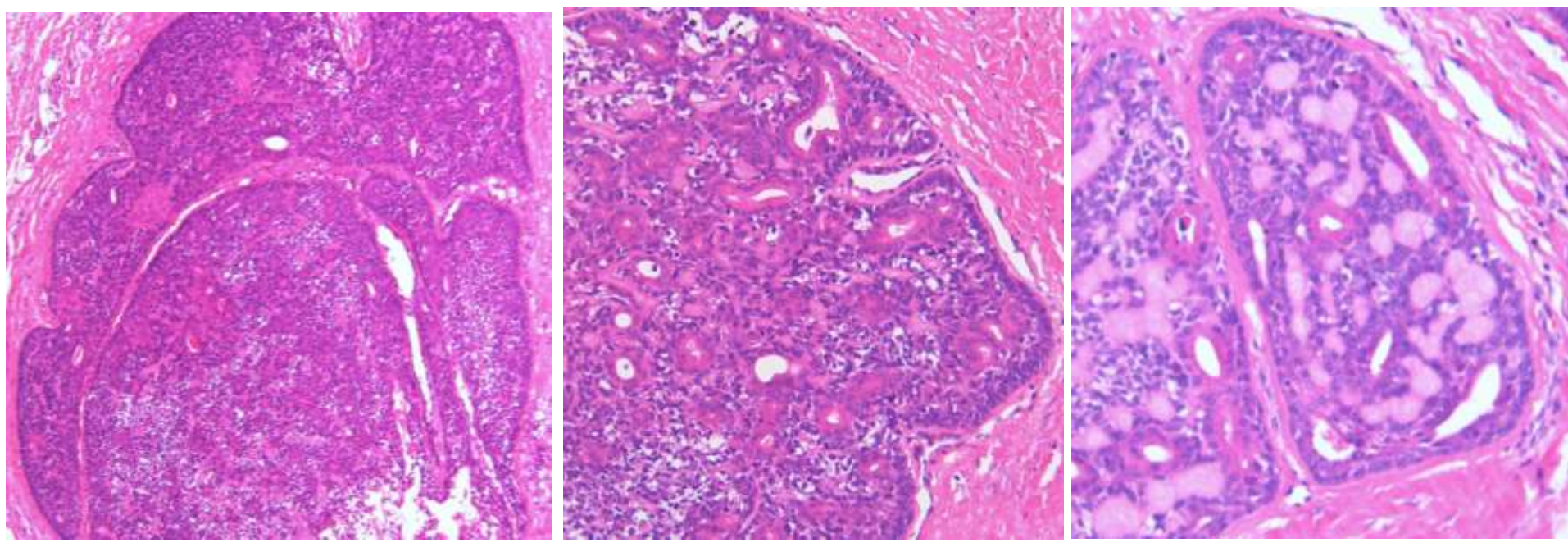
Adnexal Carcinoma

(1 point)

Subtype

Digital Papillary  
Adenocarcinoma

(1 point)





## Case 3

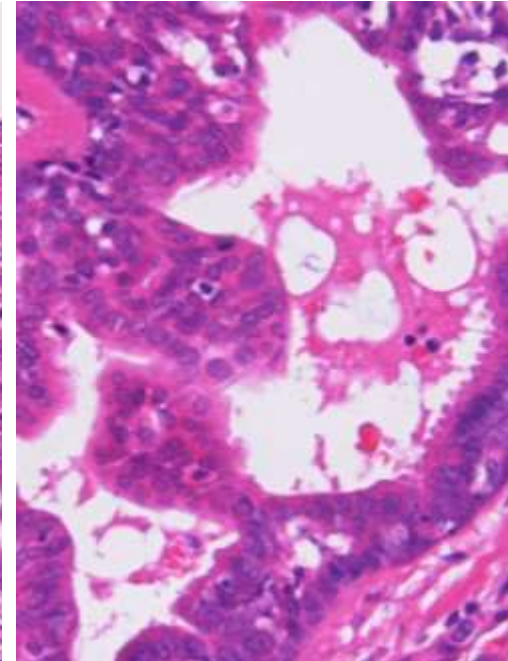
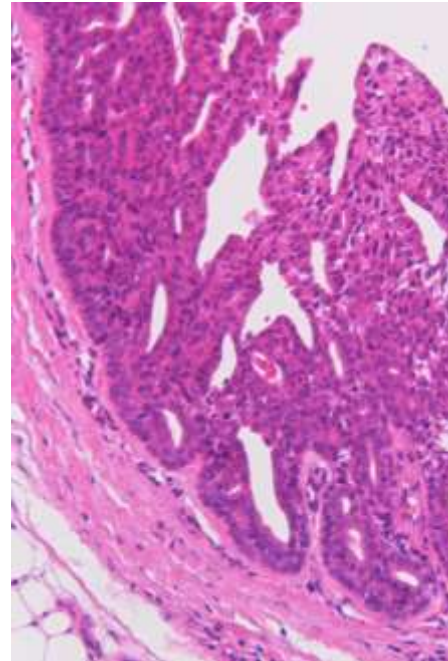
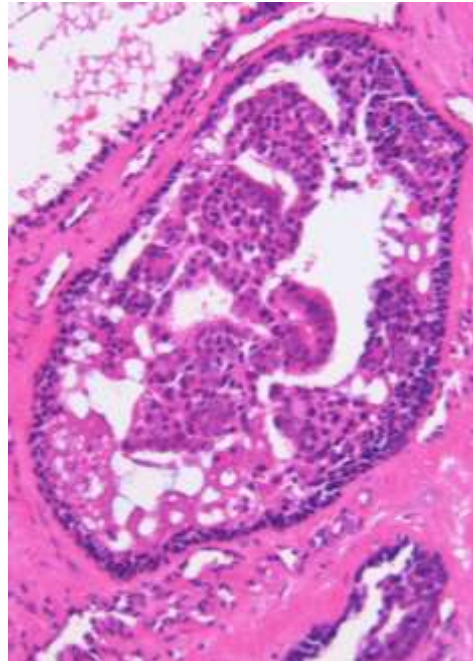
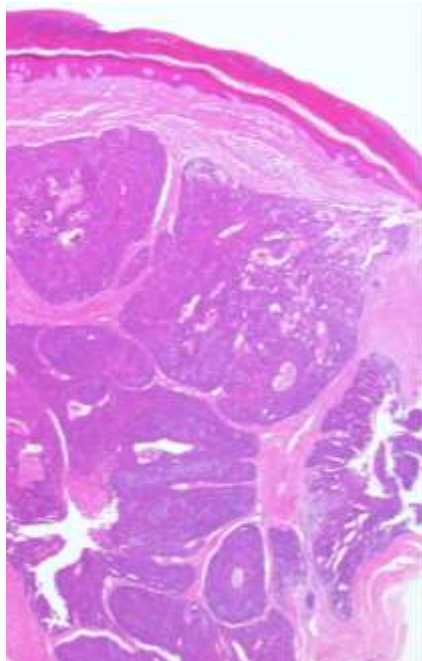
Benign v's Malignant  
Adnexal Carcinoma

(1 point)

Subtype

Digital Papillary  
Adenocarcinoma

(1 point)





# Case 1-3: Digital papillary adenocarcinoma

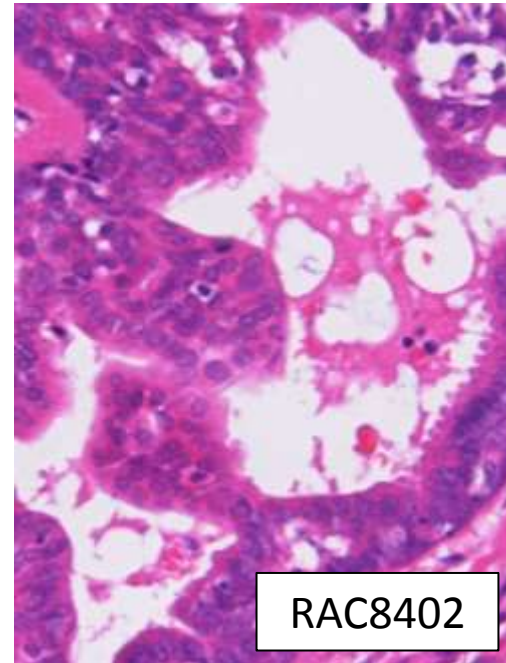
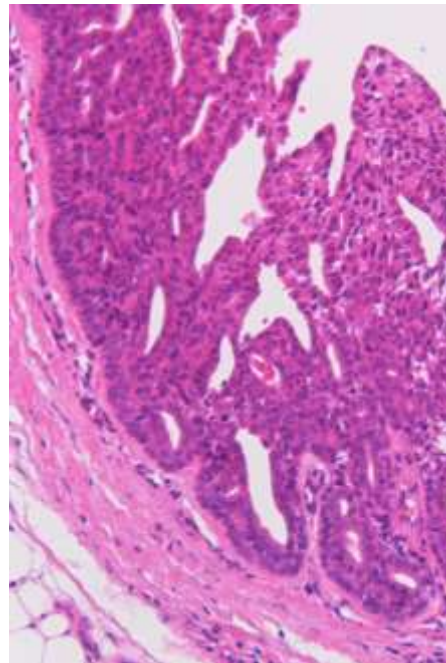
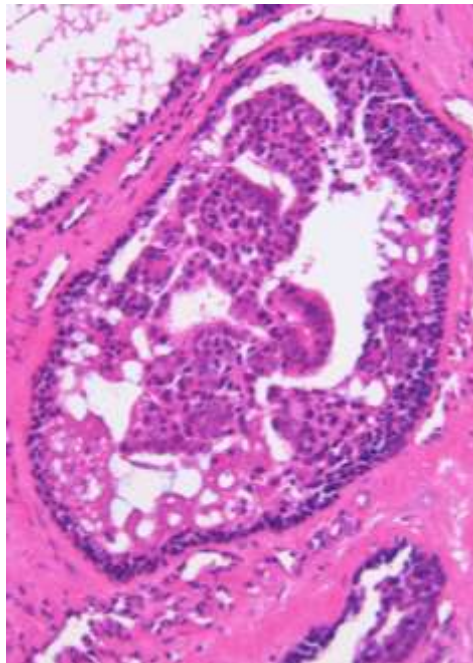
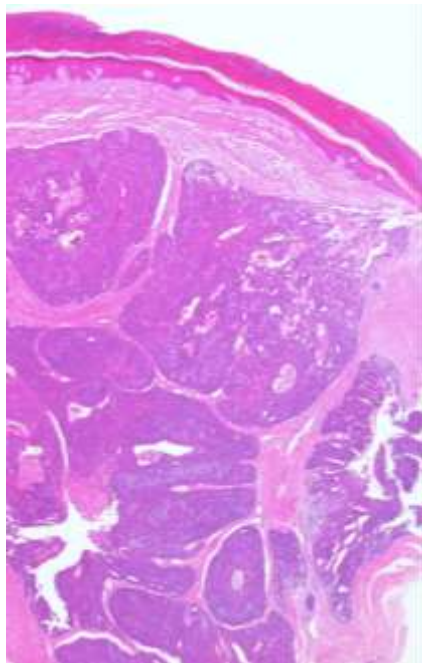
**2005: M47. Right Thumb. “Lesion” excised by Orthopaedic surgeon**

Macro: White fibrous nodule 15 x 14 x 10mm

Micro: *“...consistent with a hidradenoma (acrospiroma), best considered atypical in view of mitotic activity. The lesion is incompletely excised...Suggest complete excision.”*

## SUPPLEMENTARY REPORT

*“I have discussed this case also with my colleagues. ...the behaviour is uncertain. Metastatic deposits may occur with atypical hidradenomas. This case should be discussed at MDM.”*



RAC8402

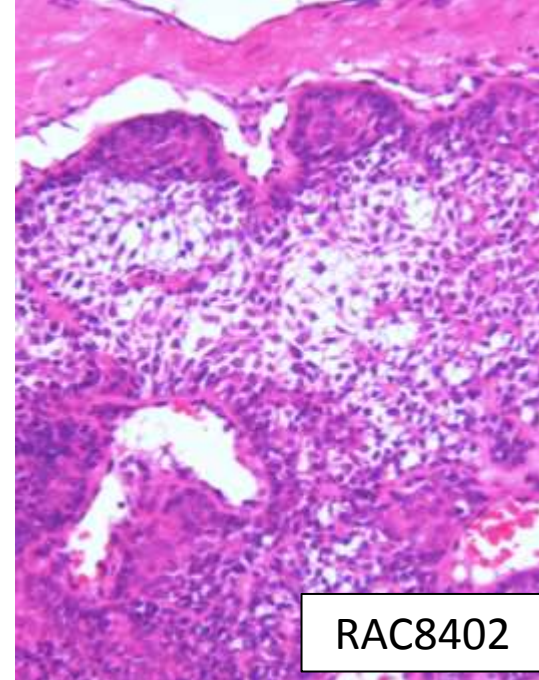
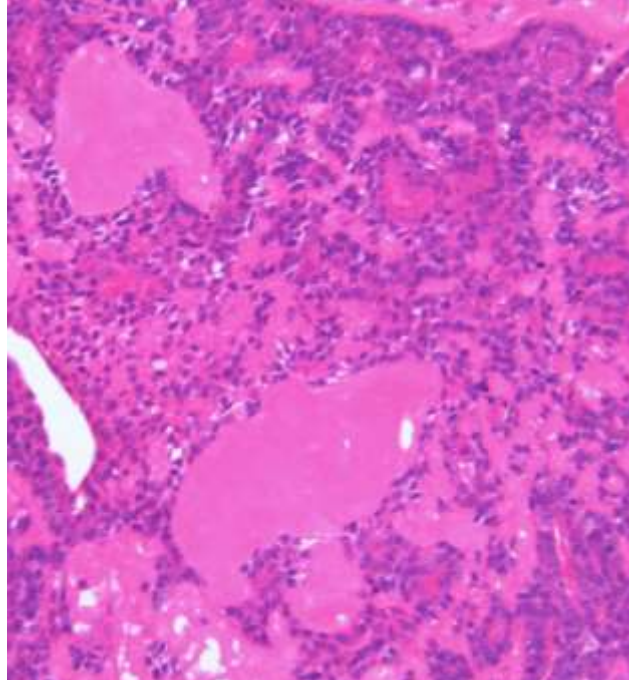
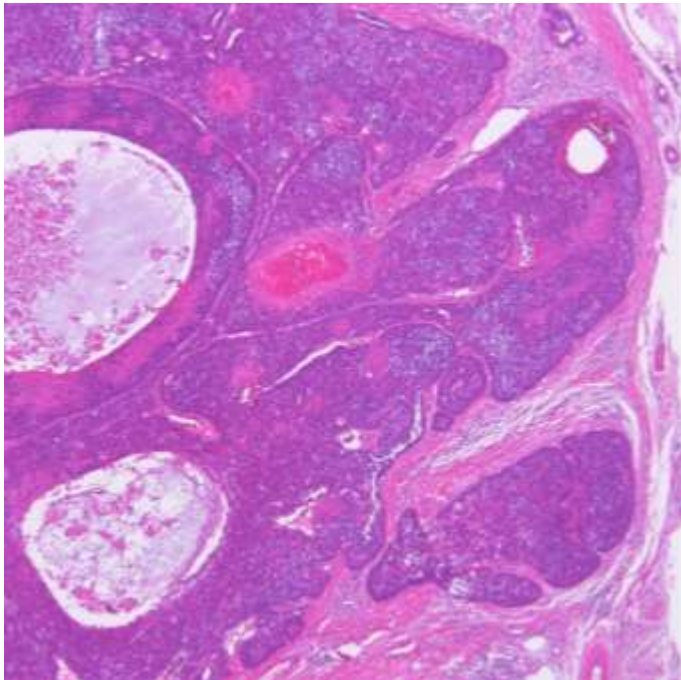
# Case 1-3: Digital papillary adenocarcinoma

## REVIEW AT EXTERNAL BONE / SOFT TISSUE CENTRE

“...eccrine differentiation...cystic spaces...tubular formation...clear cells...focal squamous differentiation...as small focus of increased mitotic activity ( $>10/10$  hpf)...no necrosis...NPVI...

*...features c/w atypical nodular hidradenoma.*

*...it should be noted the criteria for malignant nodular hidradenoma are not clearly defined”*





# Case 1-3: Digital papillary adenocarcinoma

**2019: M60.** Known Stage 4 non small cell lung cancer Feb 2018.

Cut toenail & inflamed right big toe developed growing vascular lesion, bleeding lesion. **[No mention of “atypical hidradenoma Rt Thumb”]**

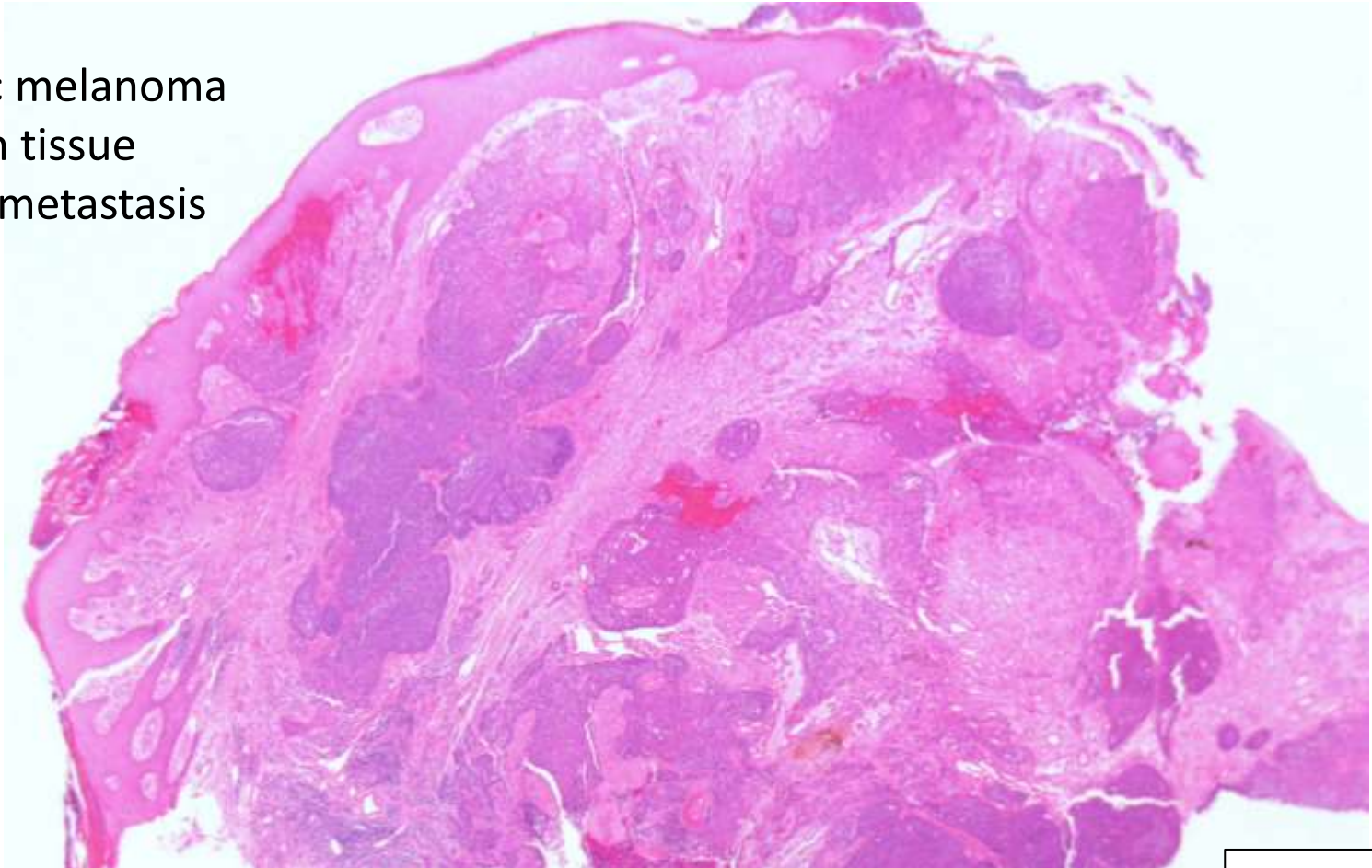
DDx:

?PG,

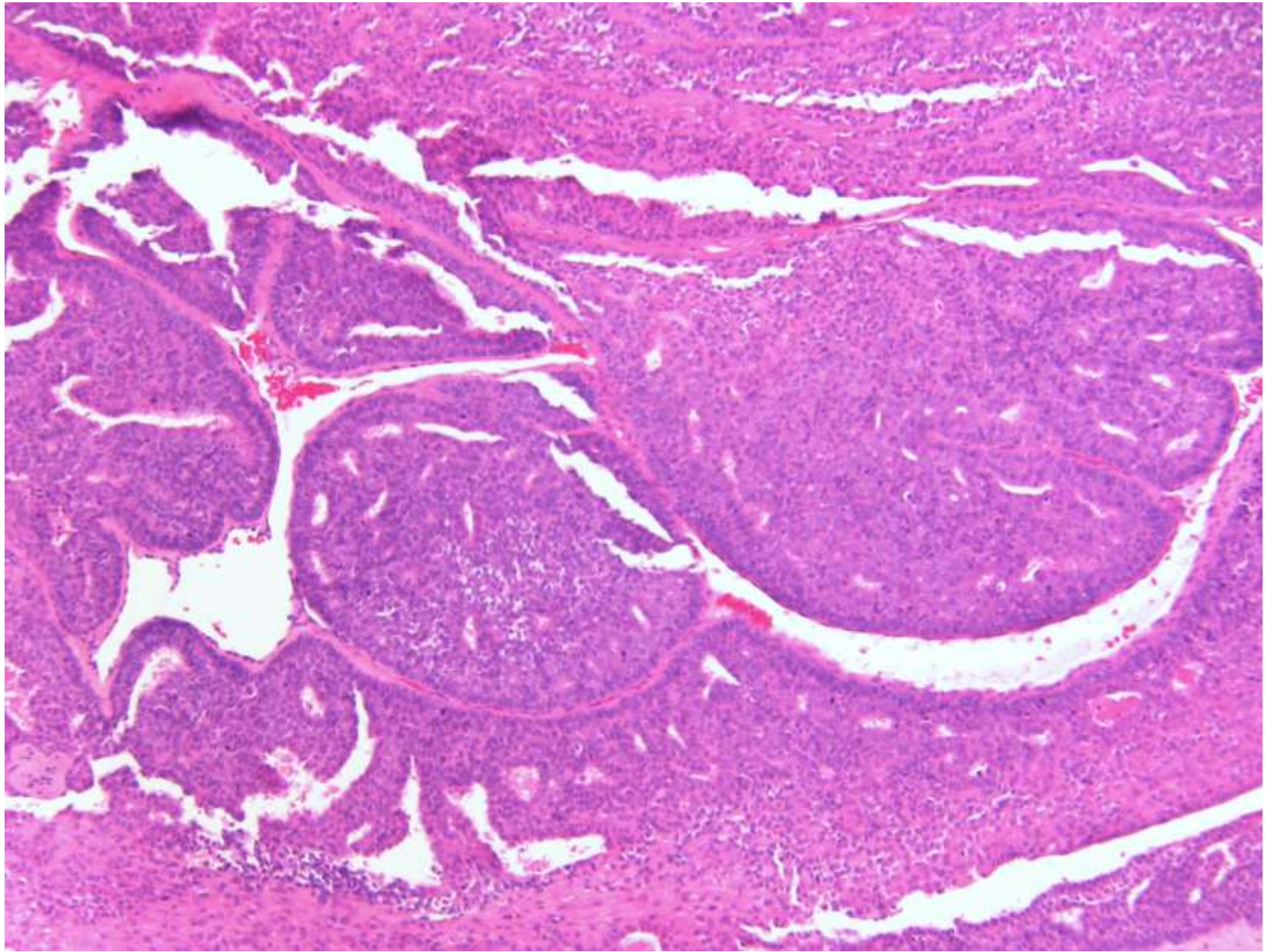
?amelanotic melanoma

?granulation tissue

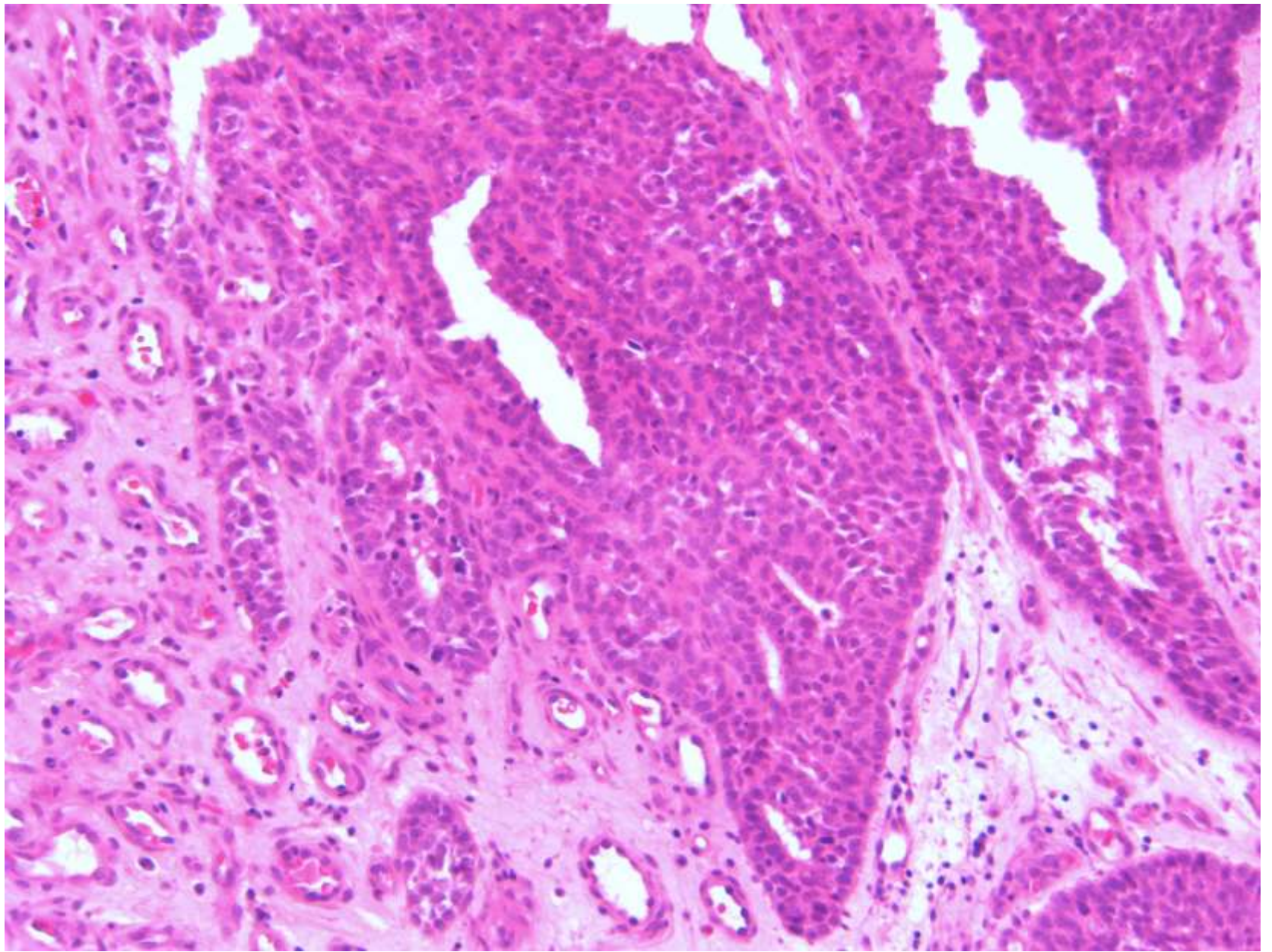
?secondary metastasis



RAC8402D









CEA

p63

BerEP4

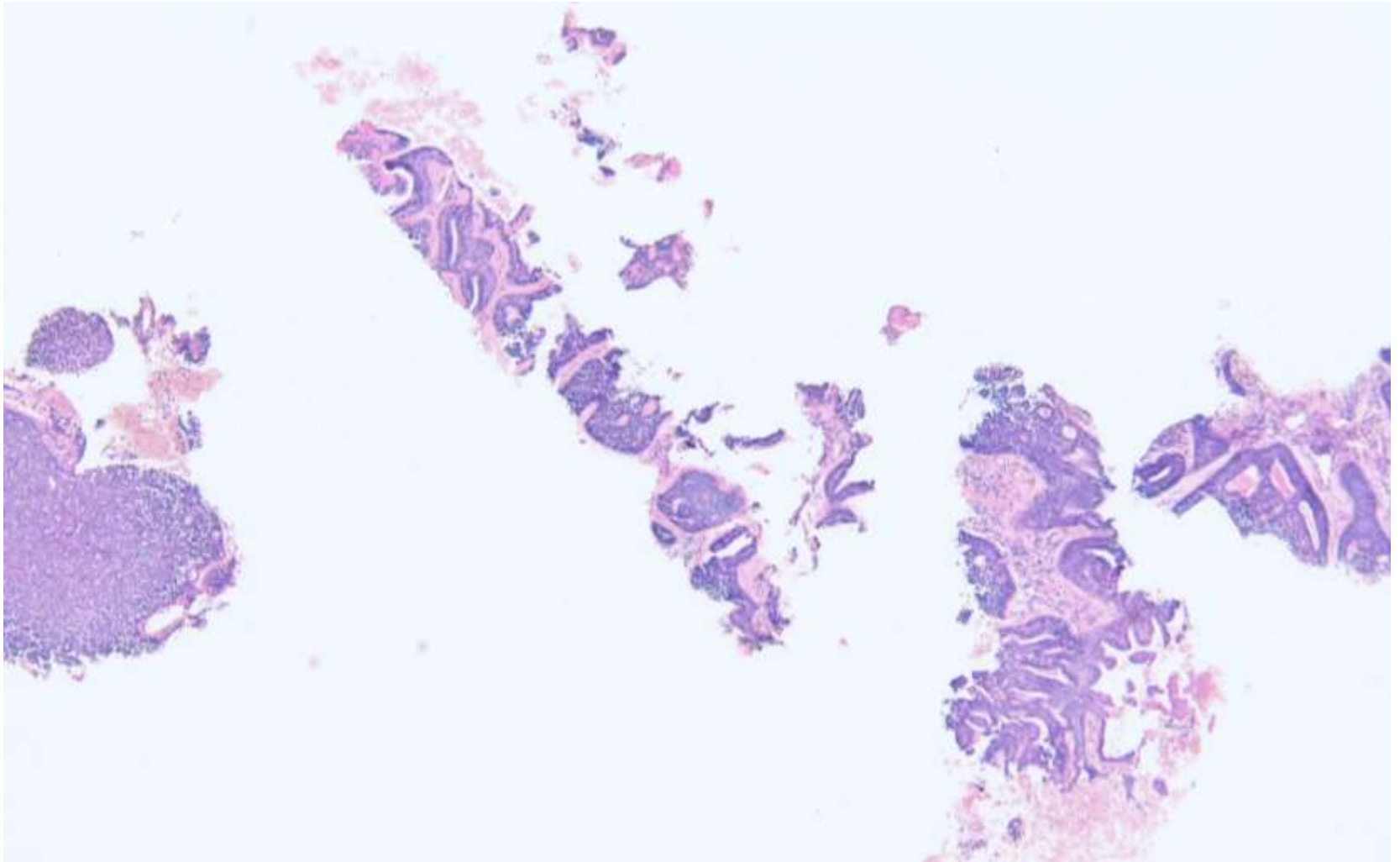
*Case with colleague – shown to RAC  
...in keeping with digital papillary  
adenocarcinoma. Excision  
recommended*

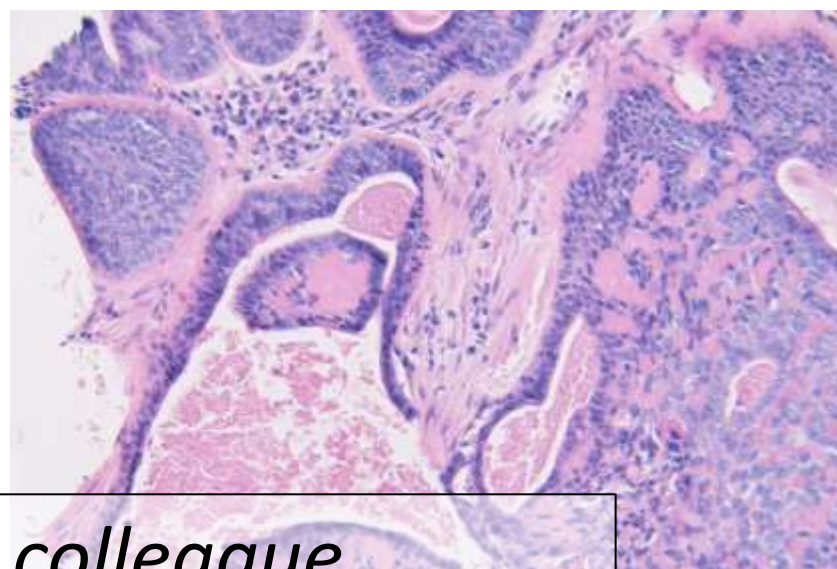
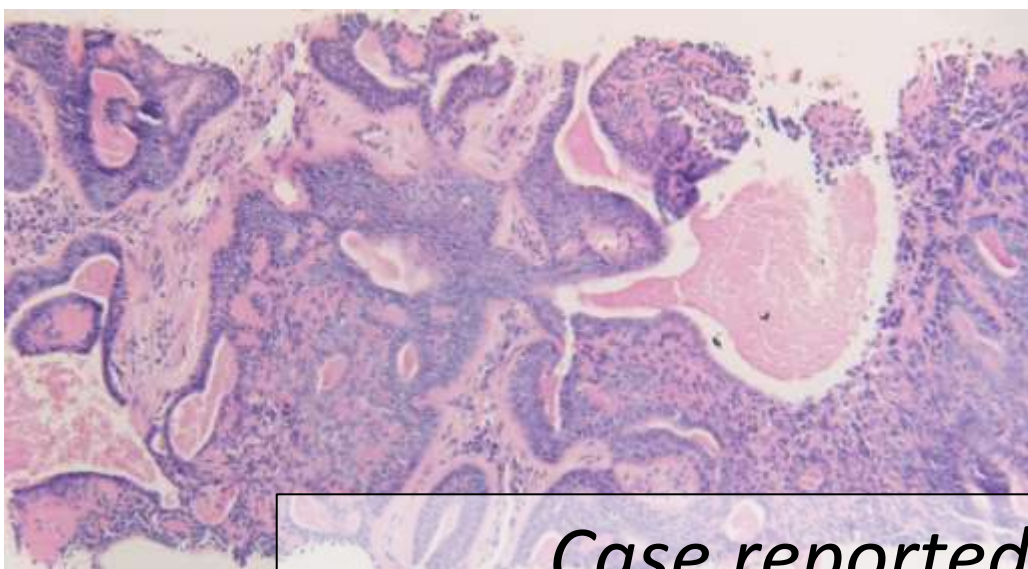


# Case 1-3: Digital papillary adenocarcinoma

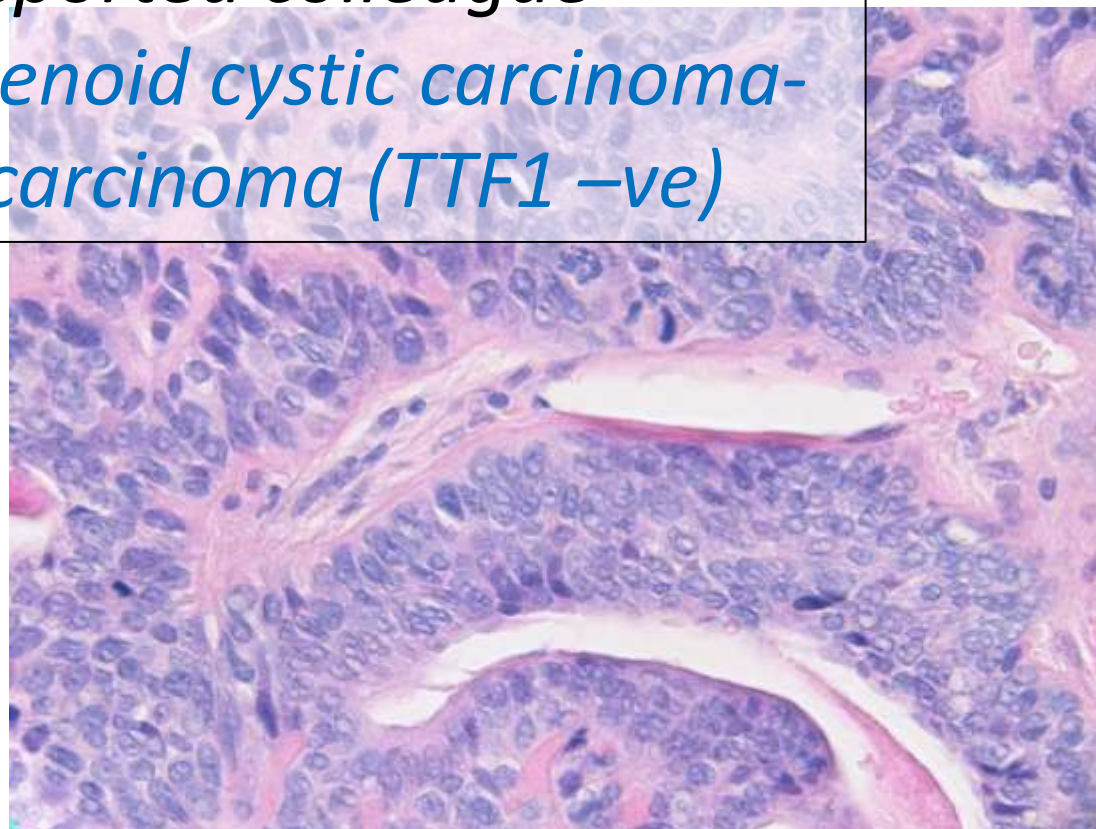
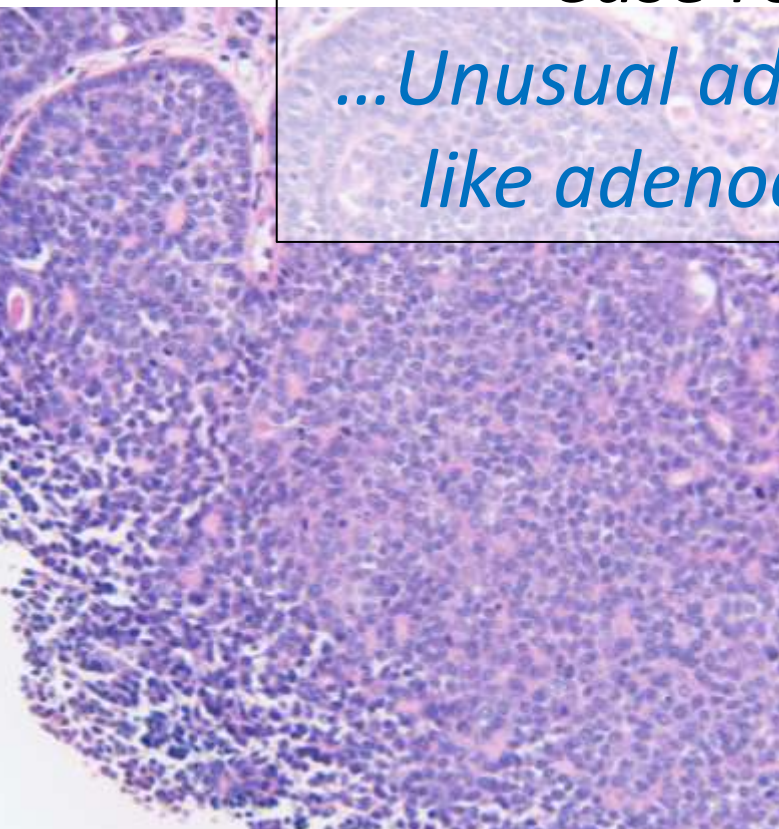
2019: MDM Discussion – Known stage 4 lung adenocarcinoma

2017: 4 “coin” lesions ?hamartomas of lung



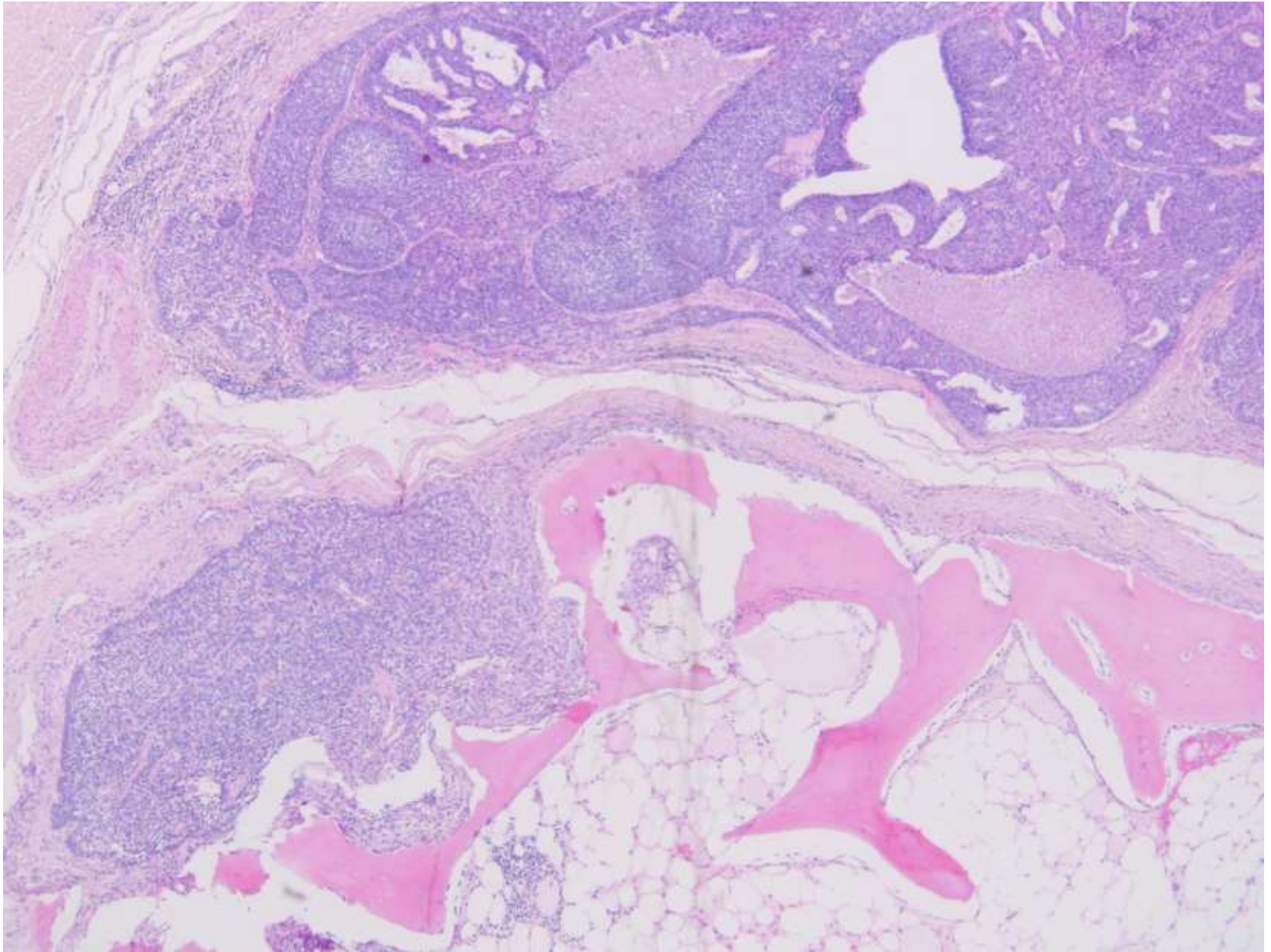


*Case reported colleague*  
*...Unusual adenoid cystic carcinoma-  
like adenocarcinoma (TTF1 -ve)*

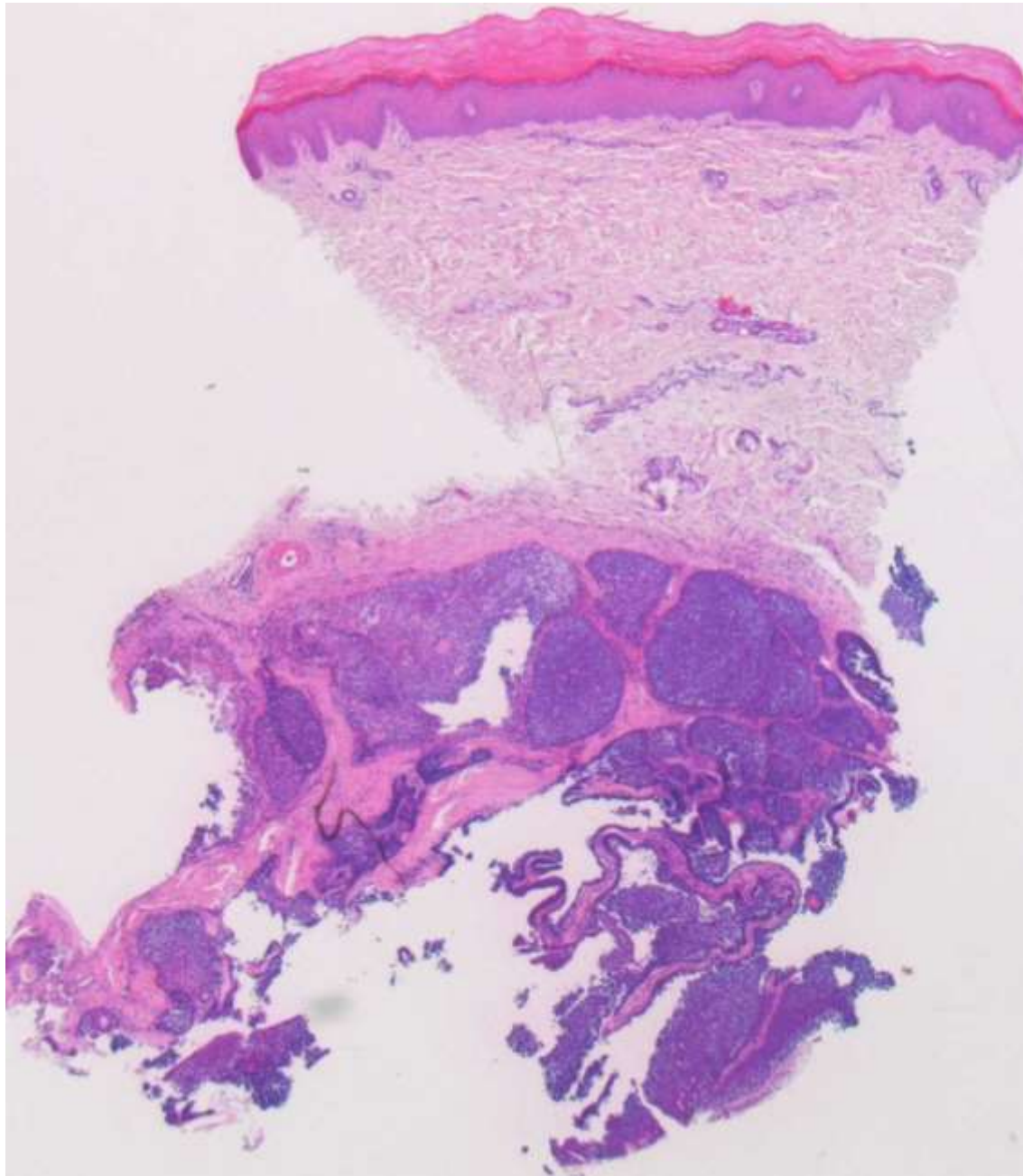




## 2019: Metastatic DPA: Bone invasion right great toe



**2019: Metastasis to right 2<sup>nd</sup> toe**





# Case 1-3: Digital papillary adenocarcinoma

2005: “Atypical Hidradenoma Rt Thumb”

2017: “... Unusual adenoid cystic carcinoma-like tumour lung

Treated for lung adenocarcinoma Stage 4

2019: Digital papillary adenocarcinoma Right big toe

2019: Digital papillary adenocarcinoma Right 2<sup>nd</sup> toe

Review of the whole case: Digital papillary adenocarcinoma of the Right thumb (2005) with late lung (2017) and subsequent aggressive toe metastases (2019)

2020: 2005 & 2019 representative blocks sent to Thomas Weisner for HPV42 testing (pending)

*Highly sensitive assay developed for virus detection in FFPE tissue*

*HPV42 contributes to pathogenesis of DPA and might be a therapeutic target*

*HPV42 can be re-classified as a high risk based on intergration of 7 crucial aminoacids of the E6 / E6AP / p53 complex*

# Case 1-3: Digital papillary adenocarcinoma (DPA)

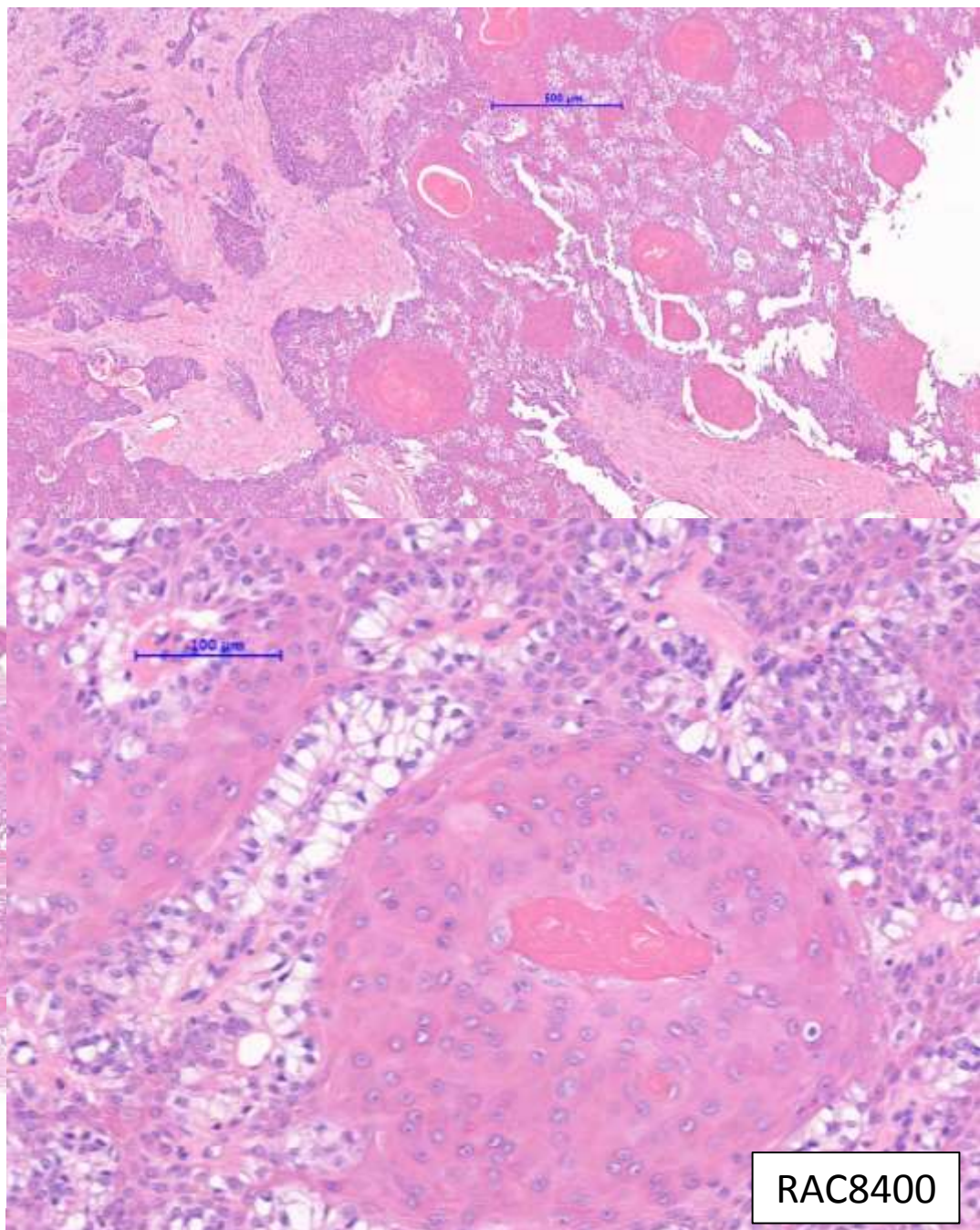
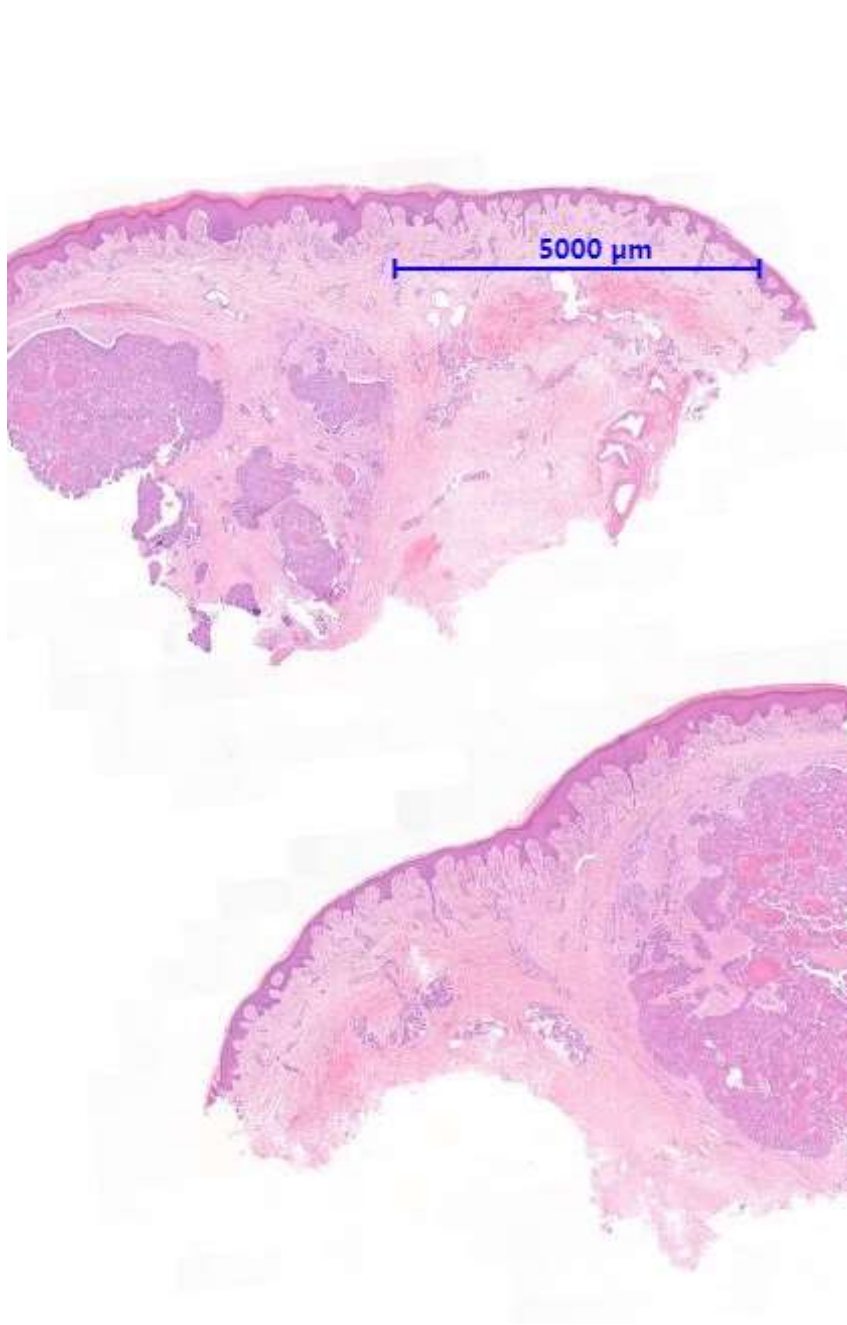
## **Learning Points**

1. *Beware the “adenoma” of the digit*
2. *Show case to dermatopathologist for second opinion*
3. *Actively ask about past medical history at MDM discussion and act on the findings*
4. *DPA have protean histological features perhaps better named “polymorphous” adenocarcinoma of the digit*
5. *These are generally “indolent” lesions and quite unpredictable but the term “aggressive” should not be applied*
6. *Recent finding: HPV42 (low risk in the setting of squamous epithelium) is high risk in the setting of the sweat duct/gland and may be specific for DPA*

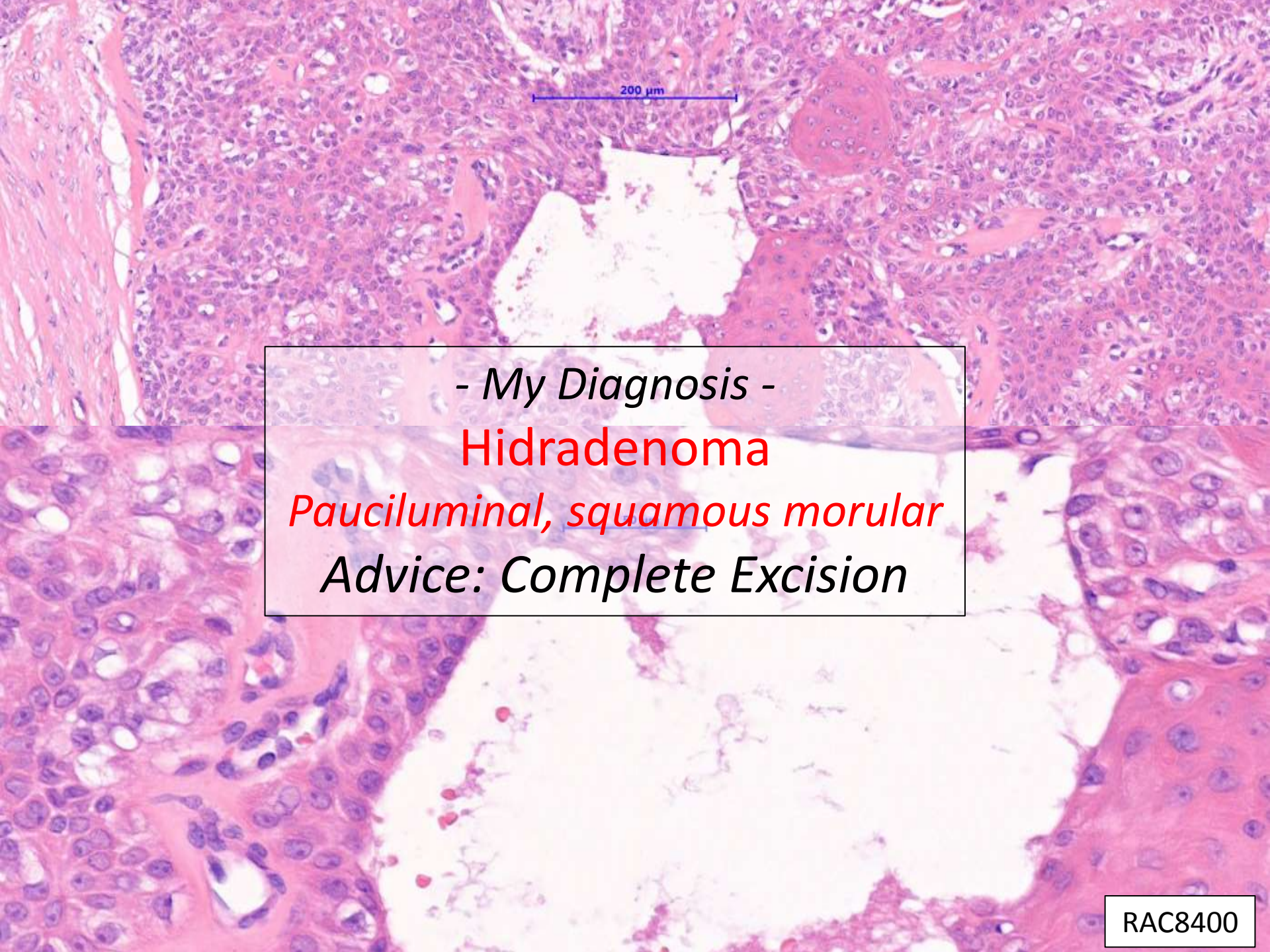


Next Case

F82. Left Ankle ?SCC (Digital referral c/o Dr Saleem Taibjee)







- *My Diagnosis -*

**Hidradenoma**

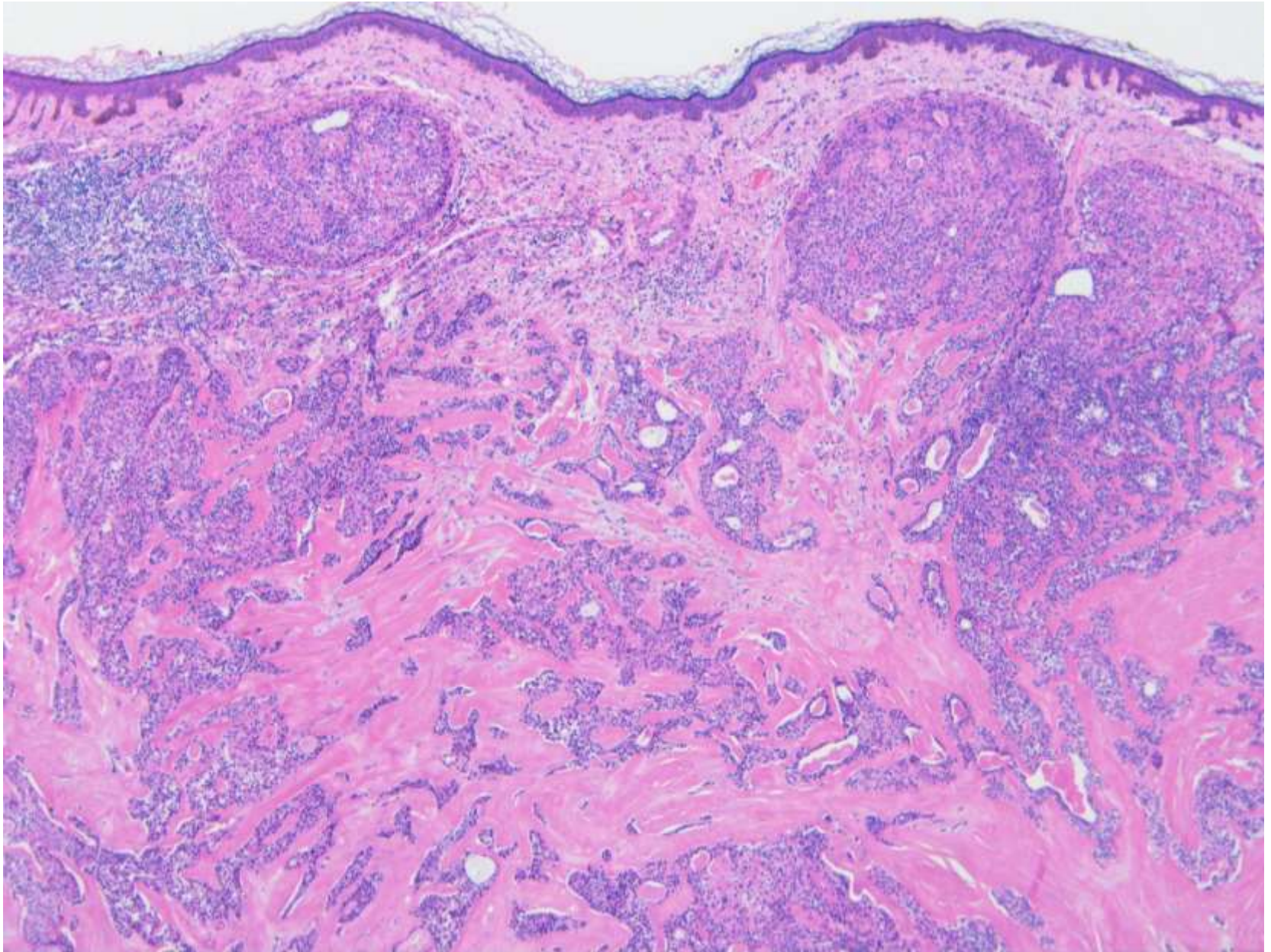
*Pauciluminal, squamous morular*

*Advice: Complete Excision*

Next Case



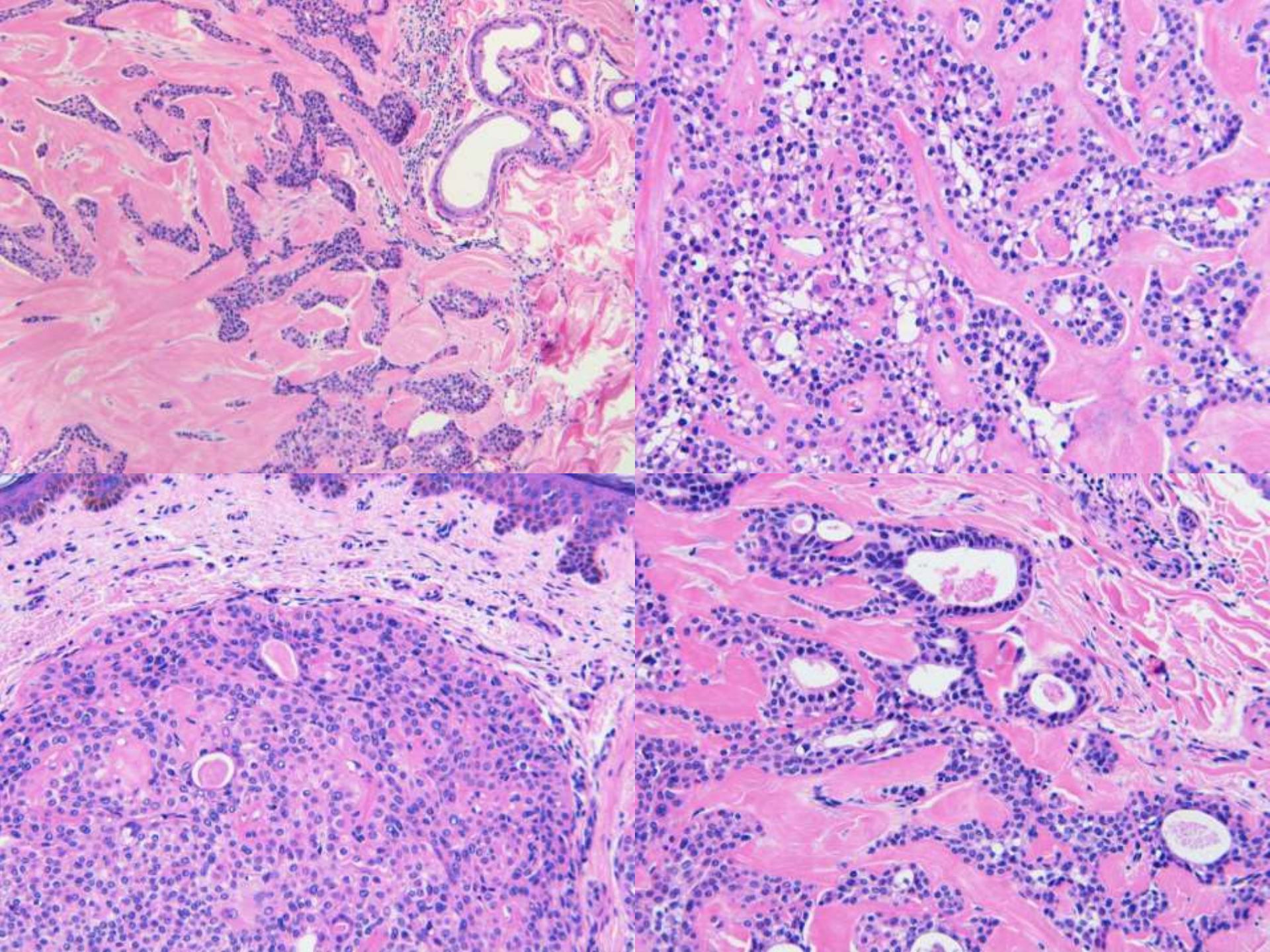
F67. Left foot. 6/12, increased in size



Referred Case c/o Saleem Taibjee

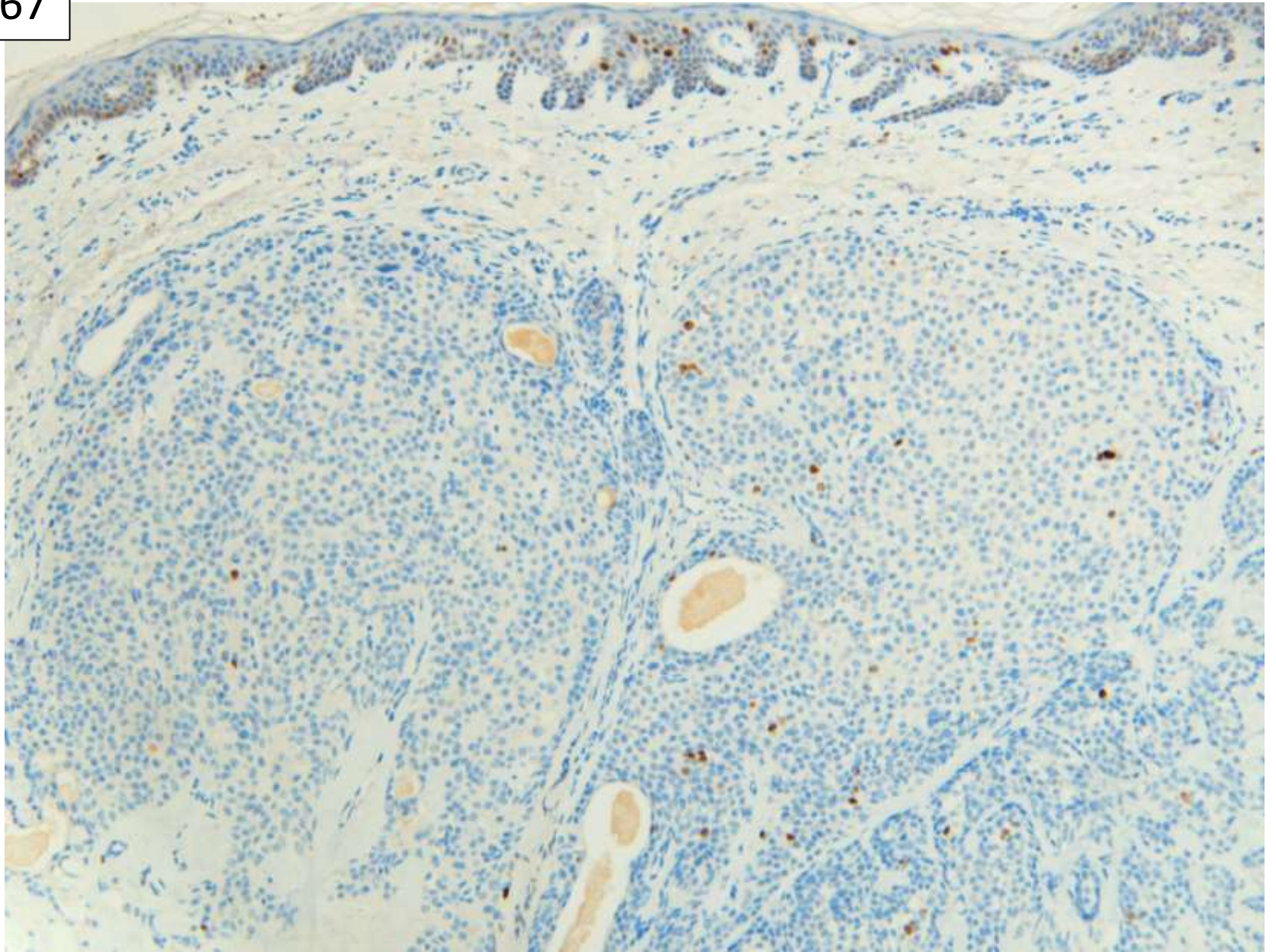
RAC7105







Ki67



s/b National Expert: Hybrid follicular, glandular elements, very bland, concerning feature is the "infiltrative" growth pattern with extension on to margin. On this basis **low grade malignancy cannot entirely be excluded**, and despite the lack of likely metastatic potential, local recurrence is a possibility as indicated in the original report. Conservative re-excision may be advisable.

RAC: Rounded pushing overall profile. Rather infiltrative appearing pale and clear cells central squamous morular / squamous cuticular ductal differentiation but lumina sparse (strongly by CEA and to a lesser extent EMA) with amorphous secretions strongly +ve for PAS and EMA). BerEP4 patchy moderate ++50%, CK5 diffuse, Ki67 5-10%. Amorphous stroma typical of hidradenoma and lesion extensively sclerotic centrally. Sparse calcification some rounded psammoma like but larger. Focally myofibroblastic spindle cells stroma slightly myxoid but not typically chondroid. In my opinion **a typical example of a sclerosing variant of hidradenoma**. No worrying features for malignancy but local recurrence is a possibility.

No mitoses. No cellular pleomorphism.

Dear Richard,

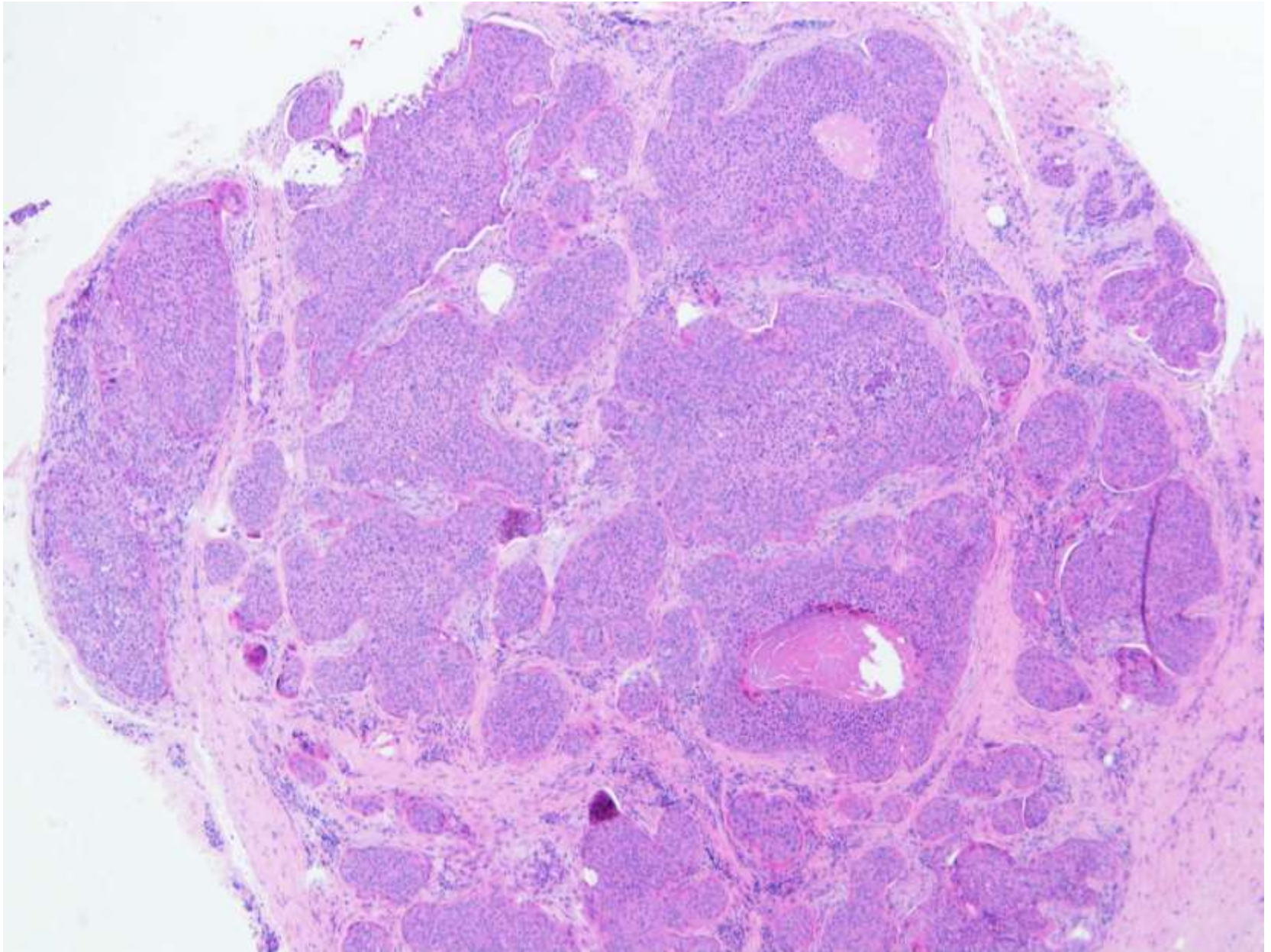
Thank you for sending me this interesting case. I completely agree with you that this is a **hidradenoma**. The sclerotic appearances are likely due to the location/ chronic repeated trauma. ***I think it is a benign lesion but after having seen cases of benign metastatic hidradenoma, I would recommend complete excision.***

Best regards, Dmitry Kazakov

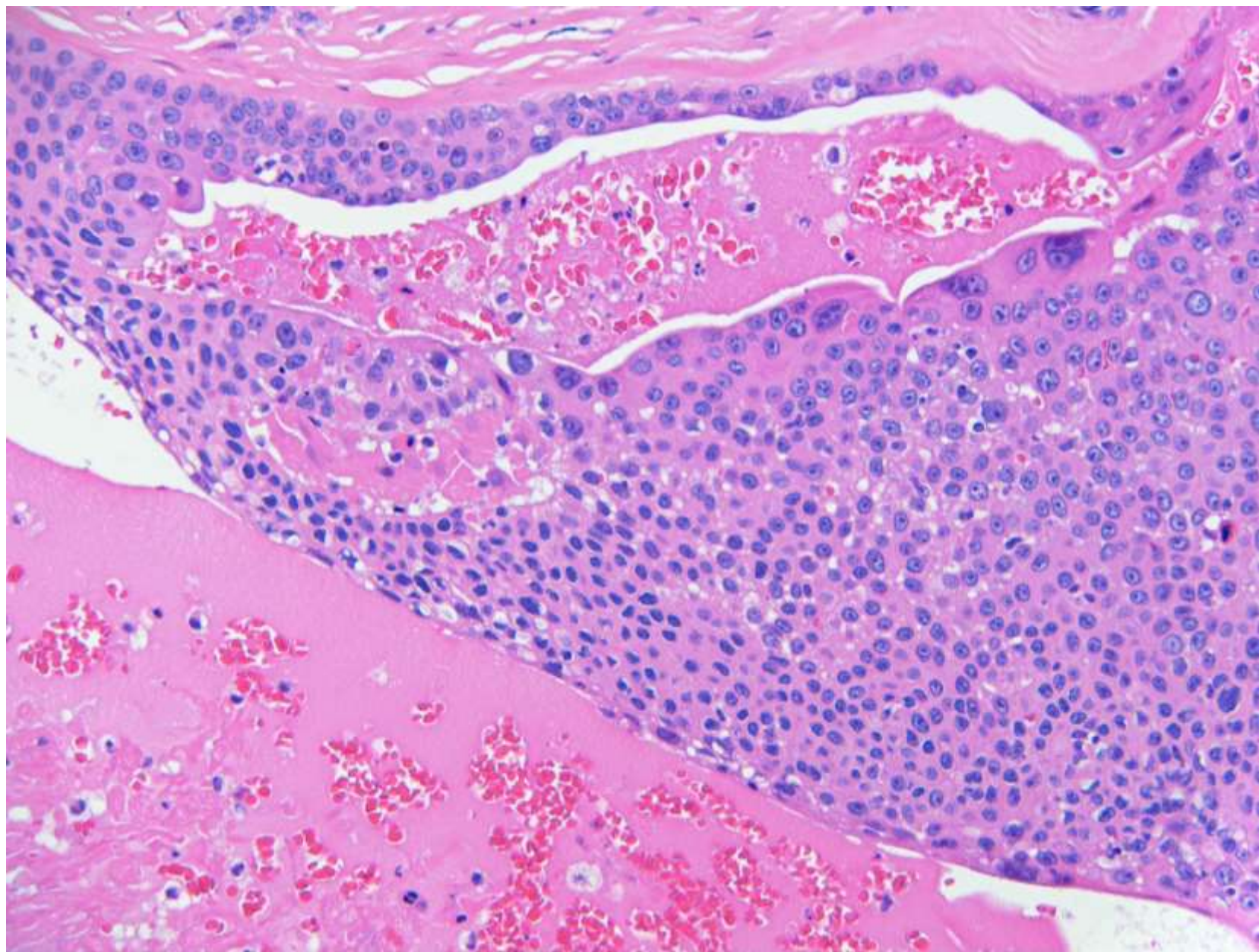


Next Case

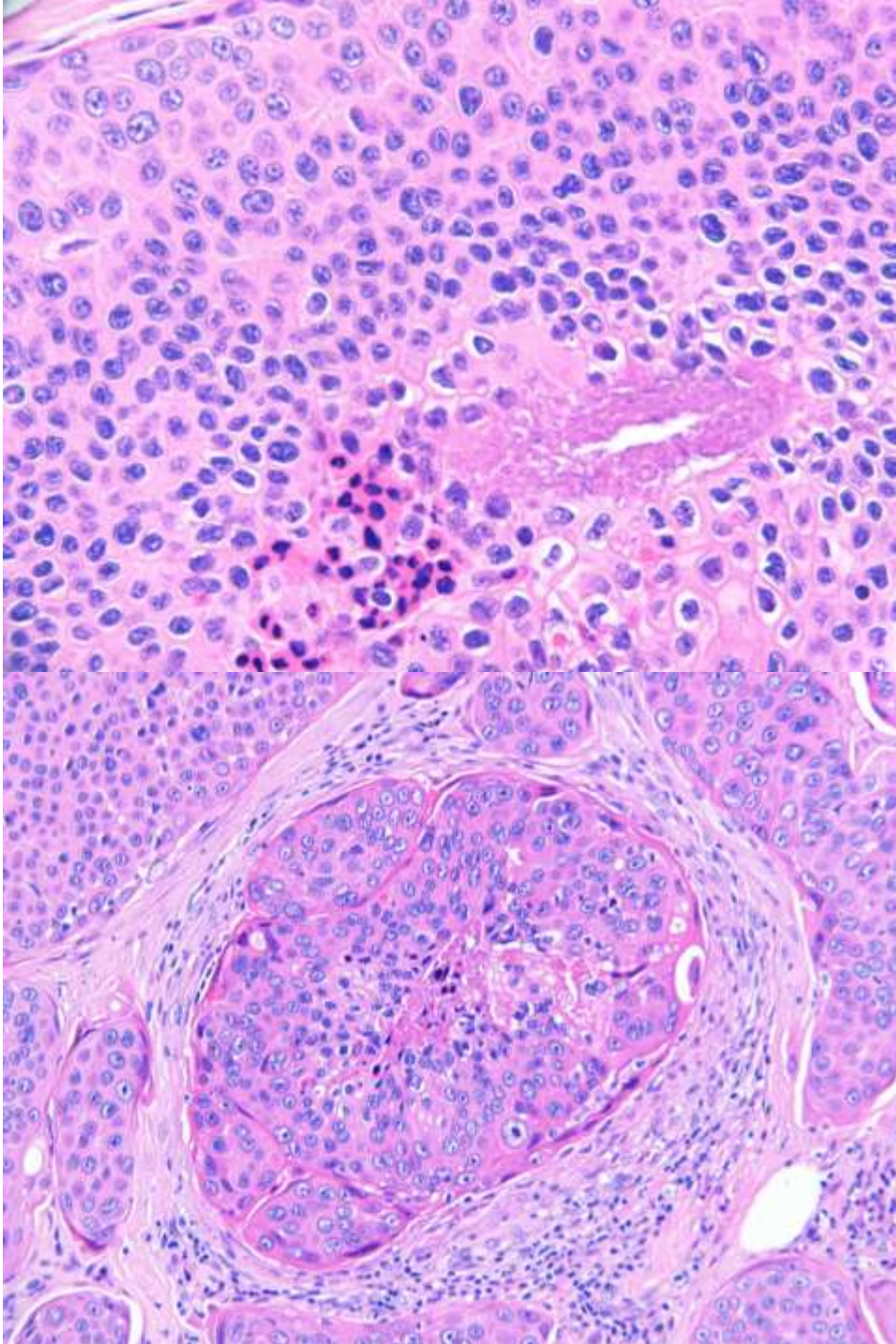
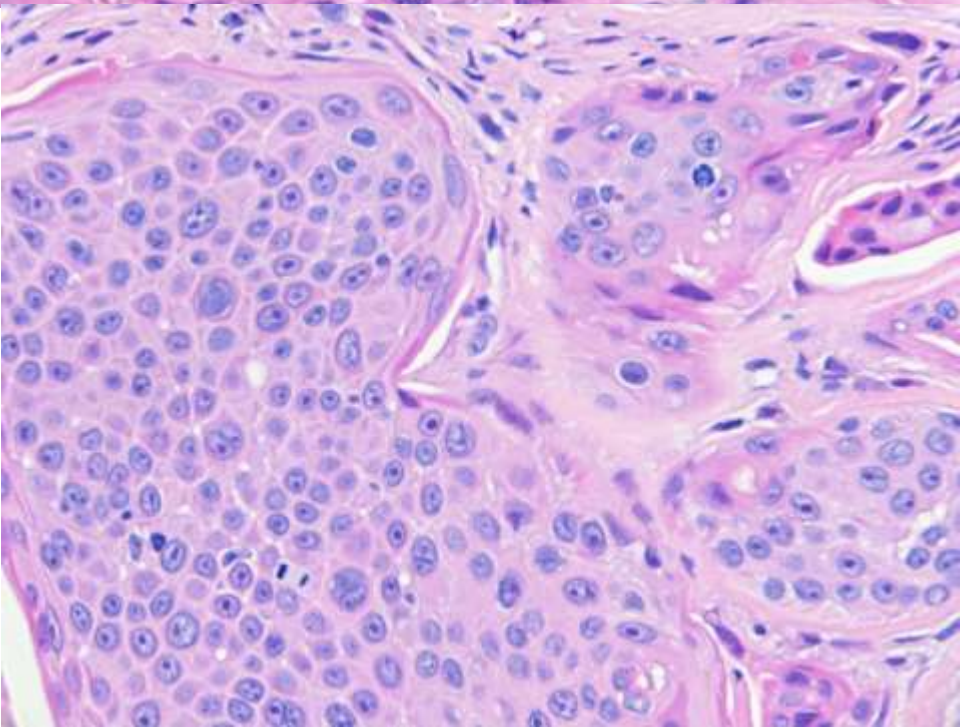
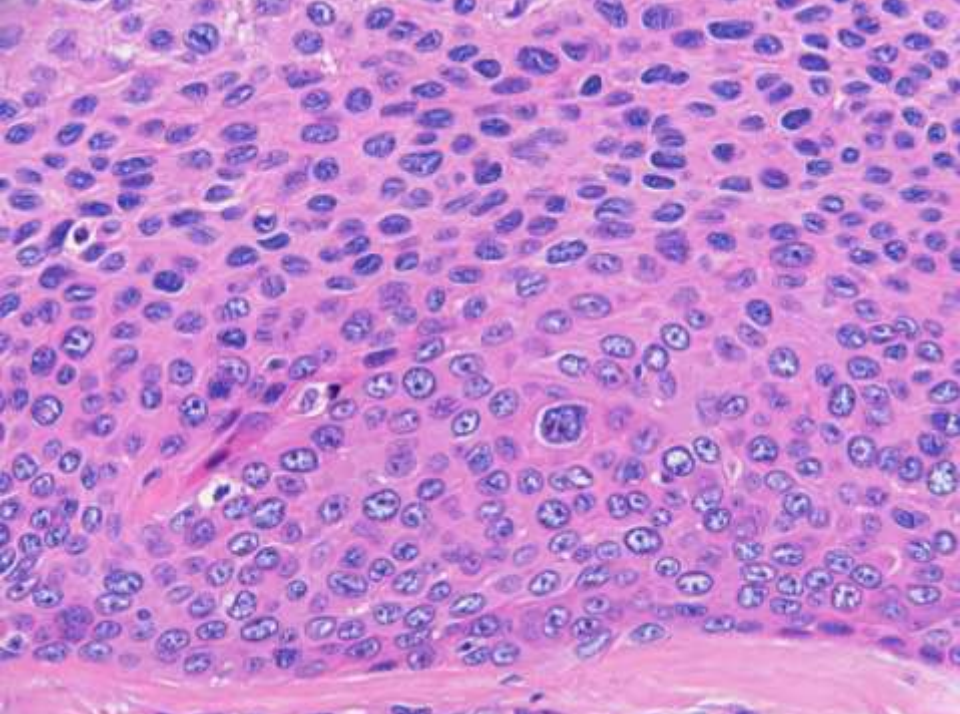
M54. Left middle finger. Soft tissue swelling ?Giant cell PVNS





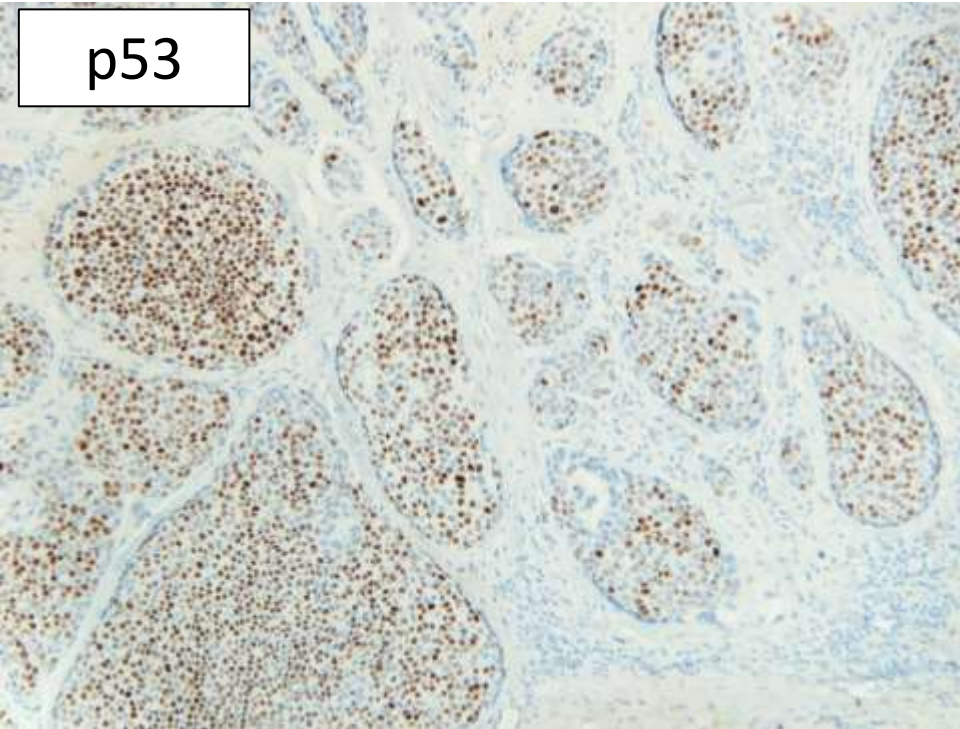




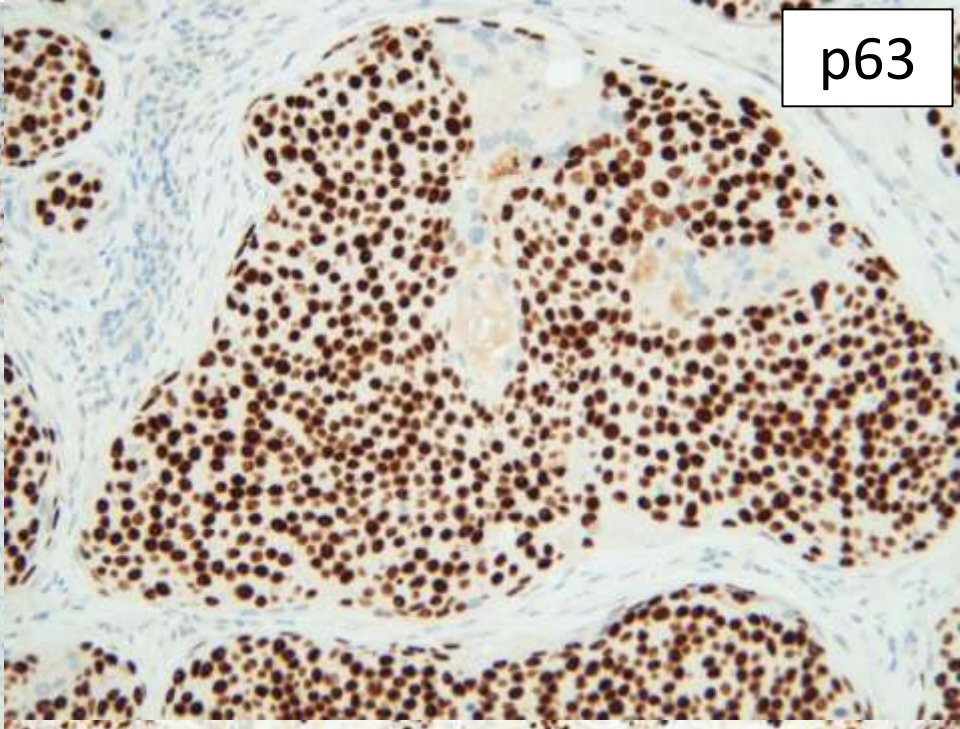




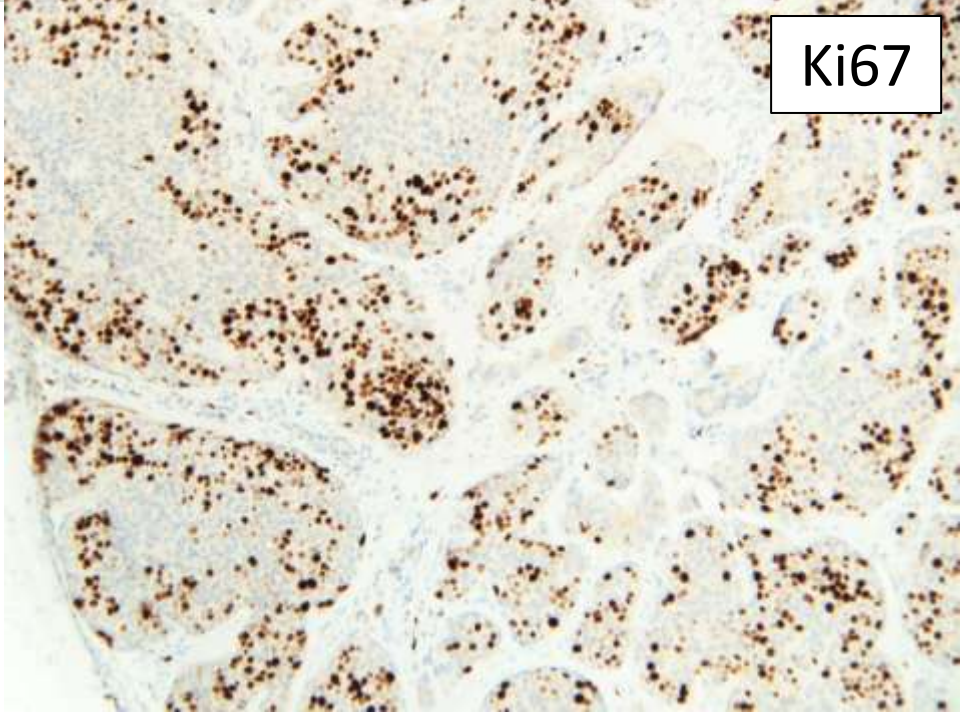
p53



p63



Ki67



- *My Diagnosis* -

**Atypical Hidradenoma**

*Advice: Ensure complete  
excision*

# Atypical / Malignant Hidradenoma

[Nazarian RM et al Modern Pathology 2009; 22: 600-610]

## Atypical

Frequent:

- Loss of circumscription

Occasional:

- Infiltrative growth
- Necrosis (focal)
- Nuclear pleomorphism

## Malignant

Frequent:

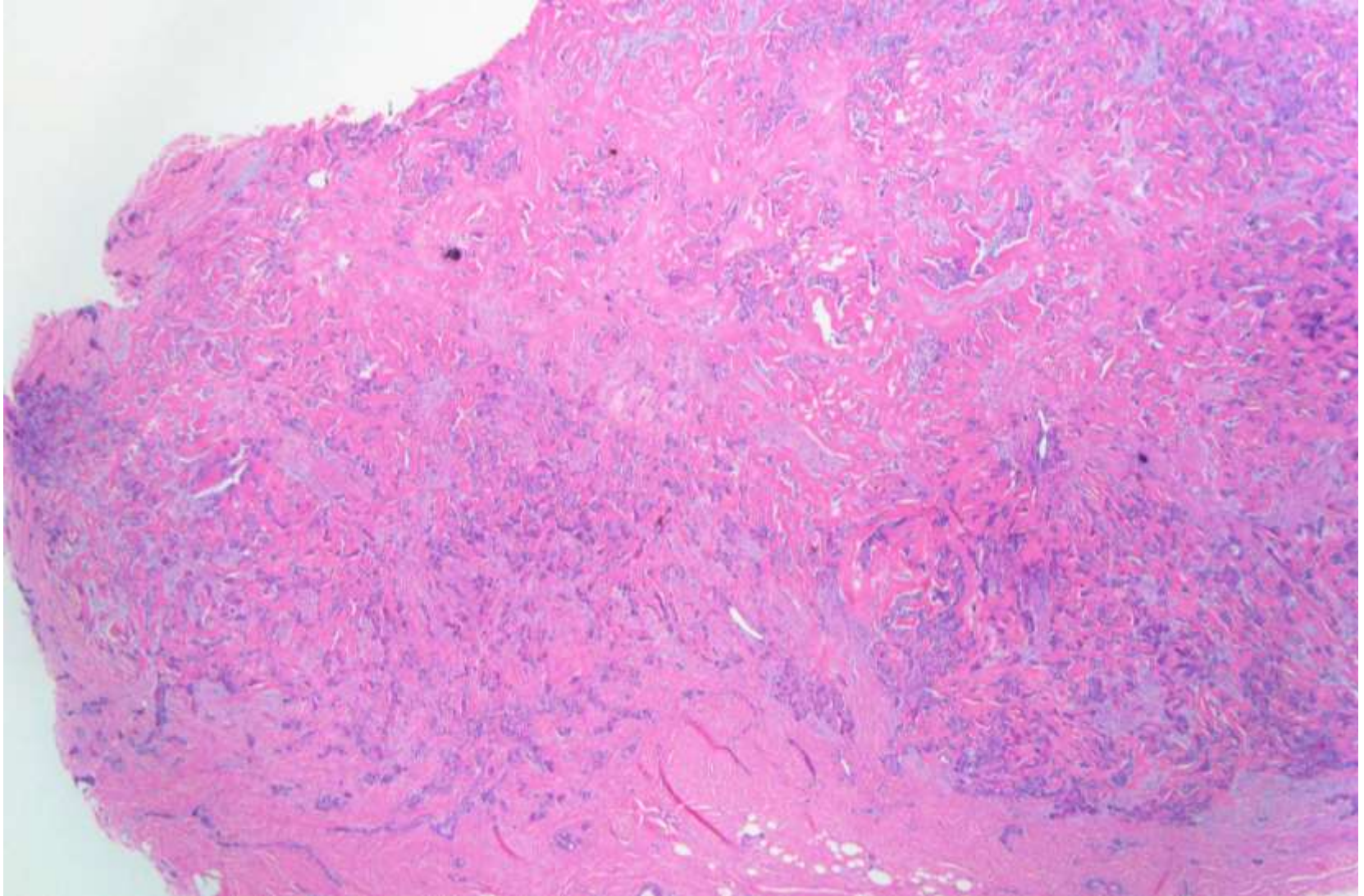
- Loss of circumscription
- Infiltrative growth
- Nuclear pleomorphism
- >4 mitoses/10HPF
- Necrosis (comedo)
- Deep extension

Occasional: PNI/LVI

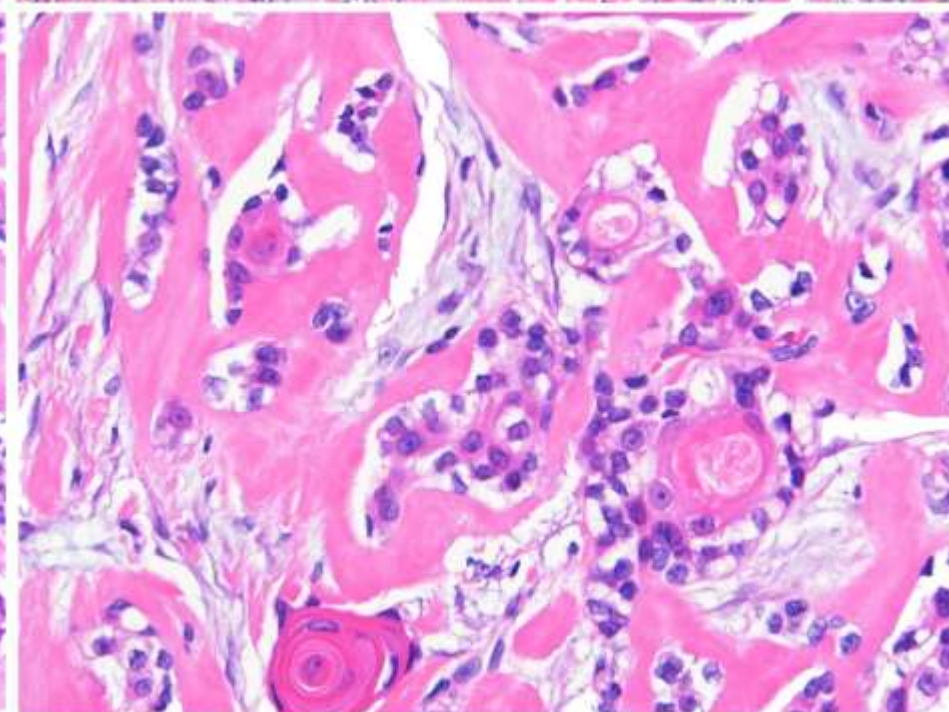
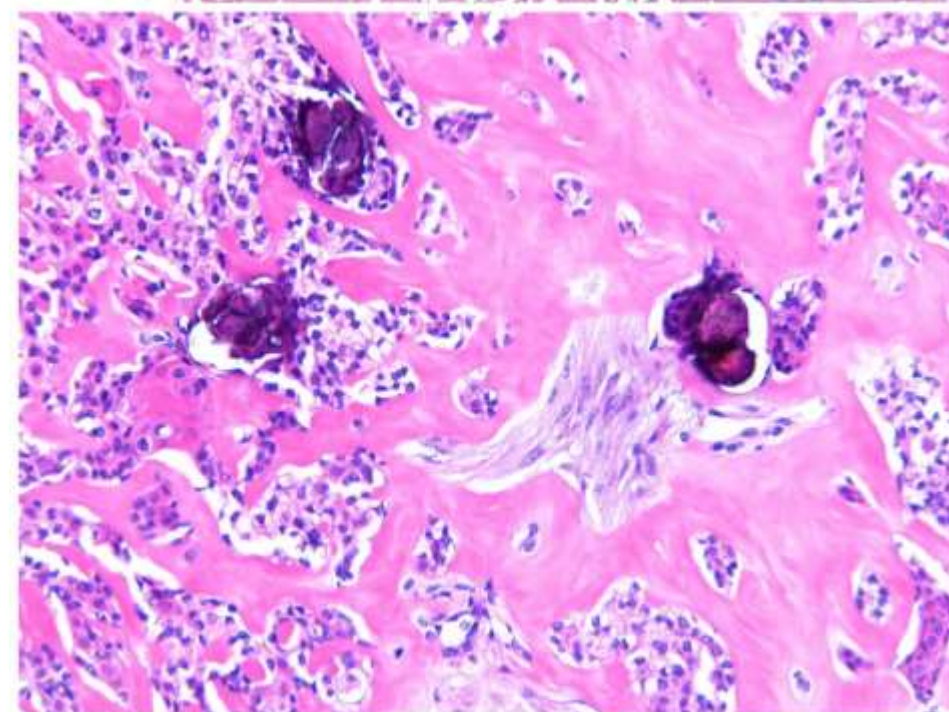
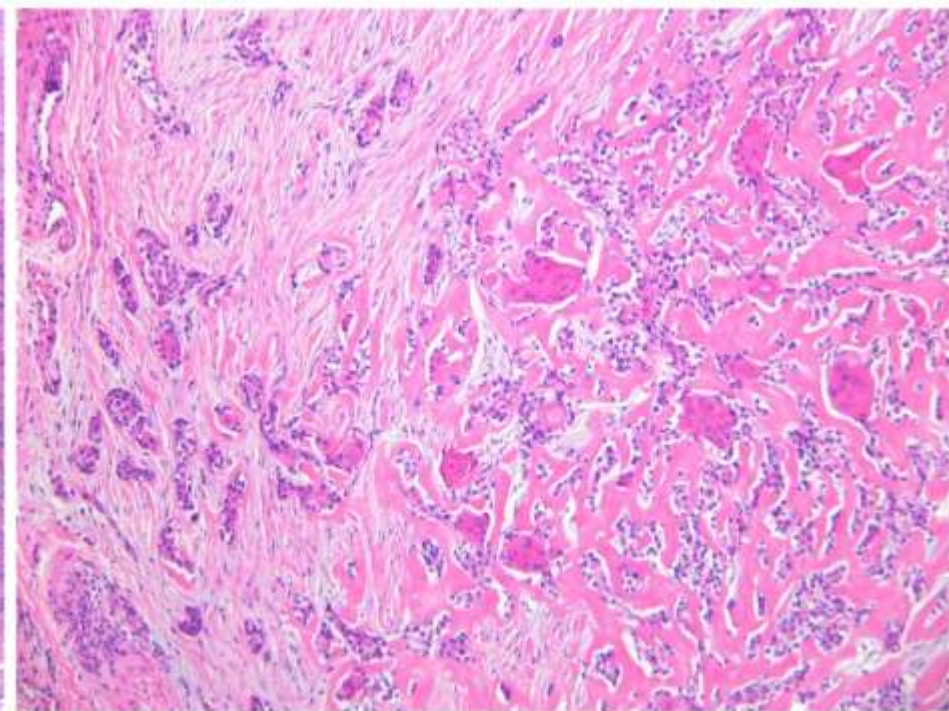
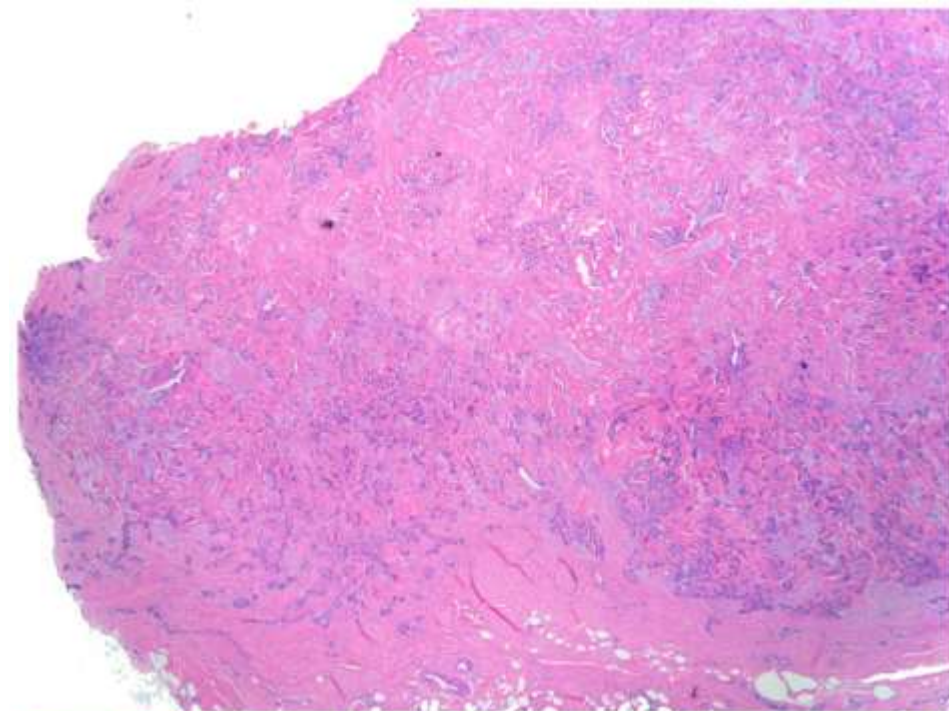


Next Case

(2015) F63. Medial big toe (Rt). Previous cylindroma ear. “Benign” eccrine tumour **left** foot (2001) - Slides n/a. Recurrent swelling medial aspect **Right** big toe ? MTPJ area ? Nature of swelling.

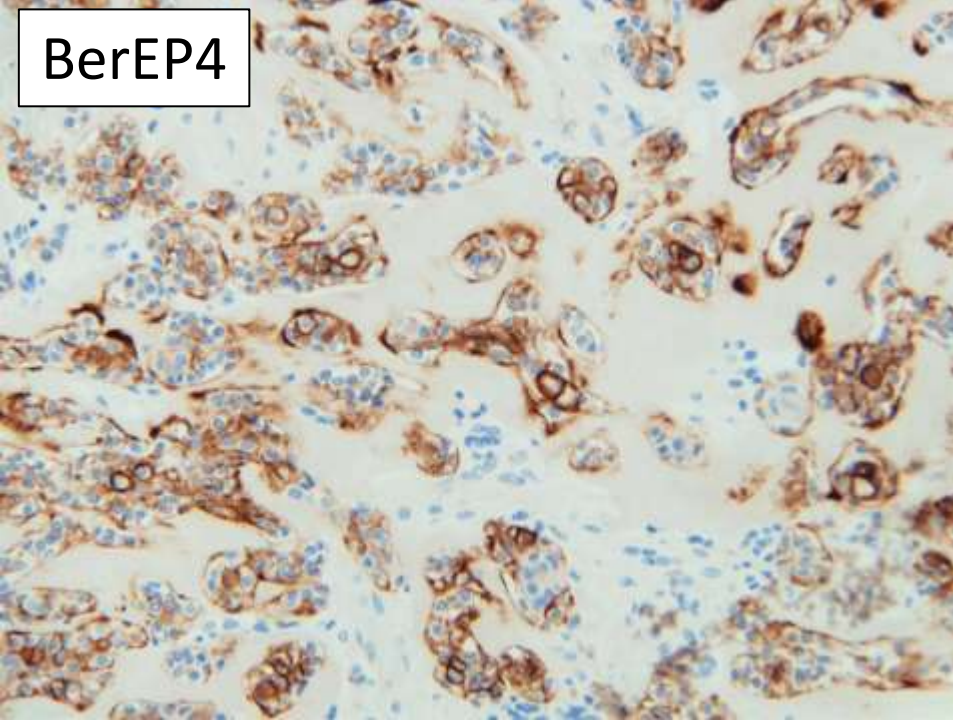




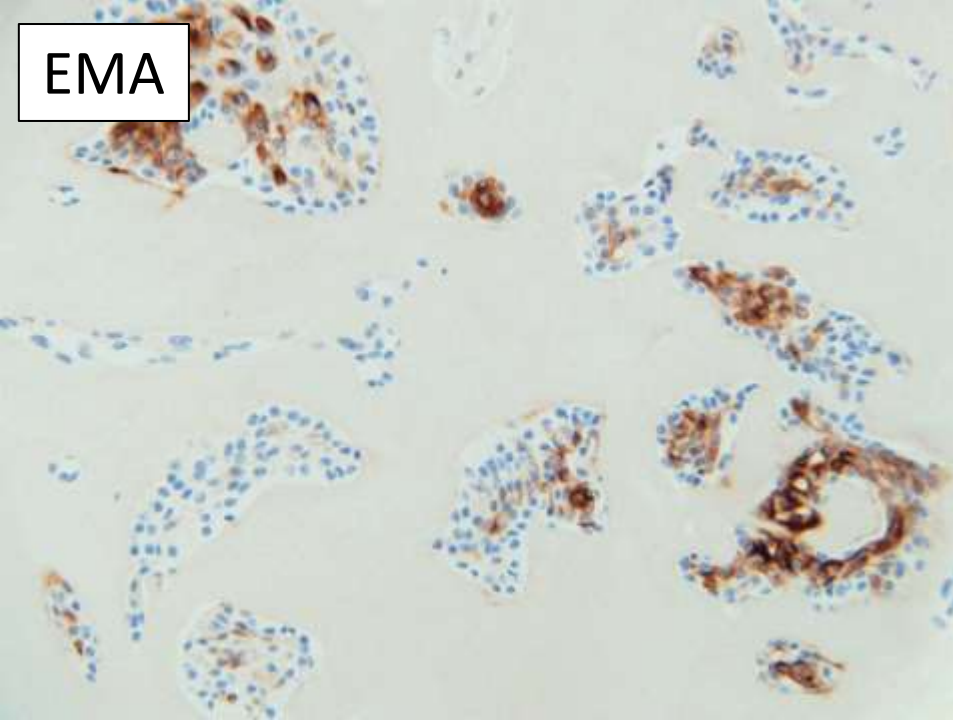




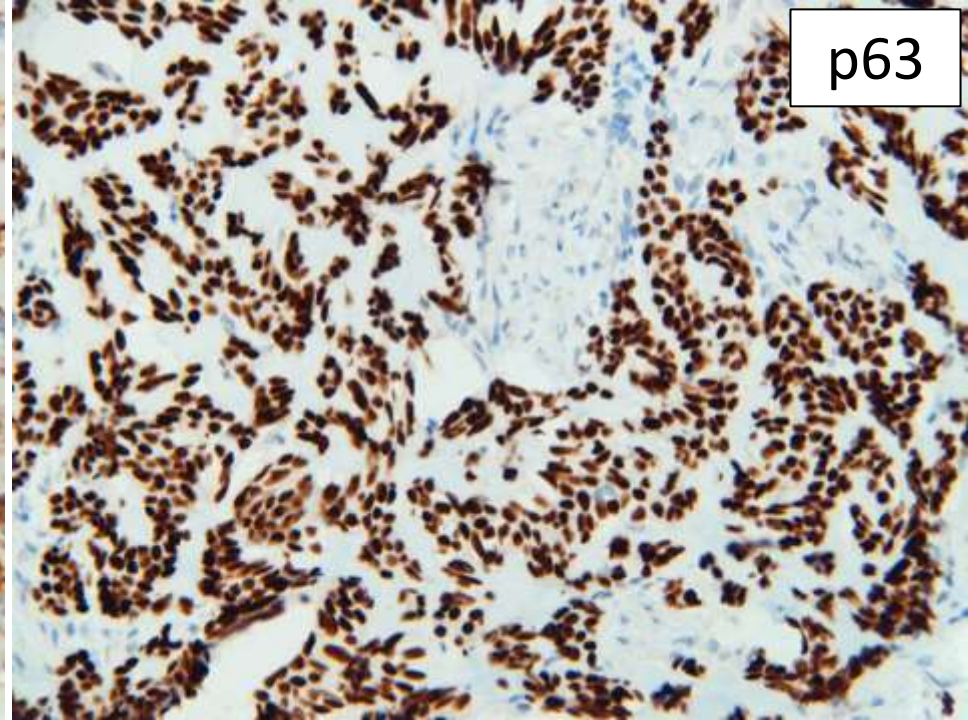
BerEP4



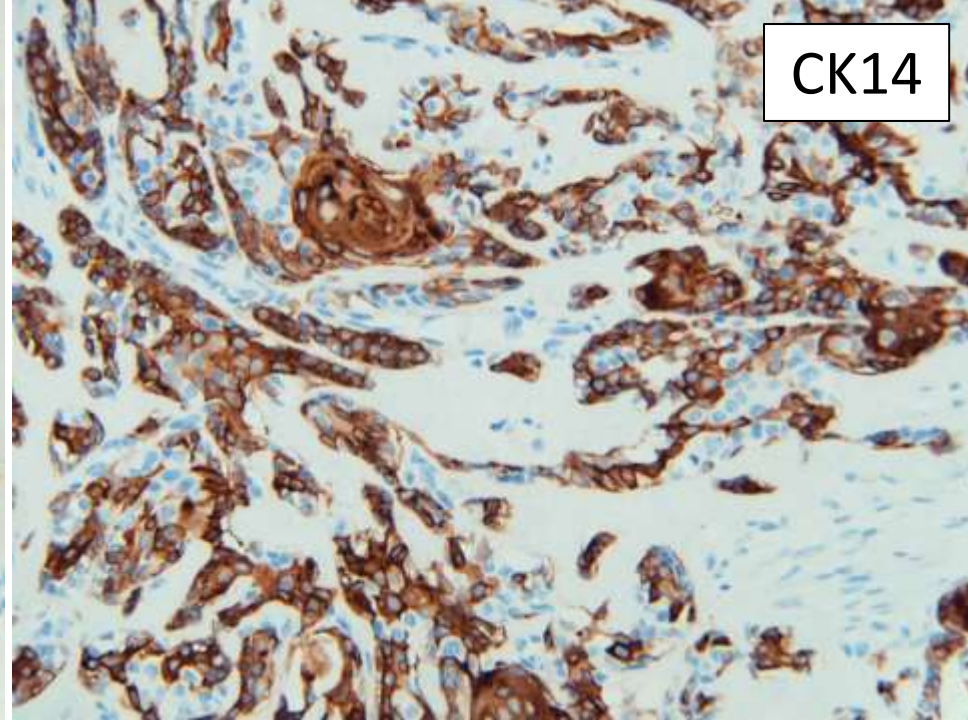
EMA



p63



CK14





A microscopic image of tissue, likely a histological section, showing numerous blue-stained nuclei. A central text box contains diagnostic information. The text is as follows:

- *My Diagnosis* -

**Hidradenoma, sclerosing, pauciluminal**

*Advice: Local recurrence may occur,  
exceedingly low risk for metastasis*

**Ki67**

**Hidradenoma, sclerosing, pauciluminal**

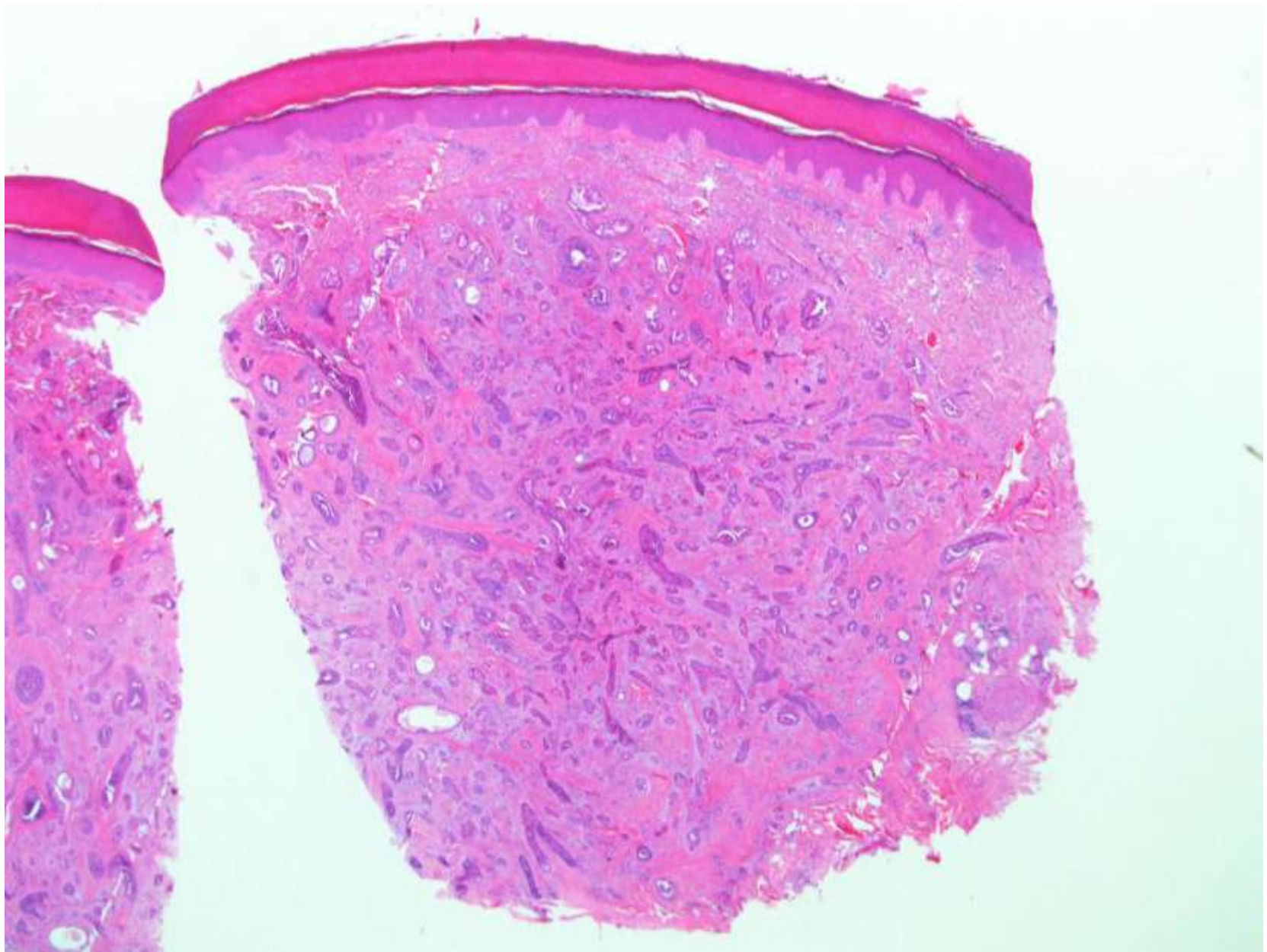
*Advice: Local recurrence may occur,  
exceedingly low risk for metastasis*

**Ki67**

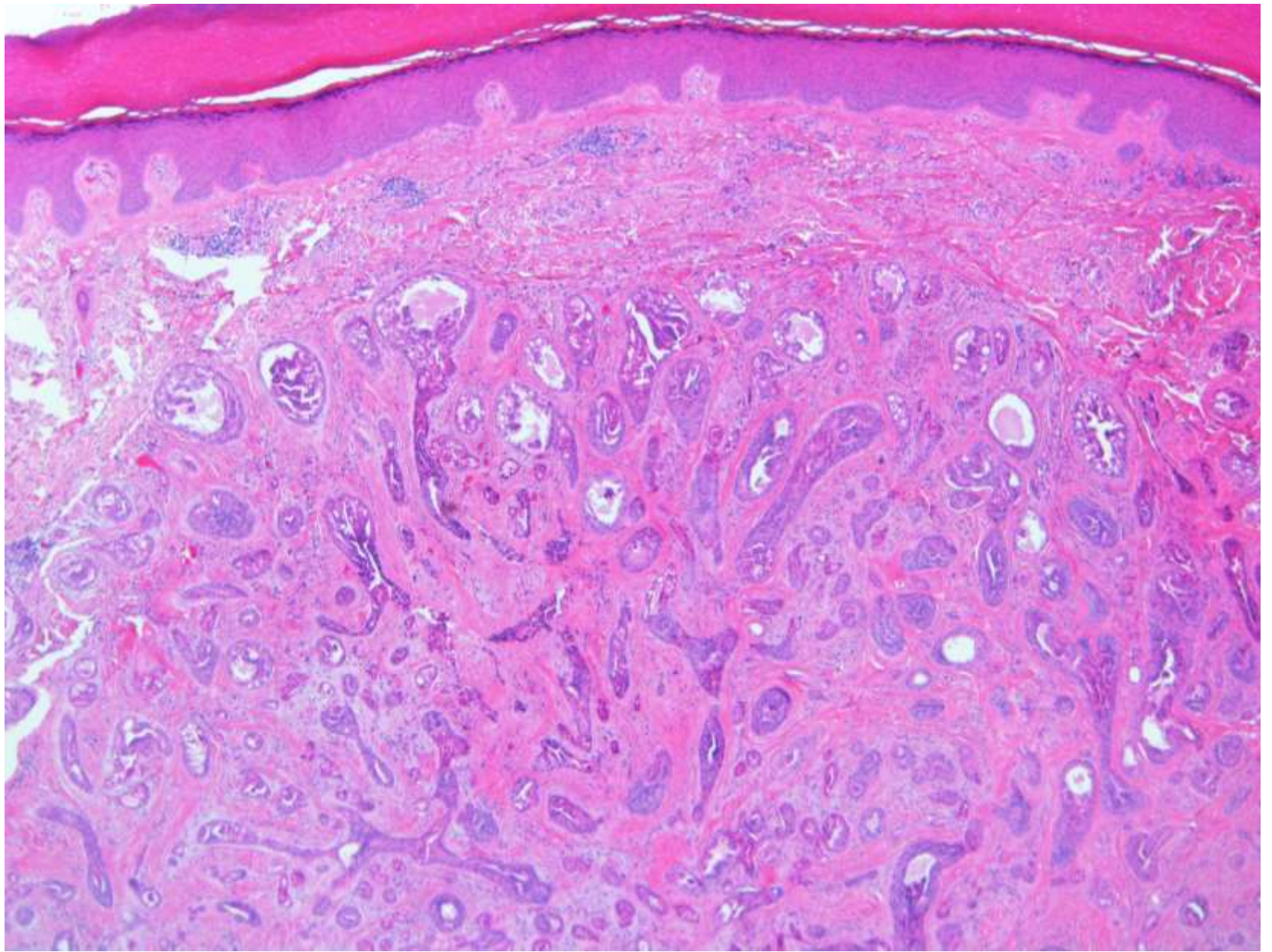
Next Case



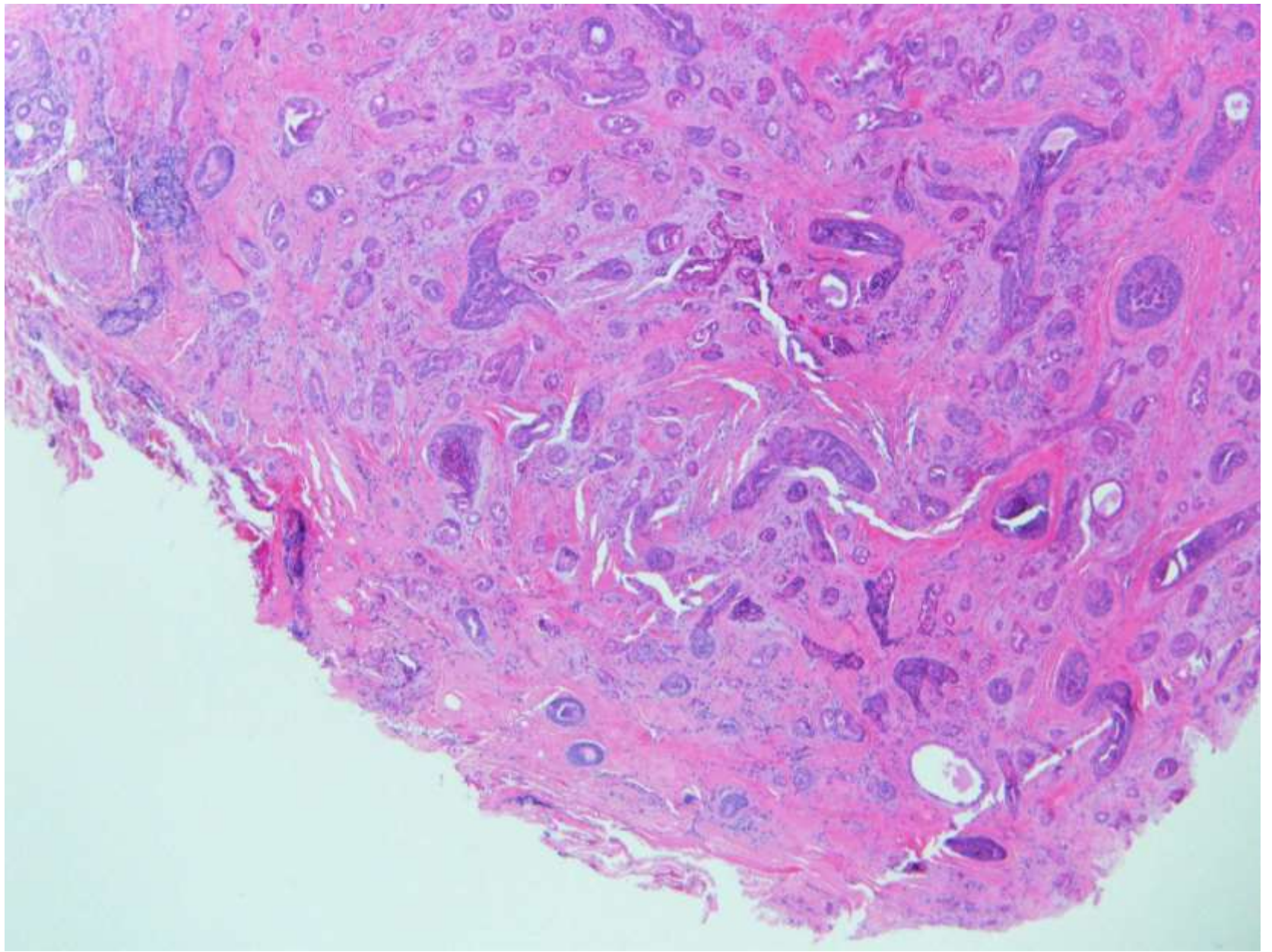
M59. 6/12 months, 13 x 11 pink nodule. ?Lymphoma. To exclude Merkel



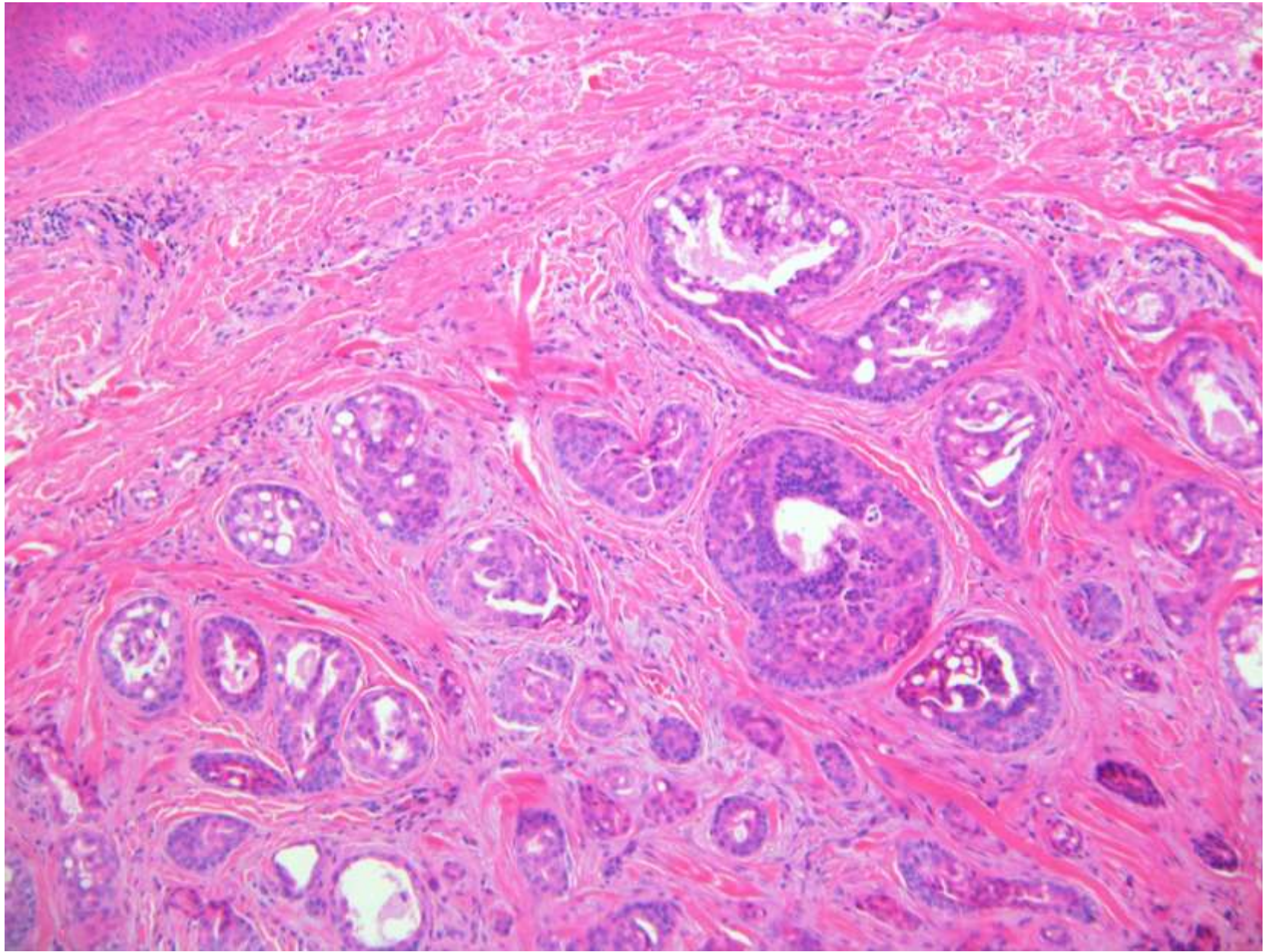




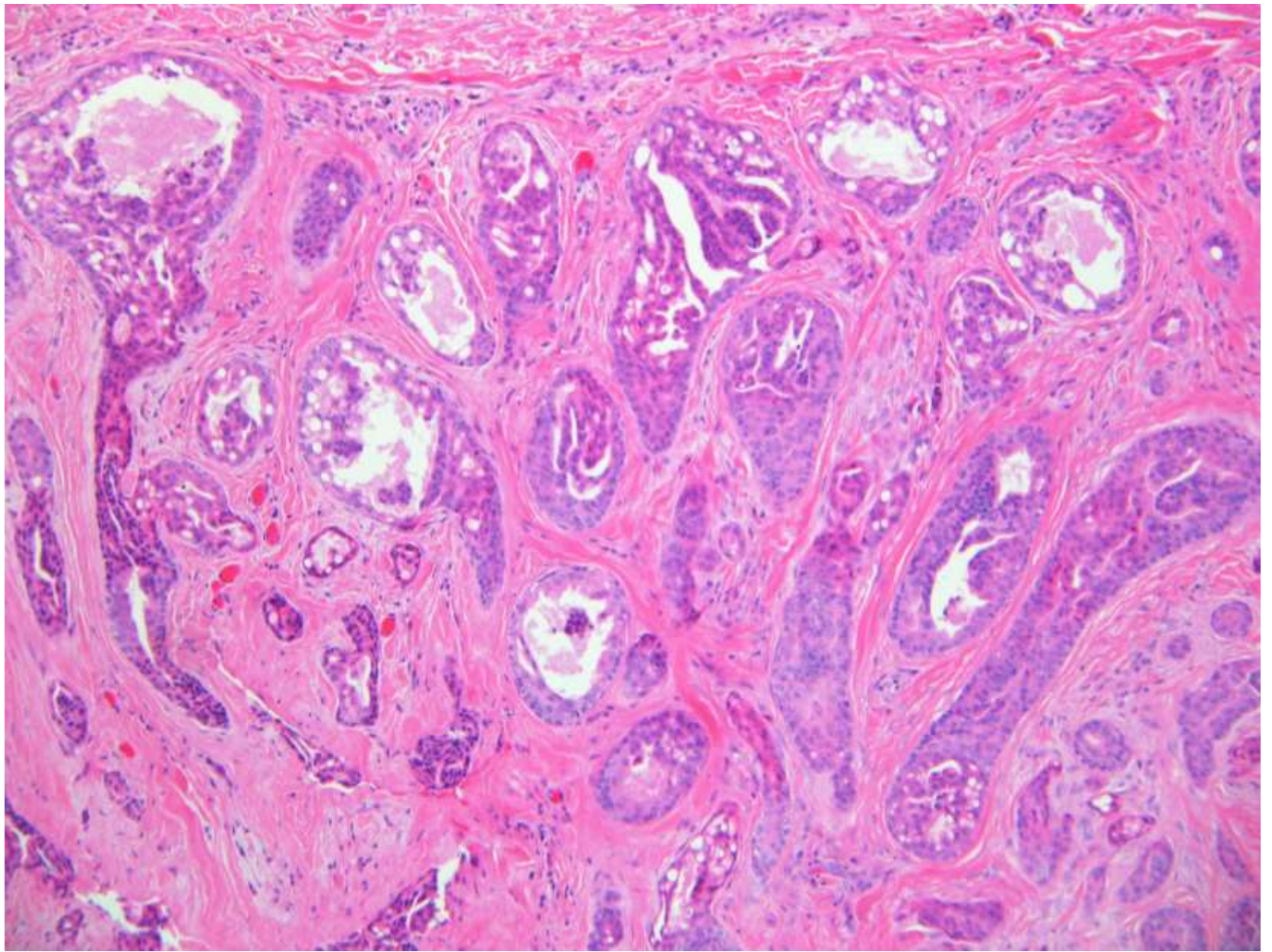




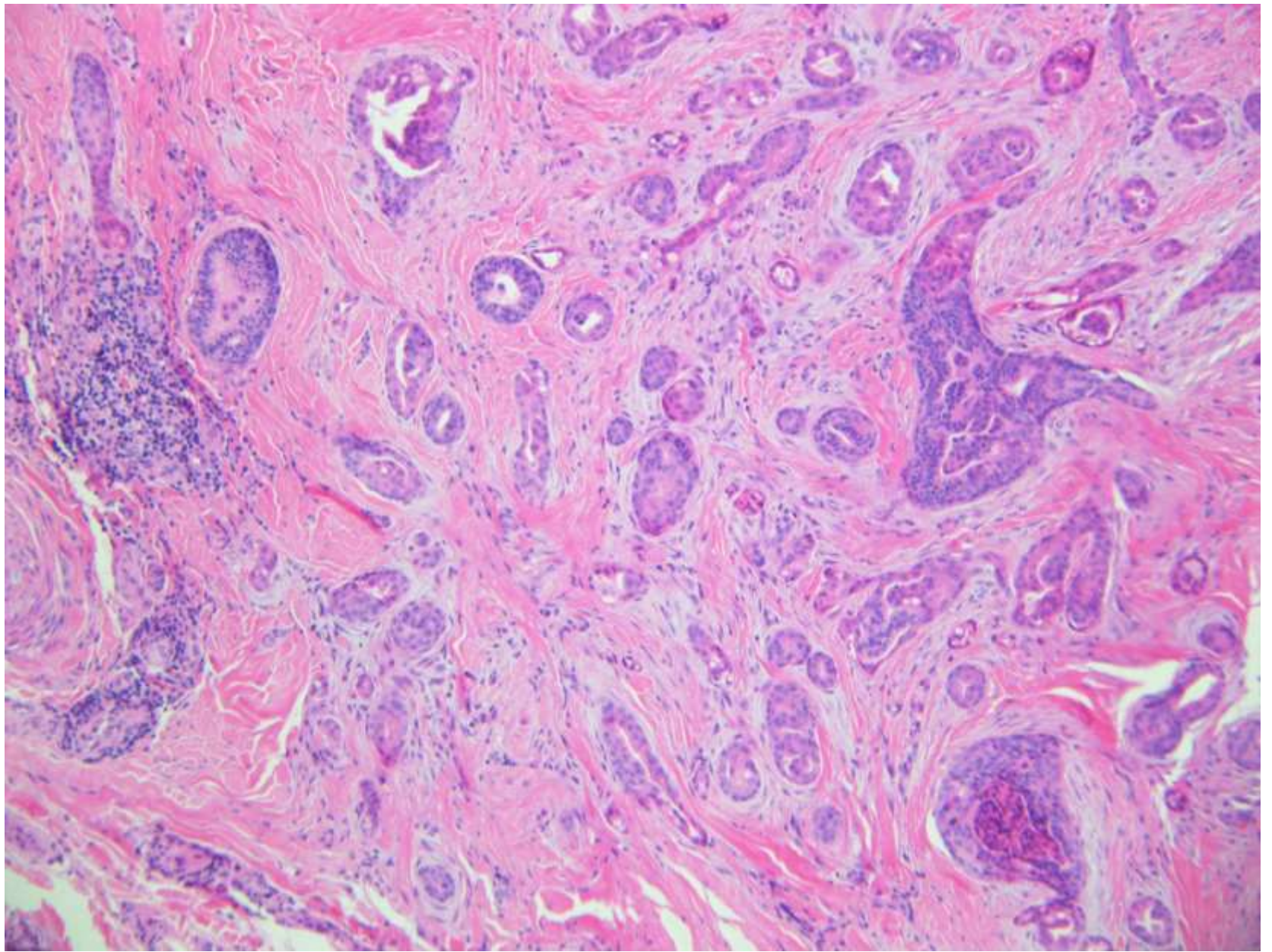




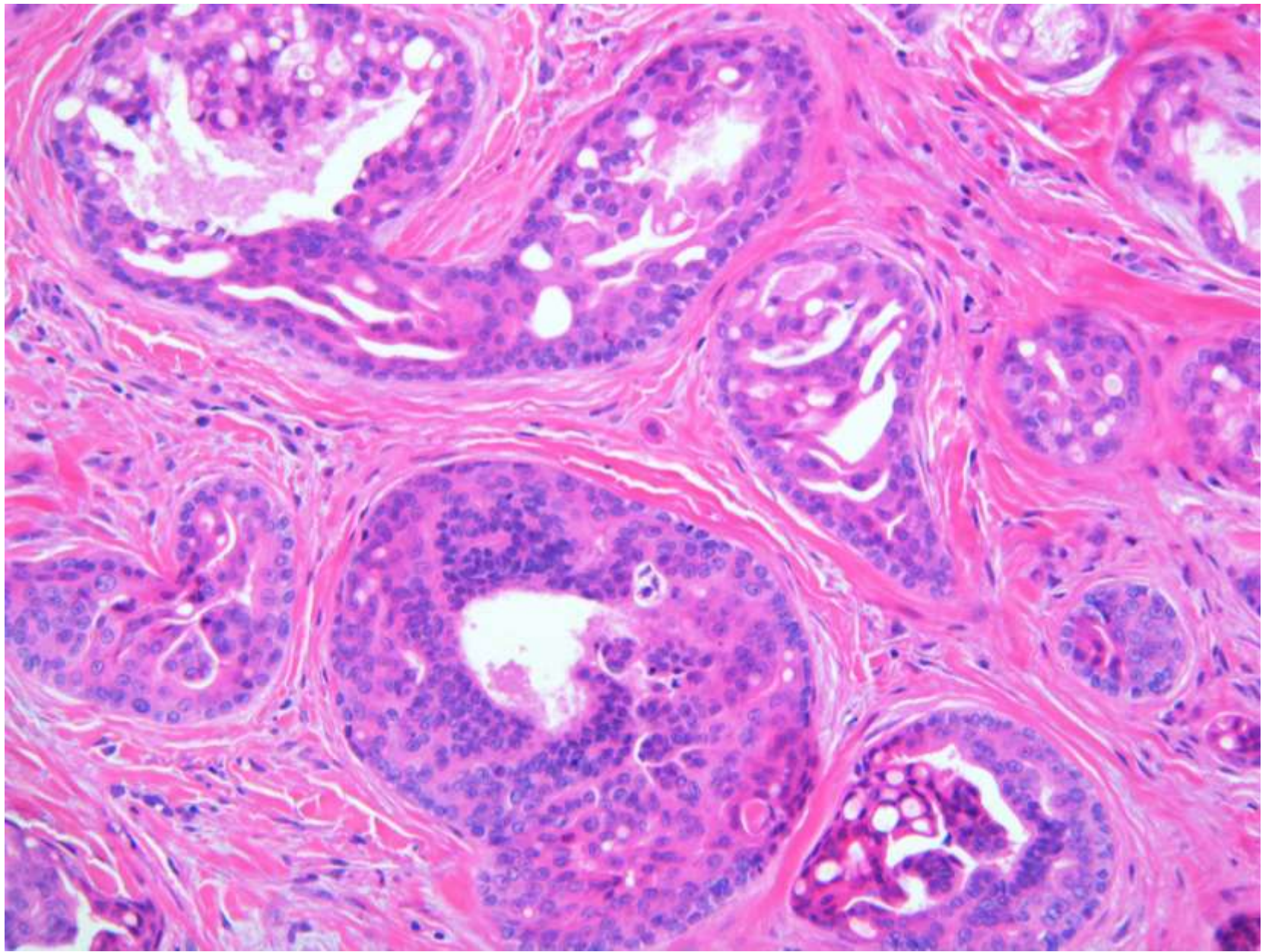




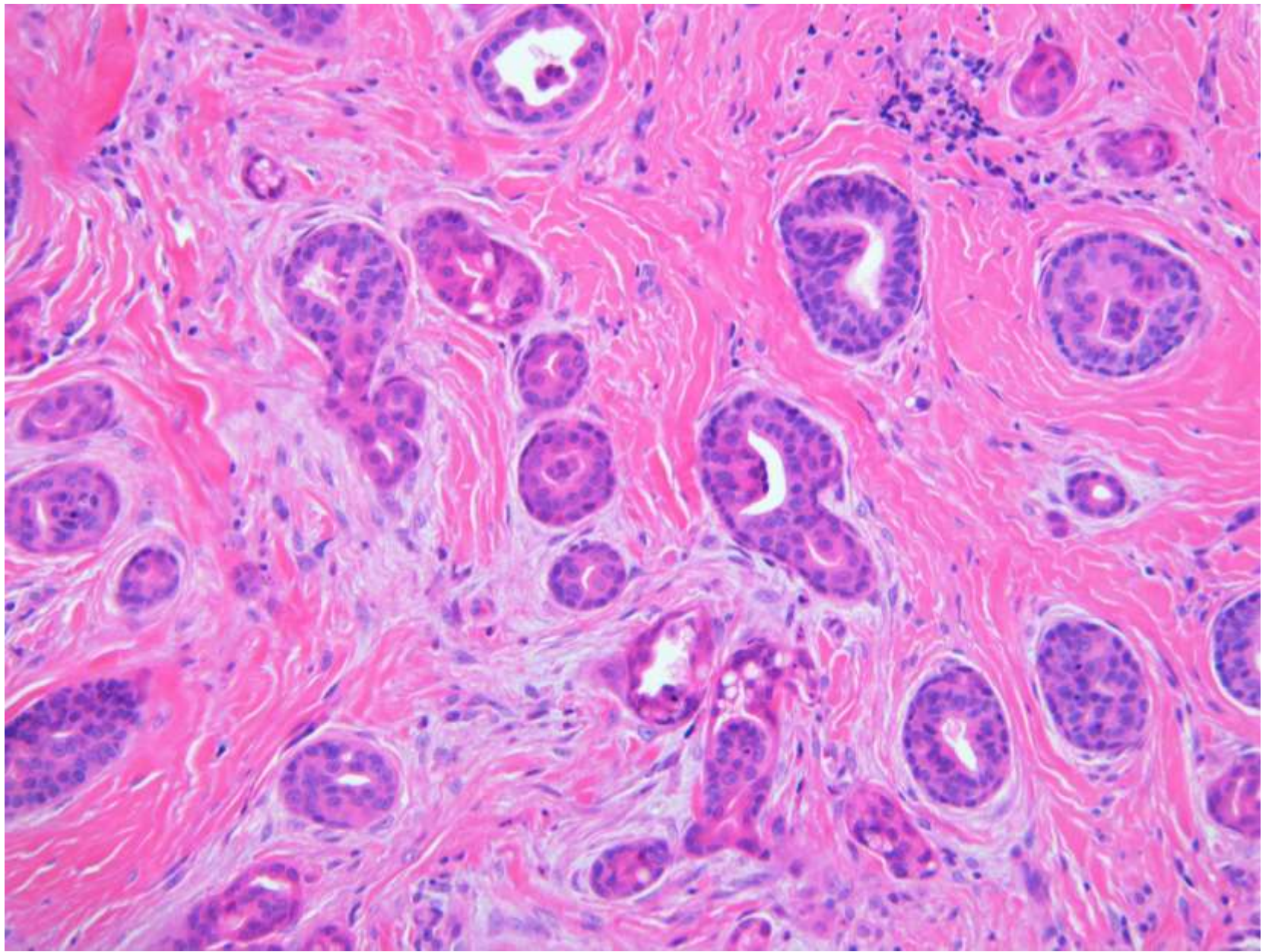




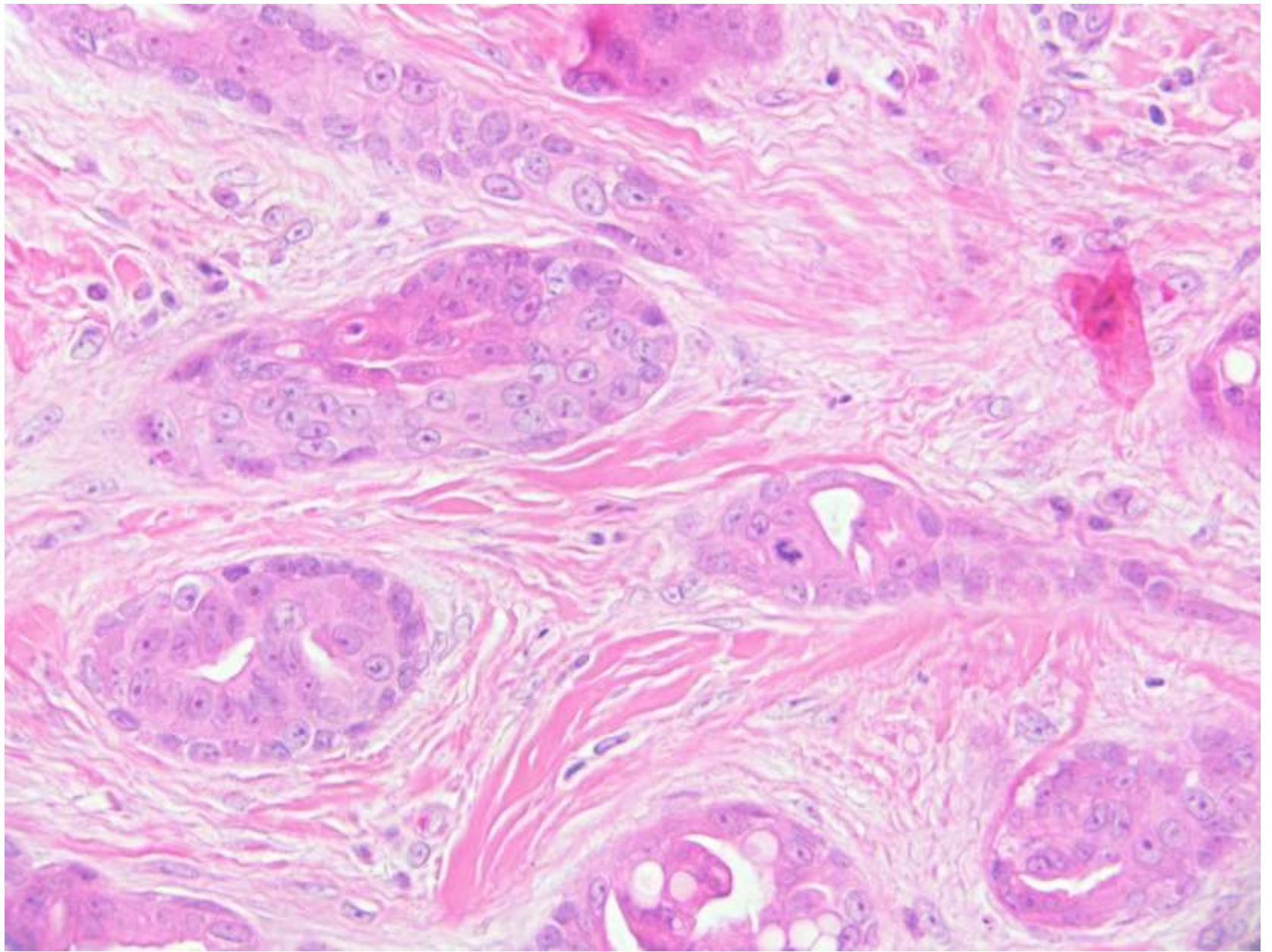




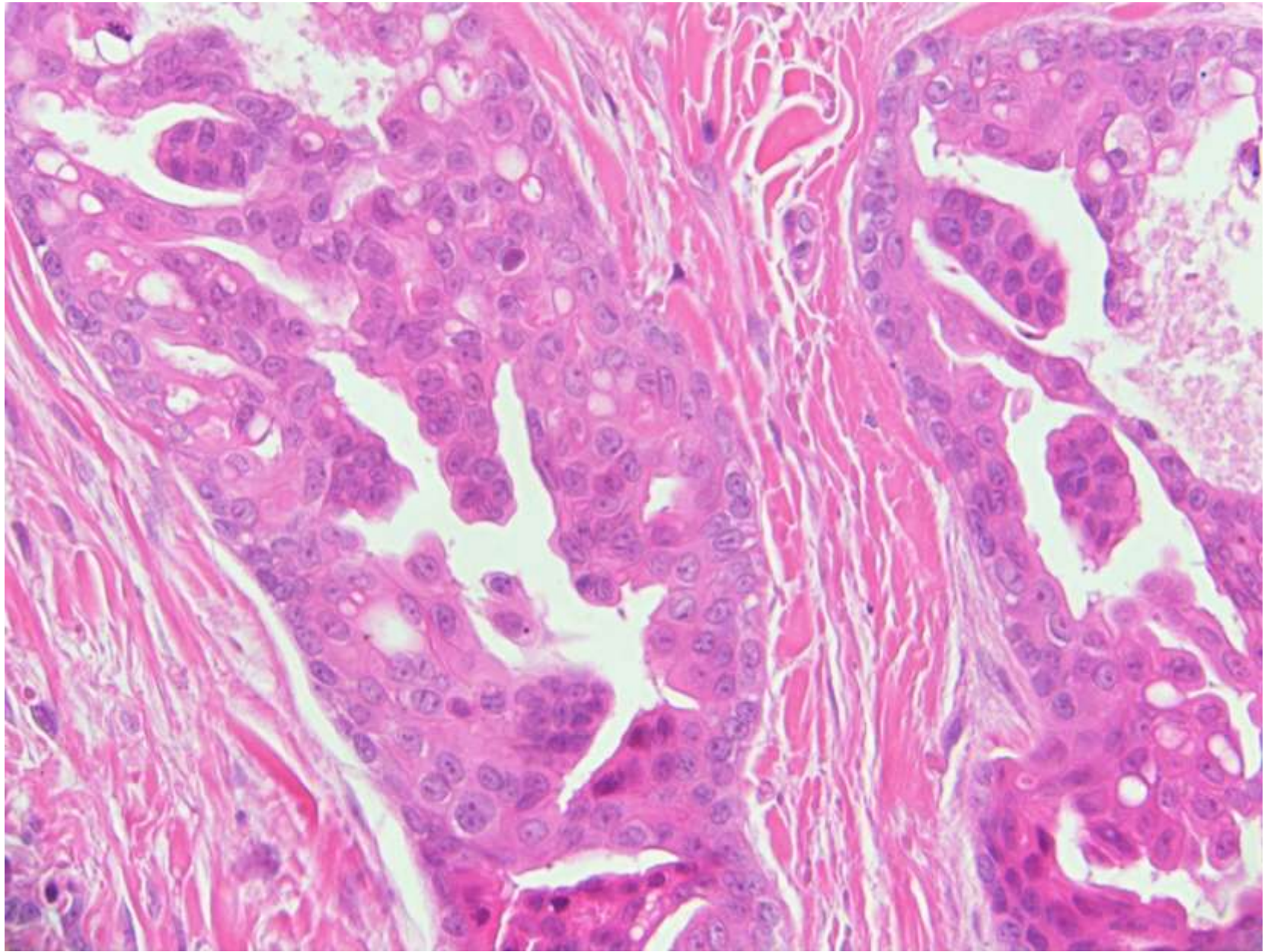




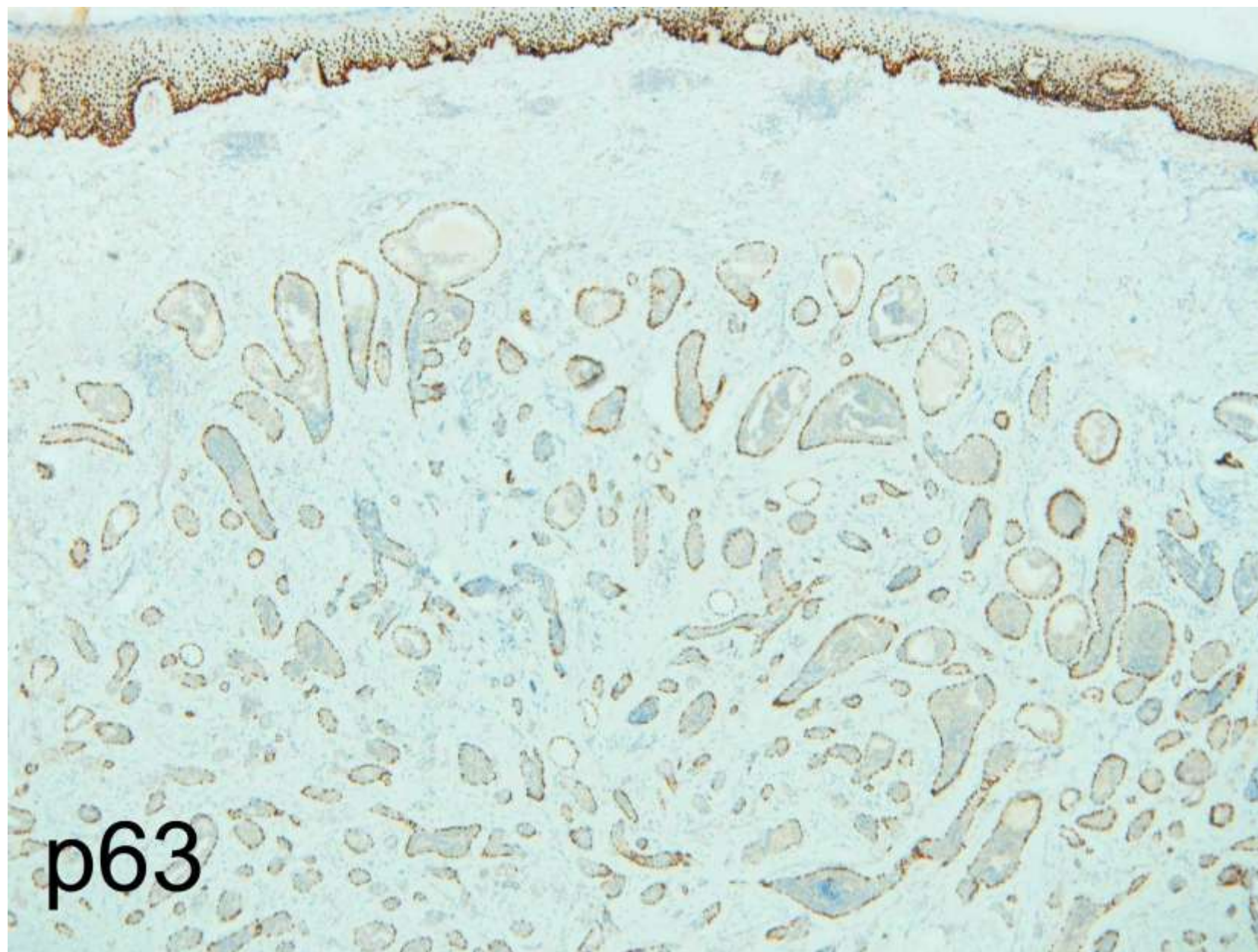


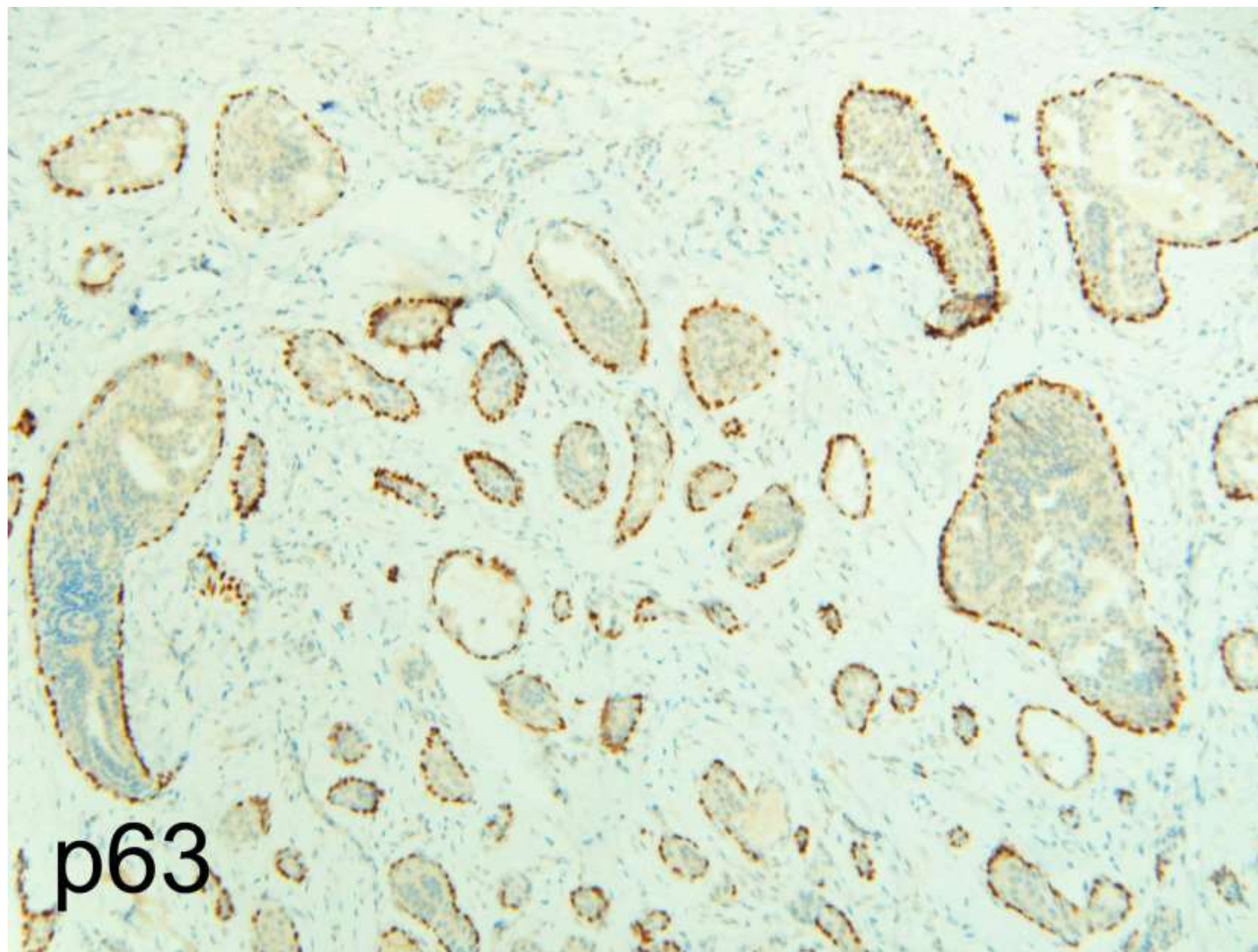




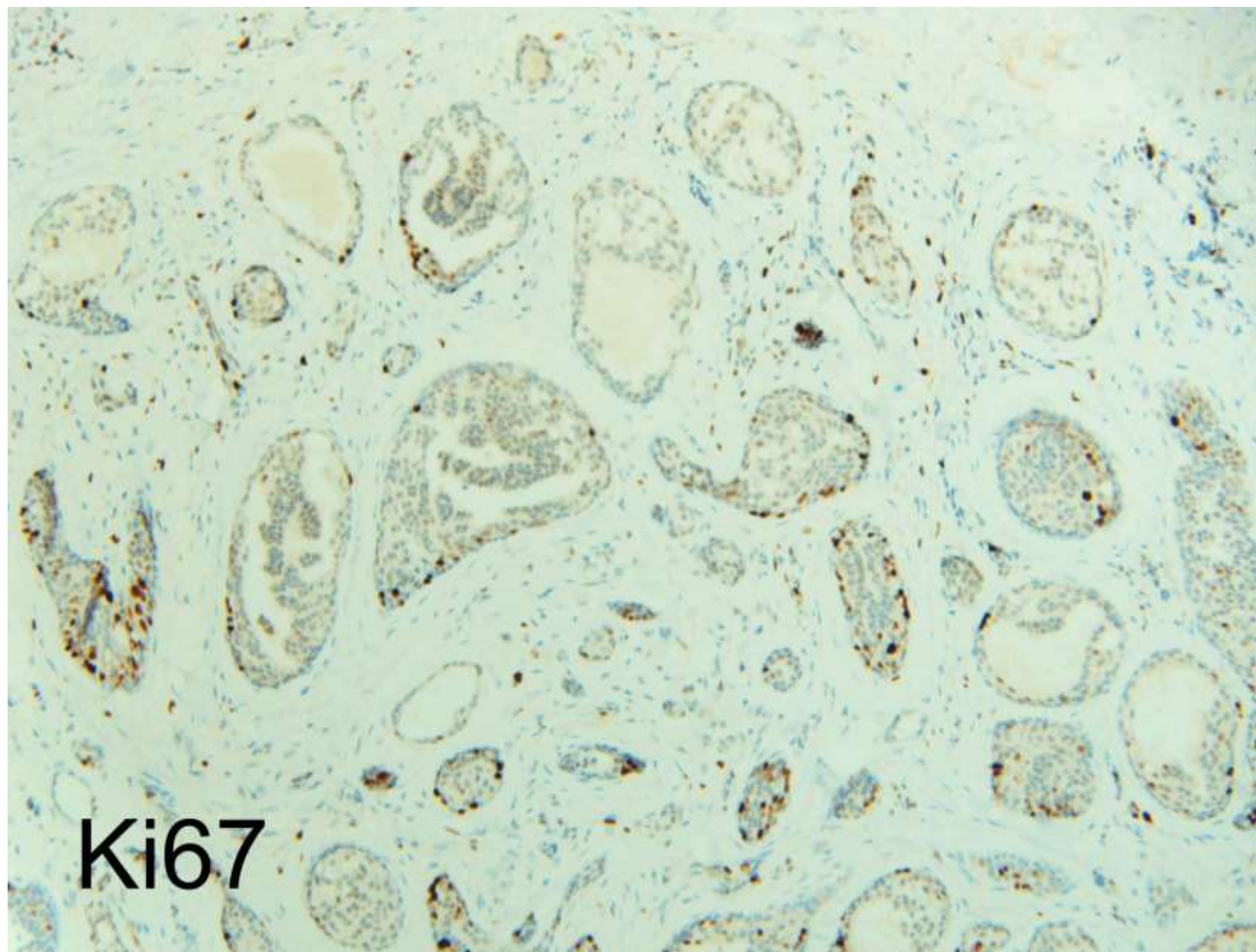












## My report:

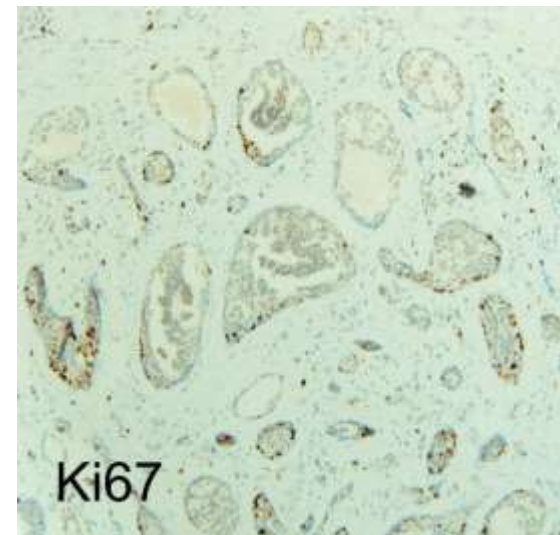
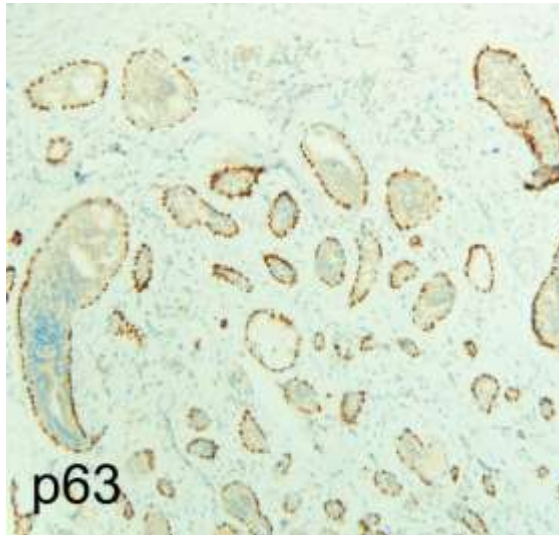
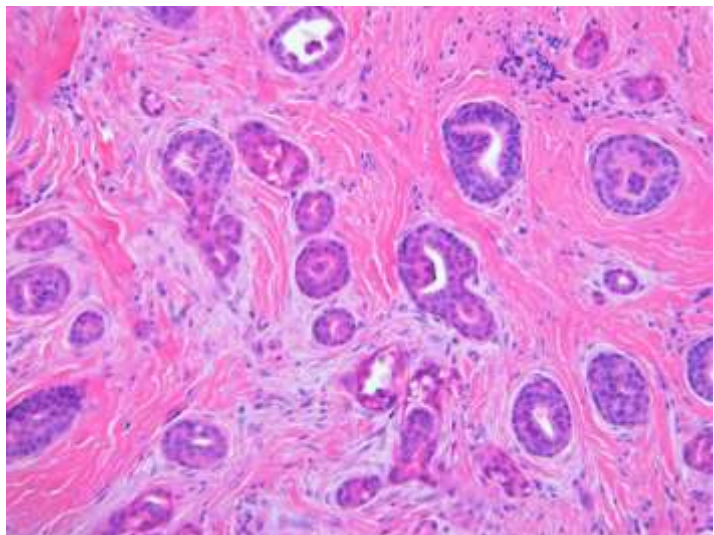
Part of a difficult low grade tubular adenomatous lesion with circumscript upper border (lower border slightly infiltrative but not fully sampled), sparse mitotic activity only and only mild cellular atypia

## Favouring tubular (papillary) adenoma

Complete excision with clear margins is recommended.

## Comments from my slide database

DDx: Digital papillary adenocarcinoma





## **Kazakov's Book: Tubular apocrine adenoma (papillary eccrine adenoma)**

Micropapillary or rarely true papillary projections

Well circumscribed, and composed of ductal structures with a distinct two cell layer

Solid areas and back-to-back glands not present.

The cystic ductal structures in DPA are usually larger and more dilated than those in tubular adenoma

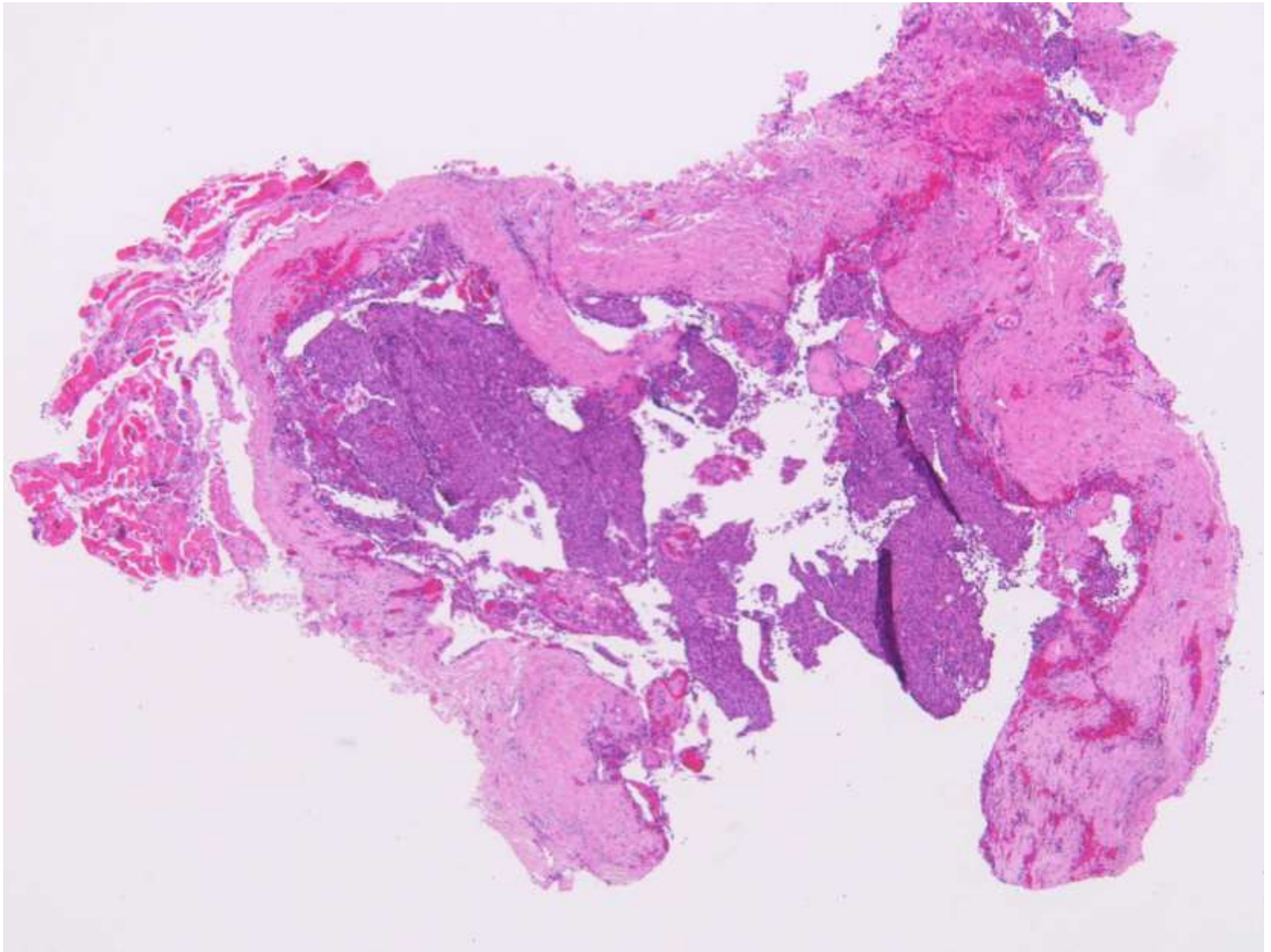
Not so polymorphous

**From Dermpathpro:** They all called it digital papillary adenocarcinoma!!

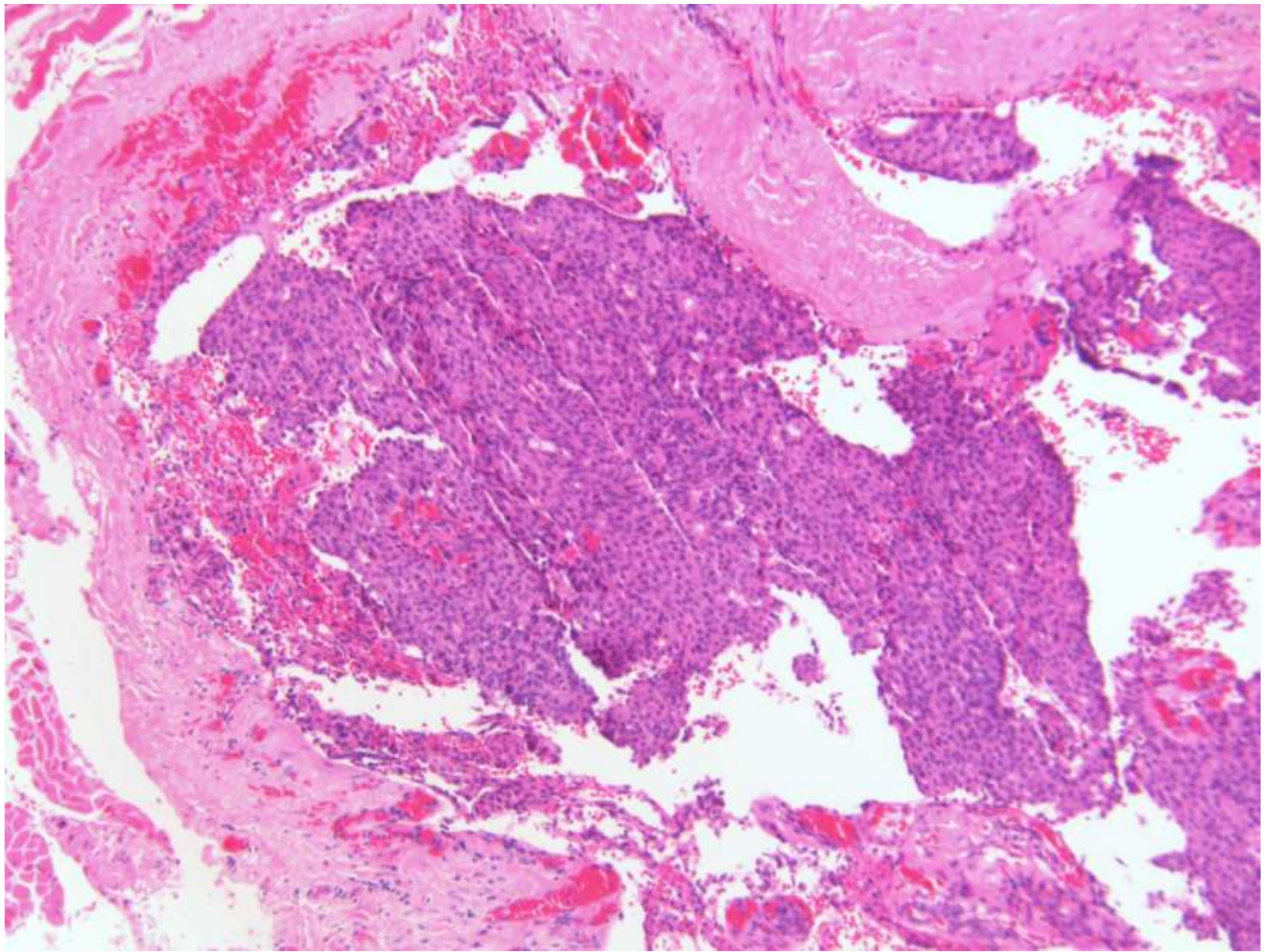
Next Case



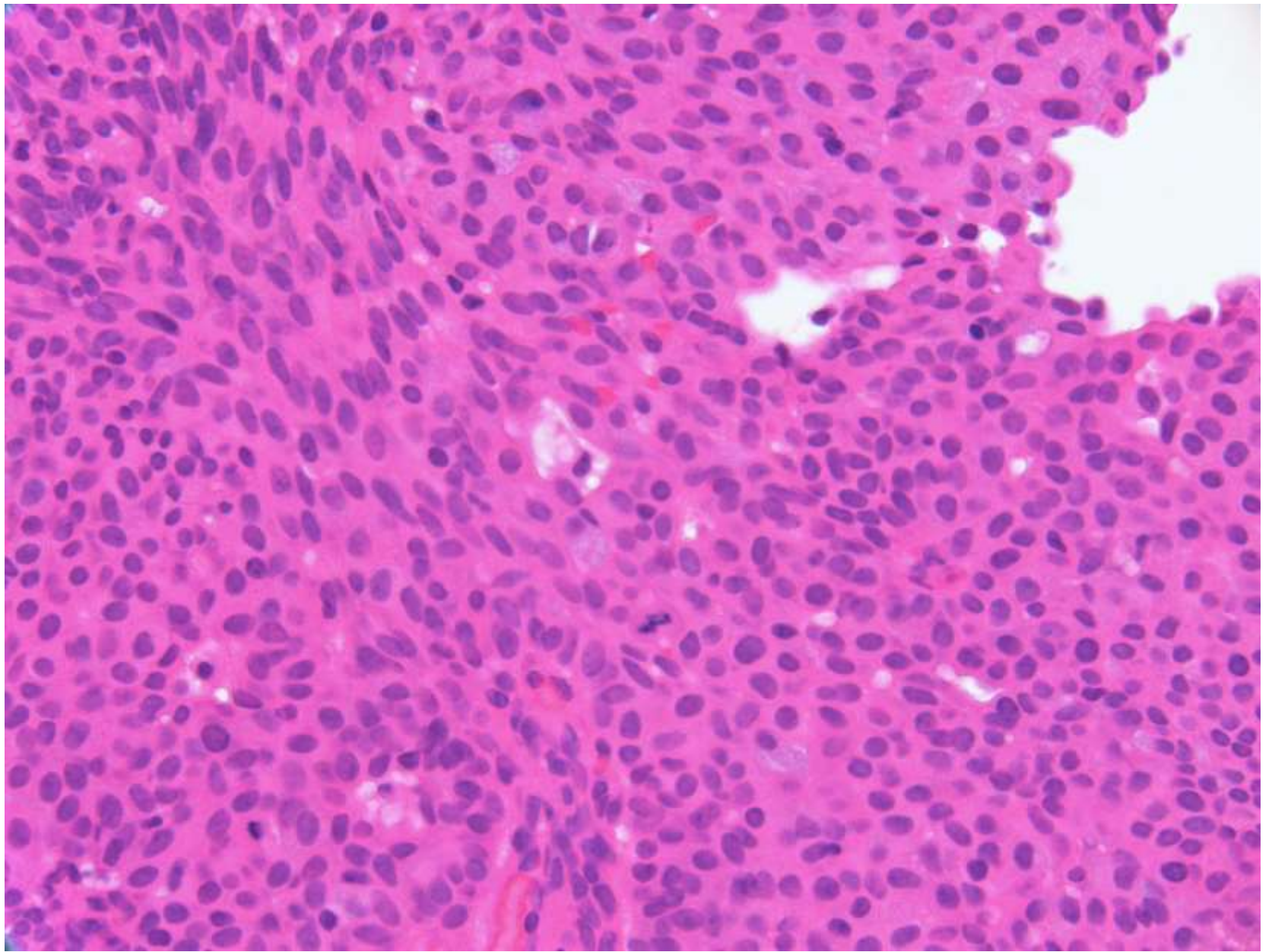
F69. Left upper lid. Cyst of Moll. Burst during last part of excision

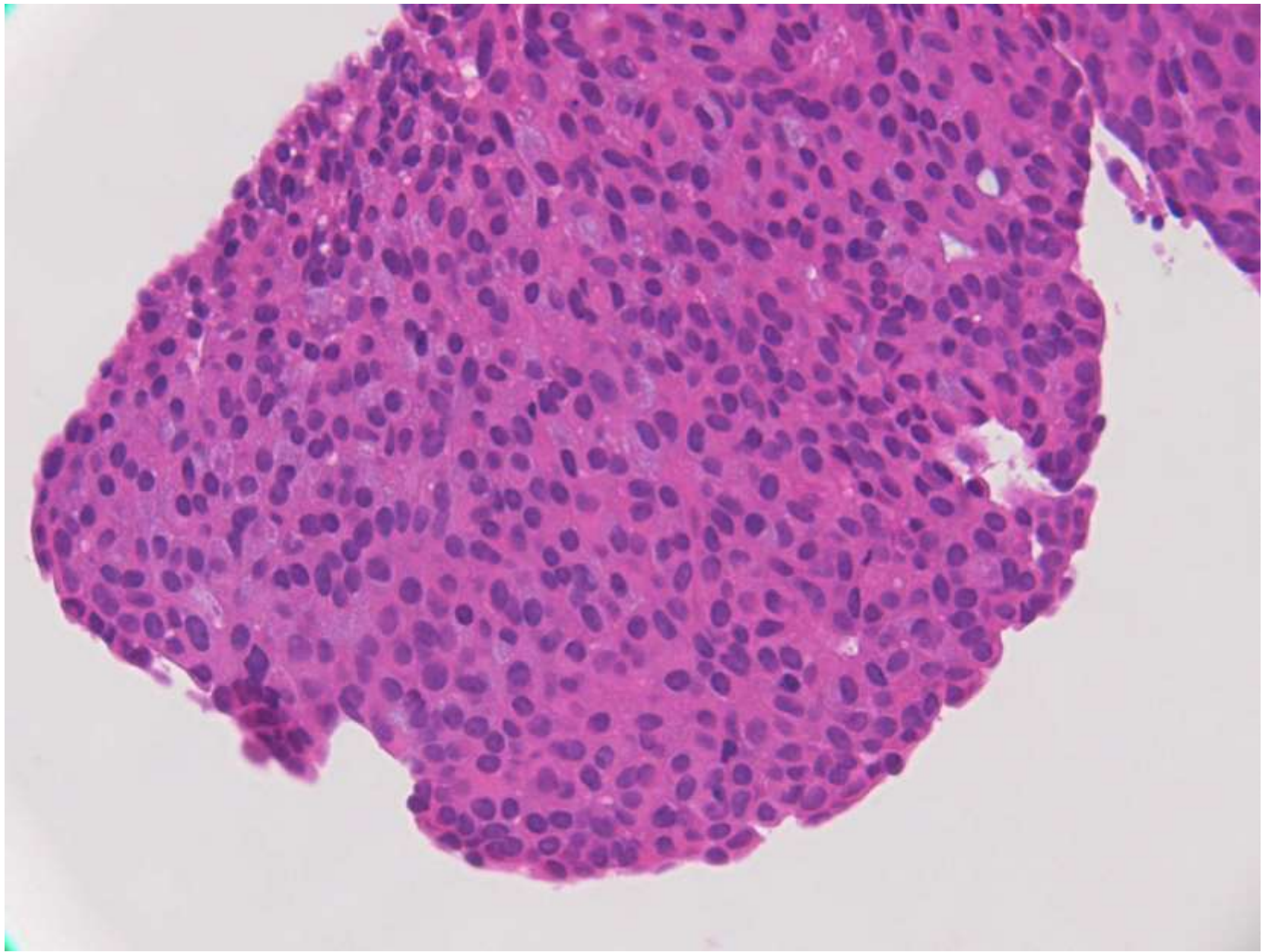






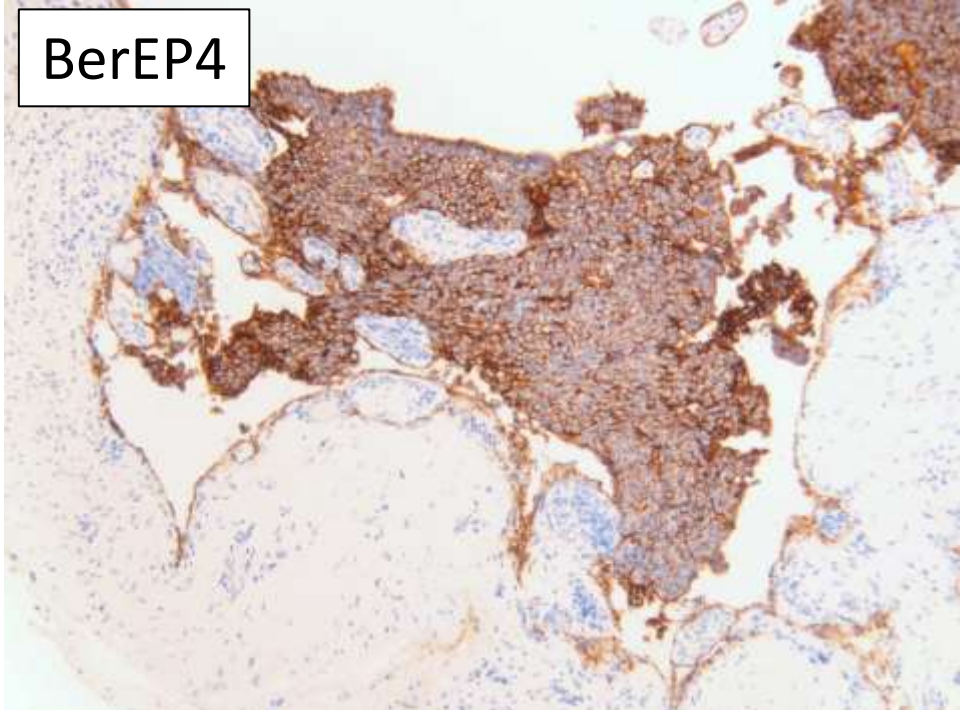




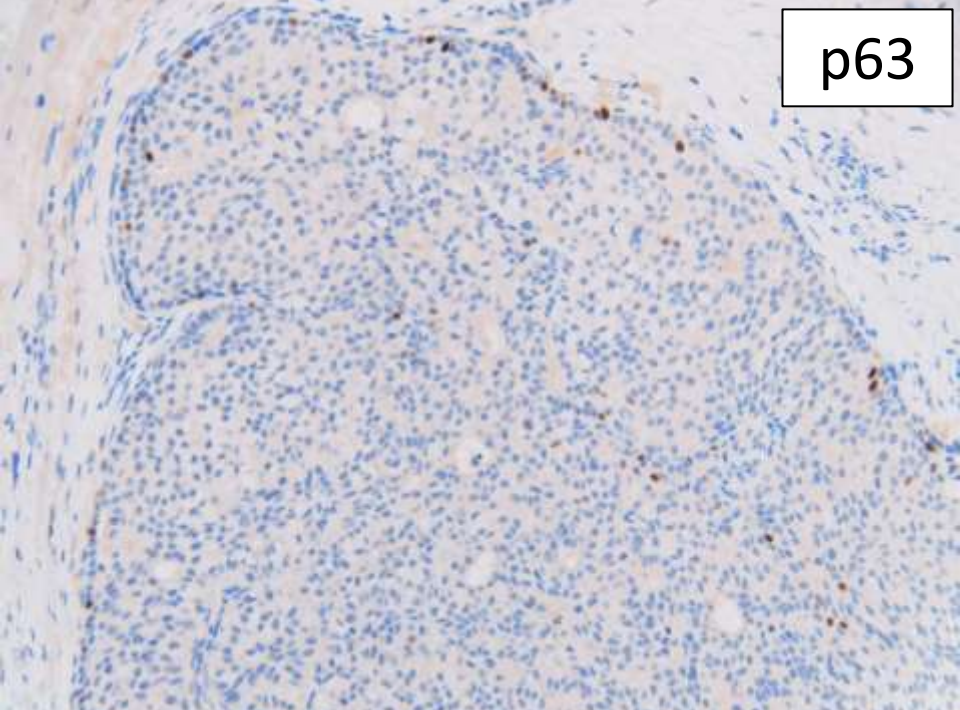




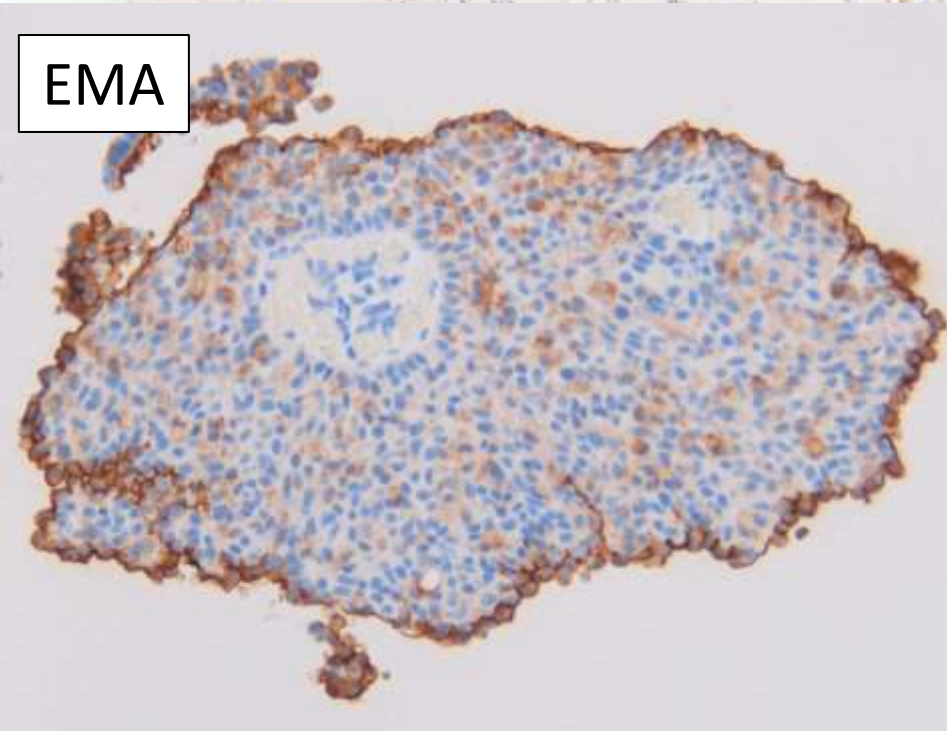
BerEP4



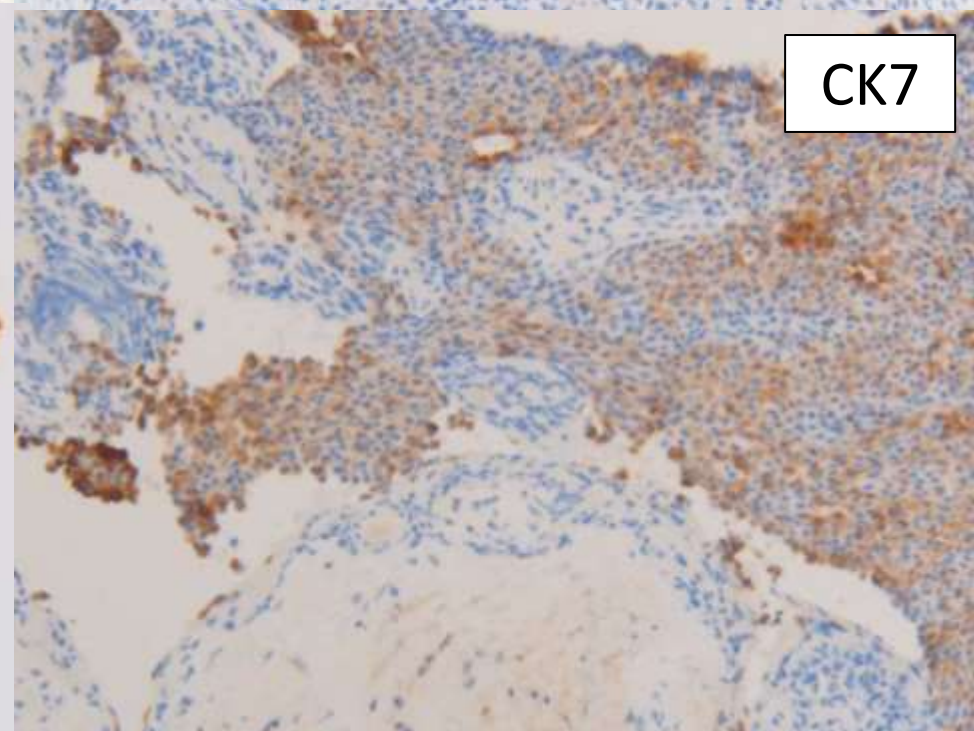
p63



EMA



CK7





Synaptophysin

ChromograninA

- My Diagnosis -

ER

Endocrine mucin producing adenocarcinoma

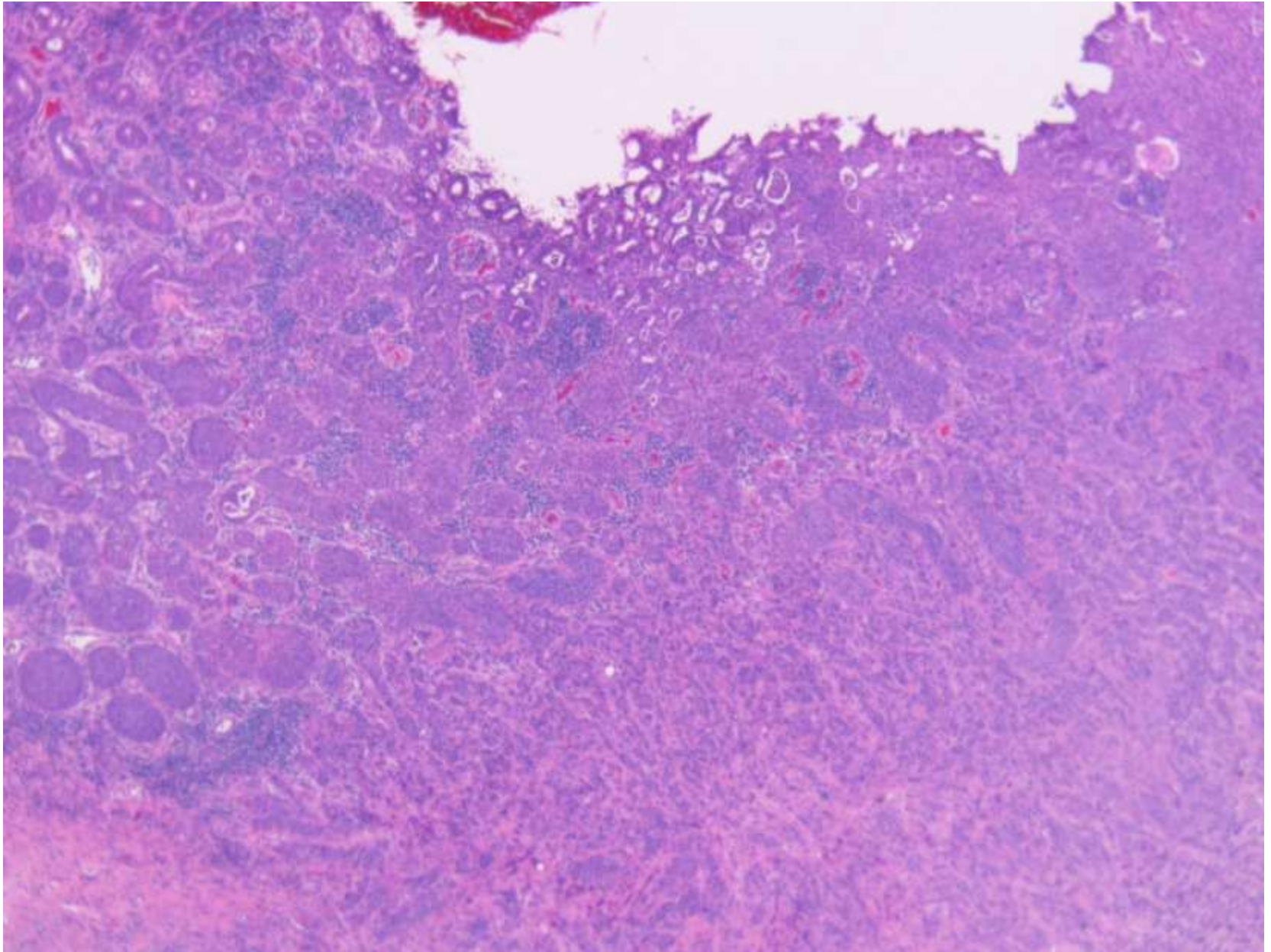
CCDFP15

*Advice: Mx as for Nodular BCC*

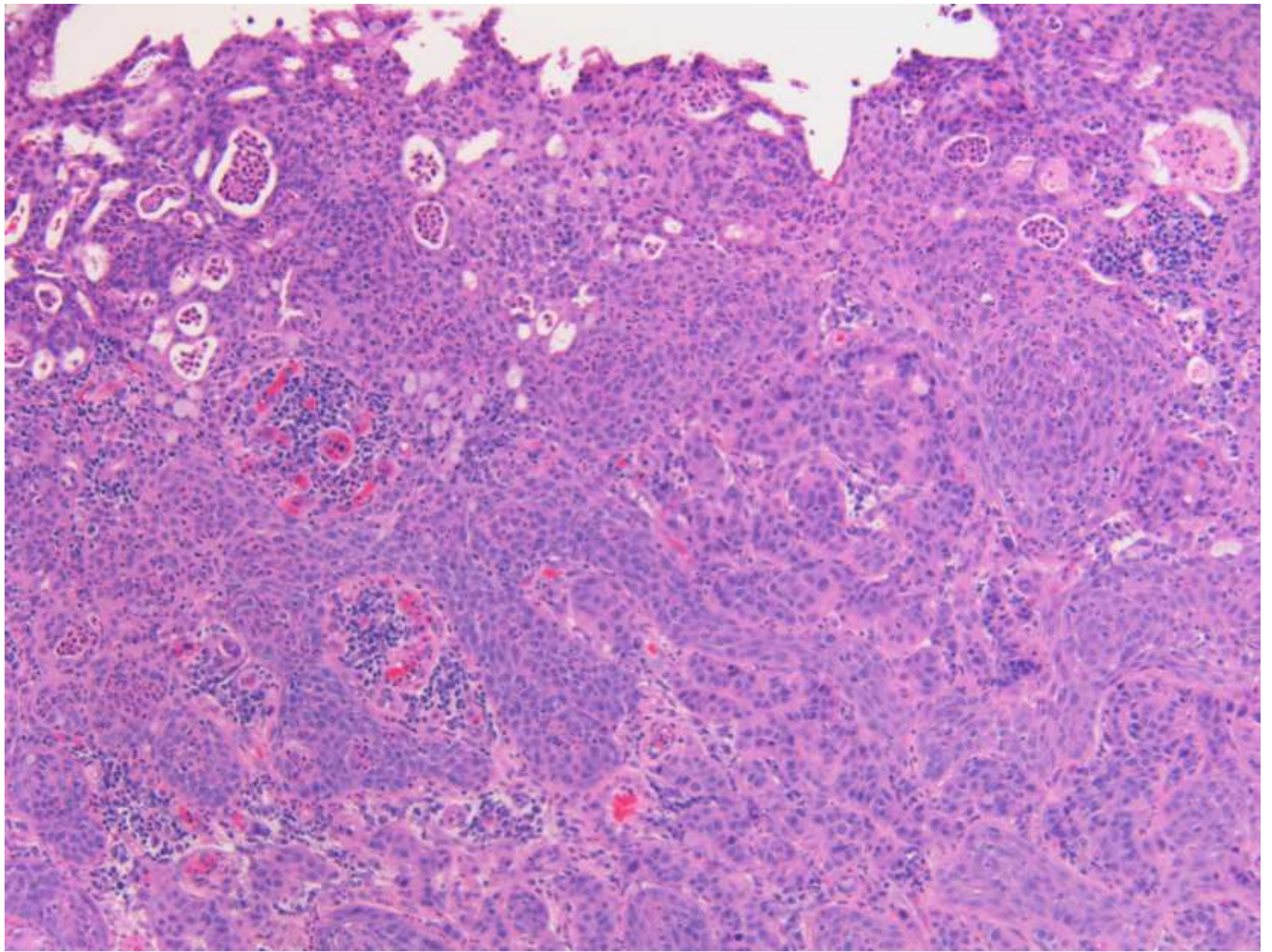


Next Case

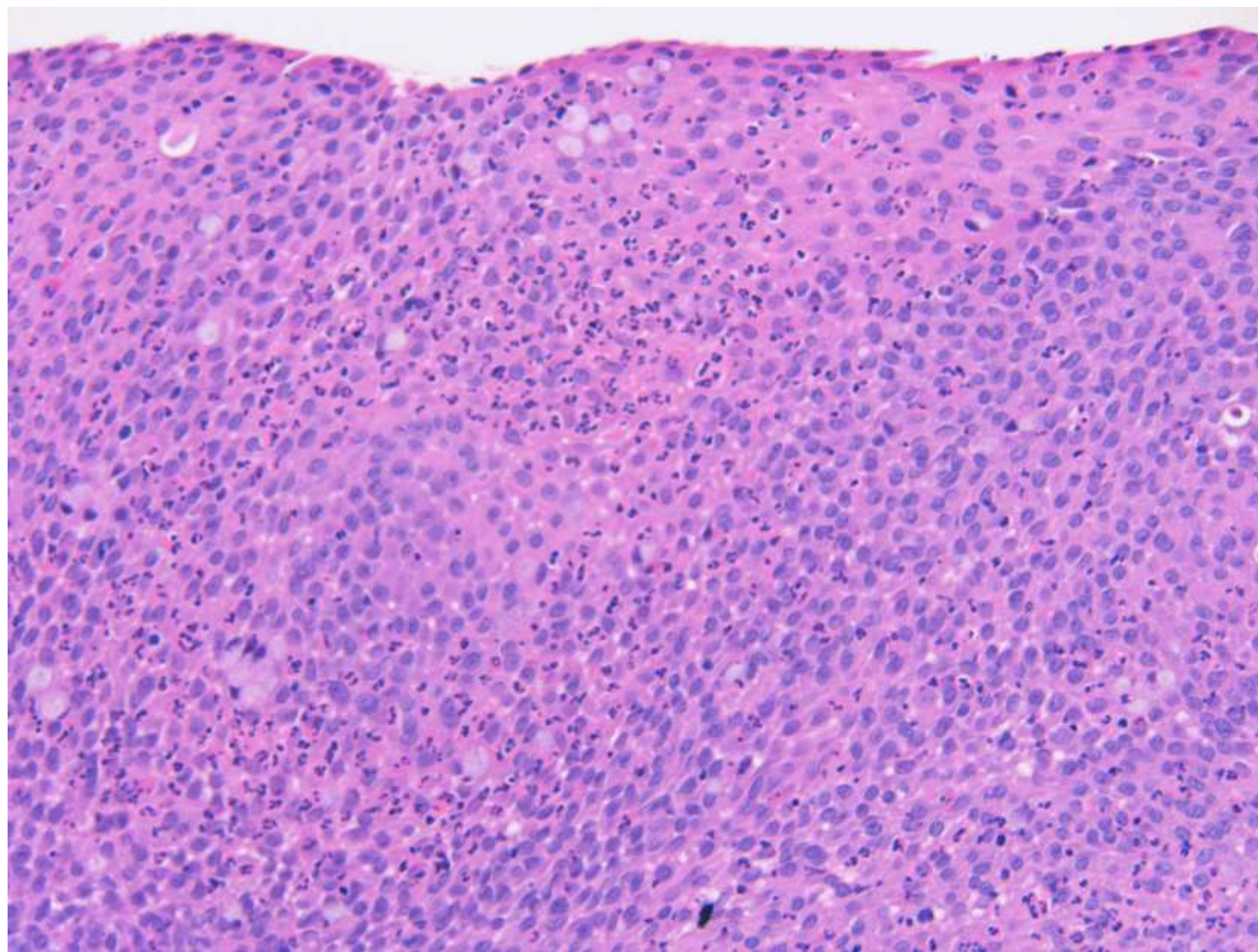
# M83 Right Upper Eyelid. Bladder cancer 2 years ago. ?secondary



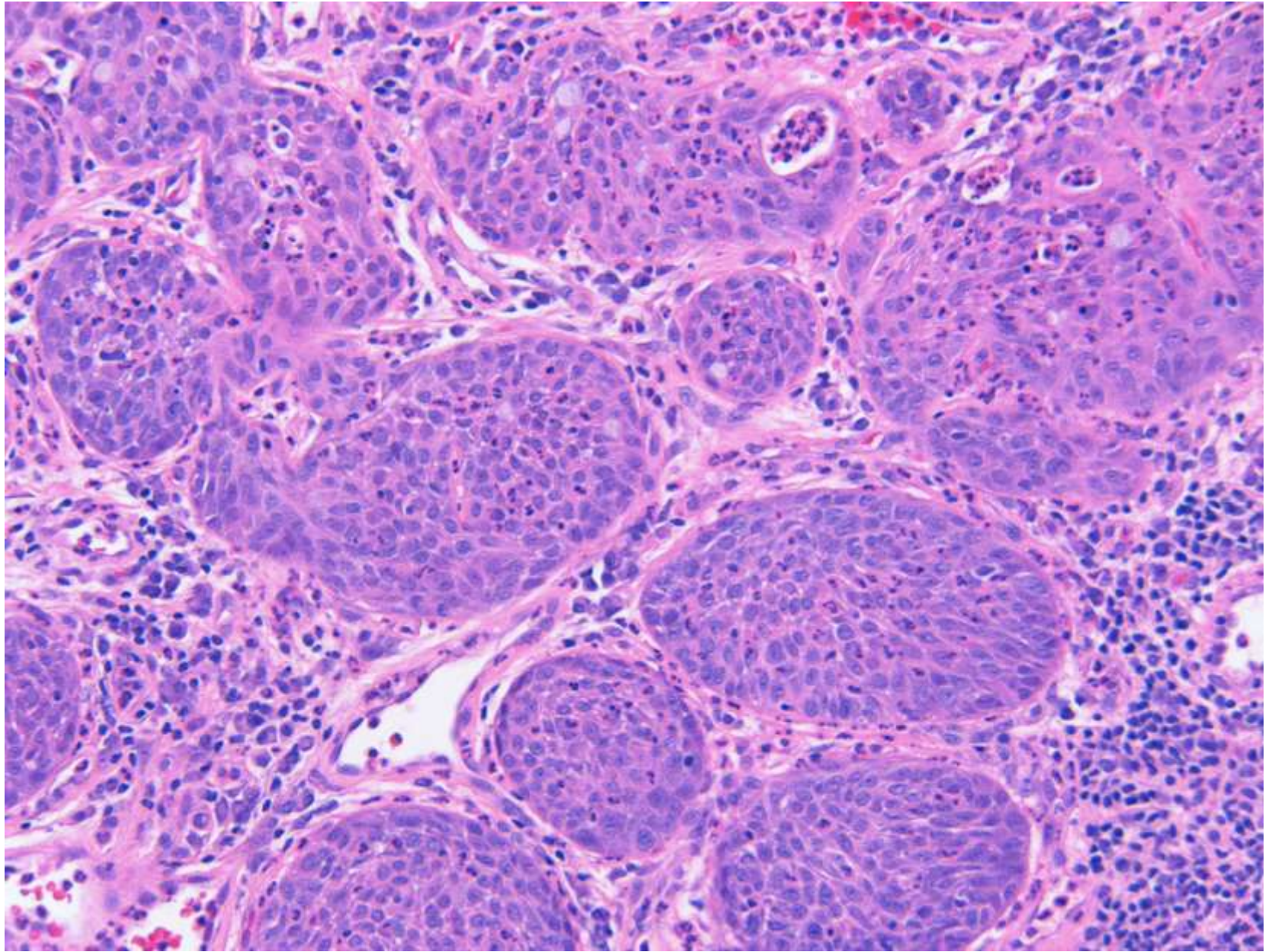




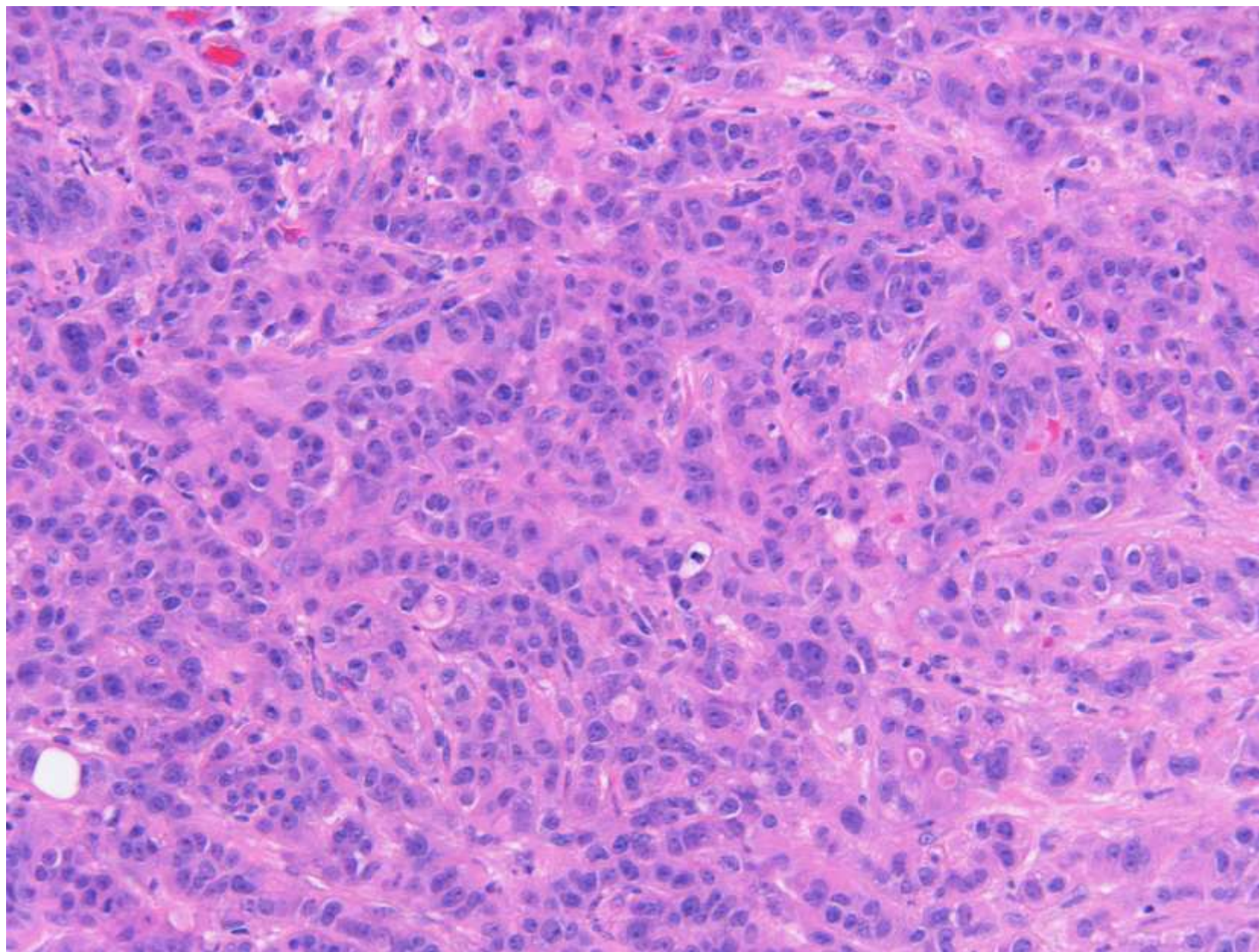




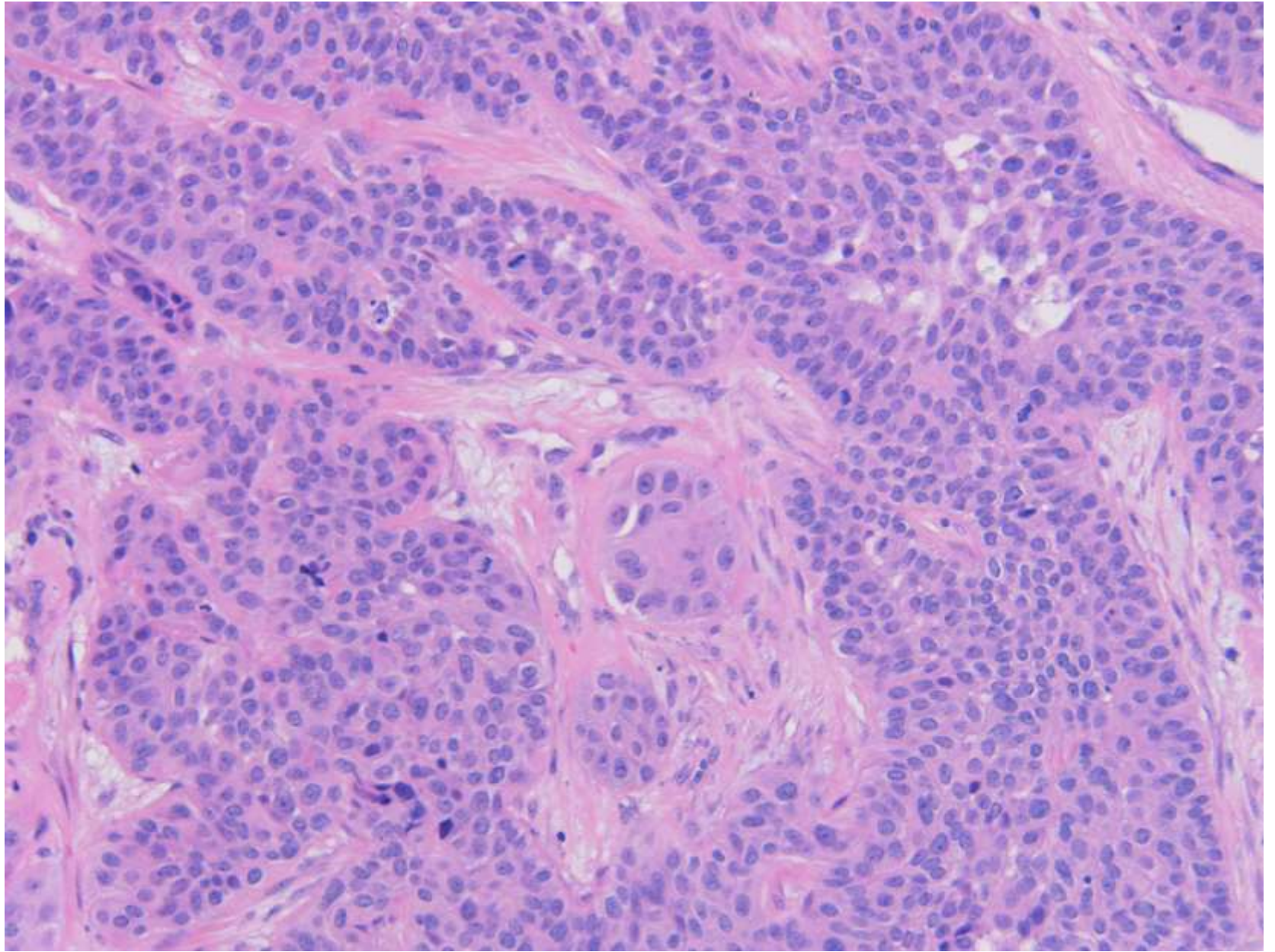








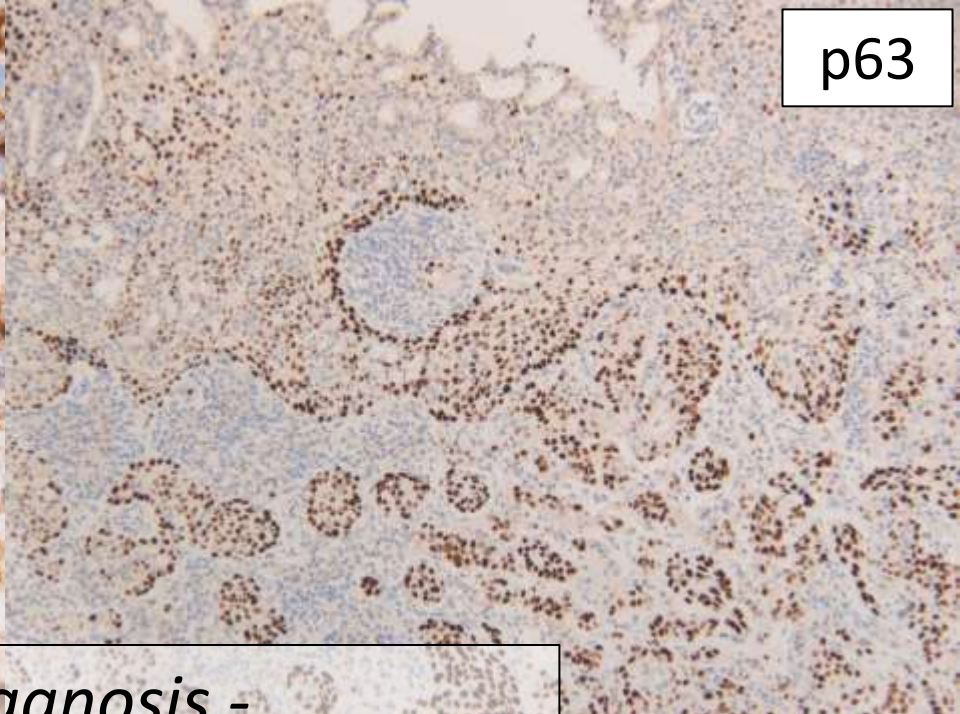
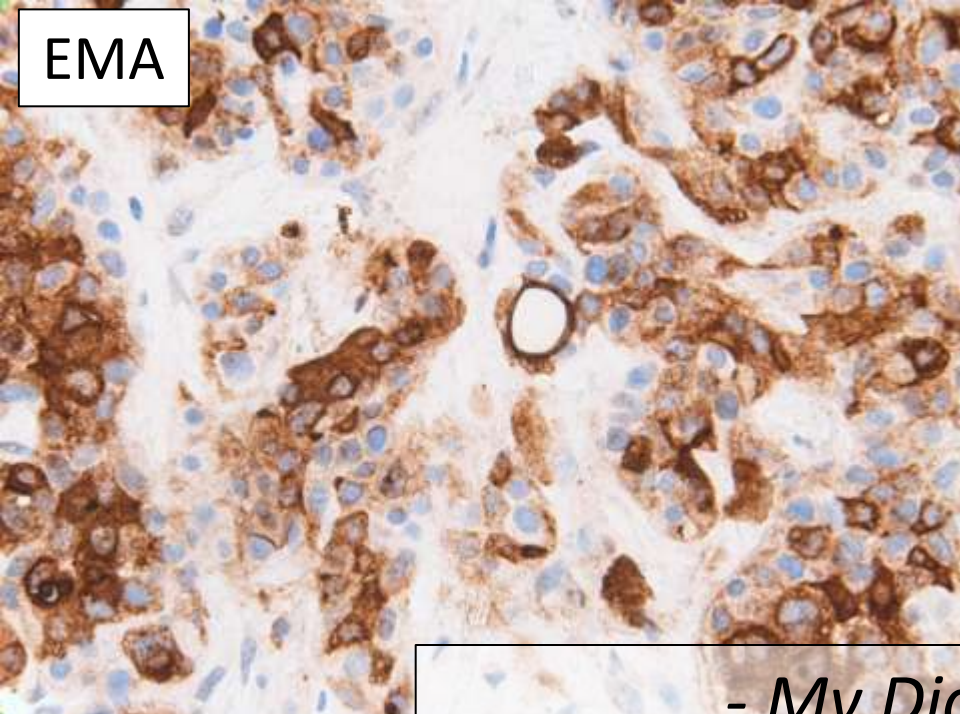






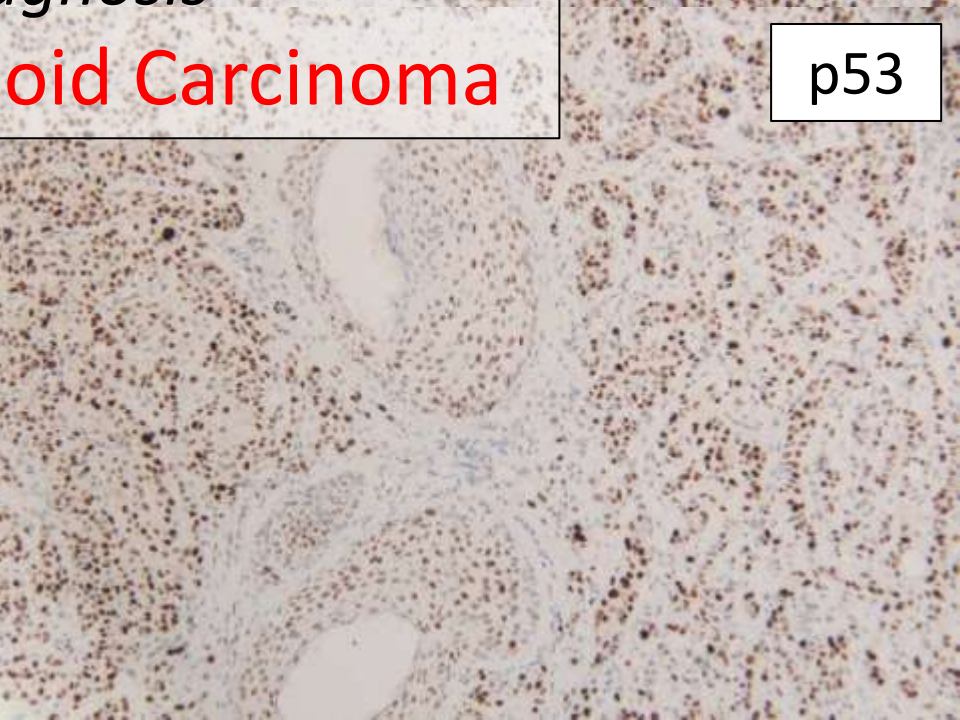
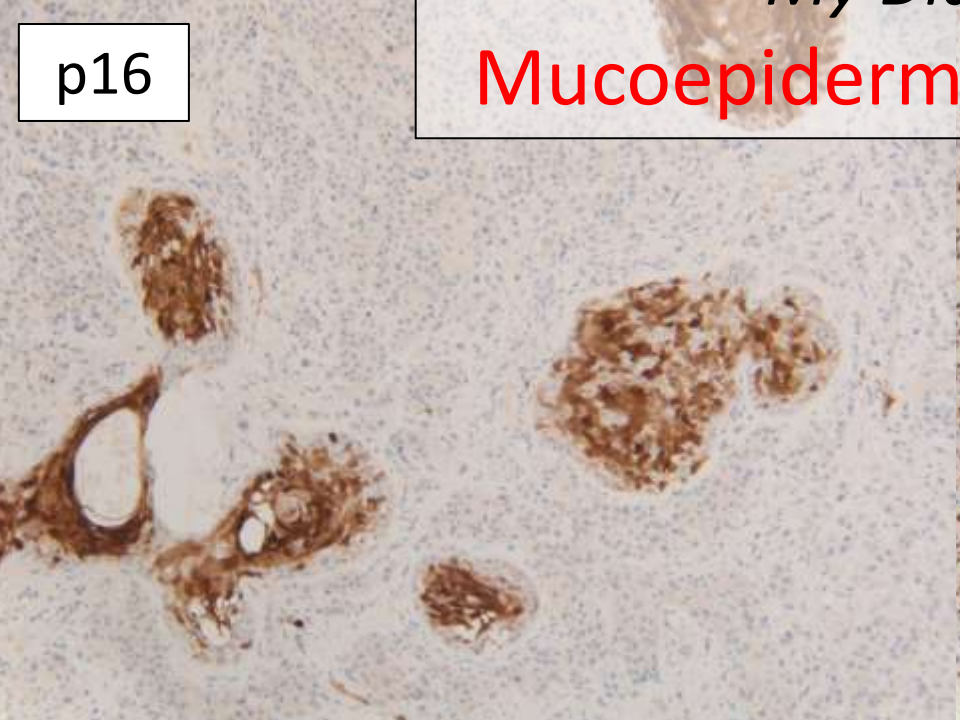
EMA

p63



p16

p53



- *My Diagnosis* -

**Mucoepidermoid Carcinoma**



## Learning Points:

Anatomic location can help make a diagnosis

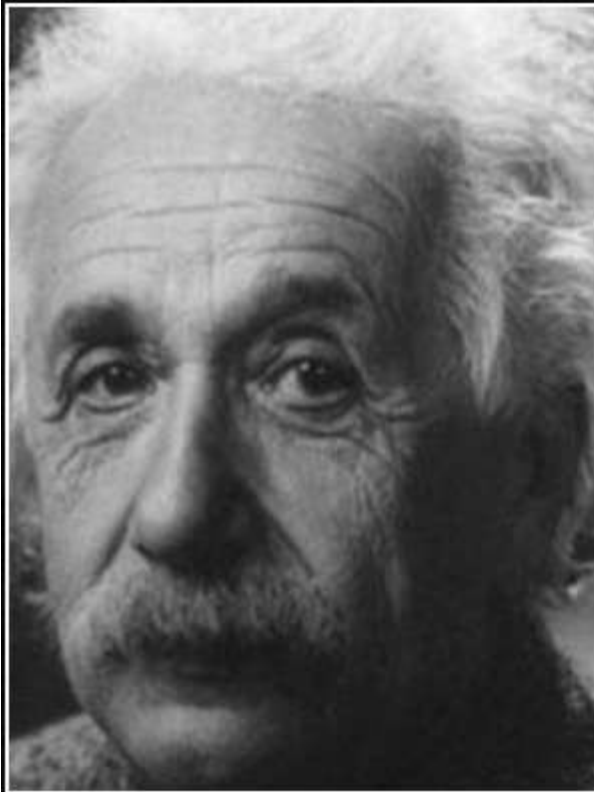
Completely circumscript tumours with malignant cytological features may be “in situ carcinoma”

Nearly all adnexal tumours that are mitotically active can grow large or undergo malignant transformation.

Most experts have seen cases of unexpected metastasis of apparently “benign” lesions

- In addition truly benign lesions e.g. hidradenomas may give rise to “benign” nodal metastases presumably due to implanation of tumour cells in to lymphatics at incomplete removal or biopsy

On this base I prefer to see adnexal tumours completely excised with clear margins.



I believe in intuitions and  
inspirations...I sometimes FEEL that I  
am right. I do not KNOW that I am.

— *Albert Einstein* —

AZ QUOTES



## **Acknowledgements: Part 2**

Ben Fletcher

Bruce Gee

Simon Tso

David Snead

Scott Sanders

Sixto Batitang

Dimitry Kazakov

Vivek Mudaliar

Veena Shinde

Nitin Khirwadkar