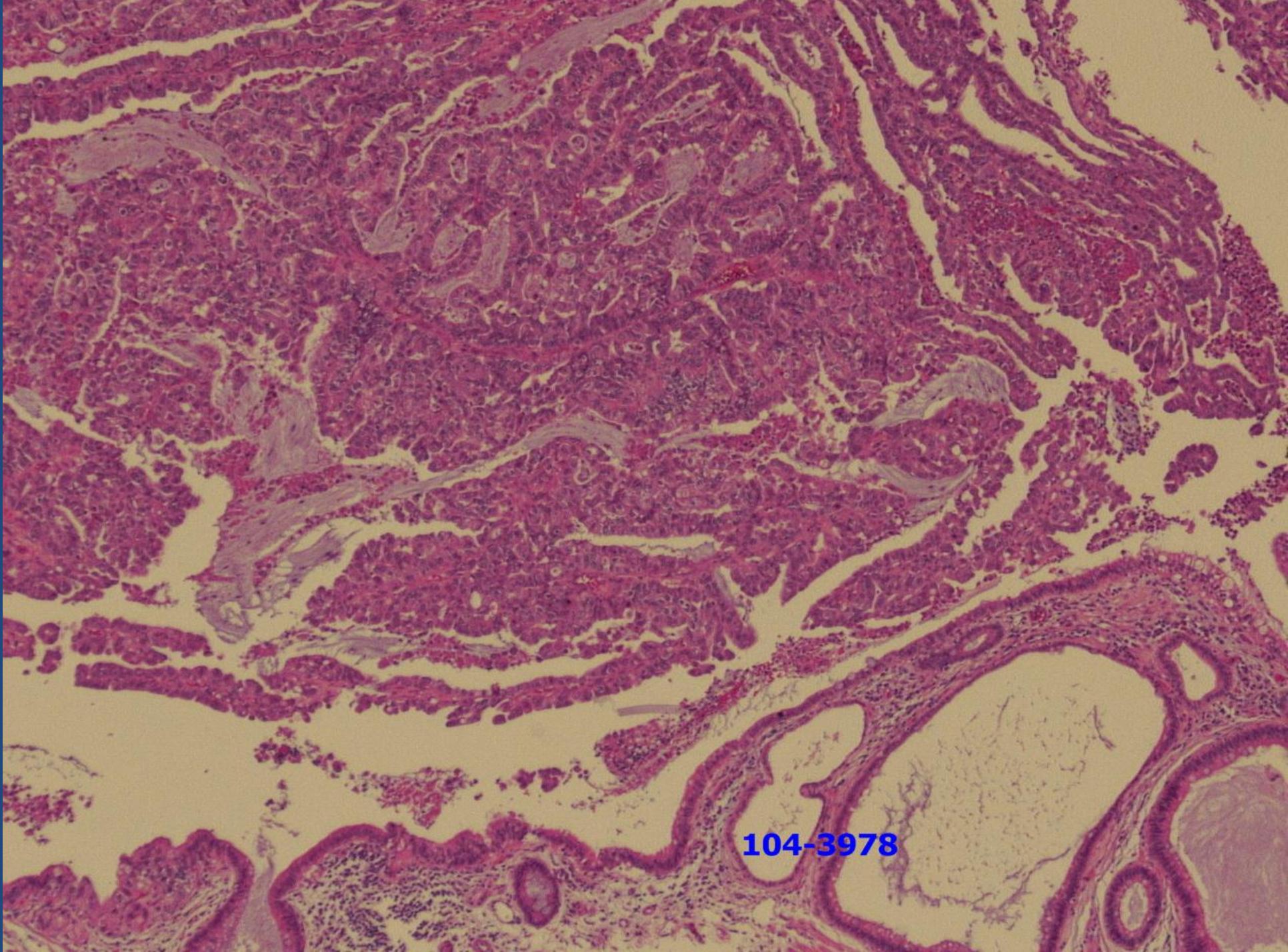
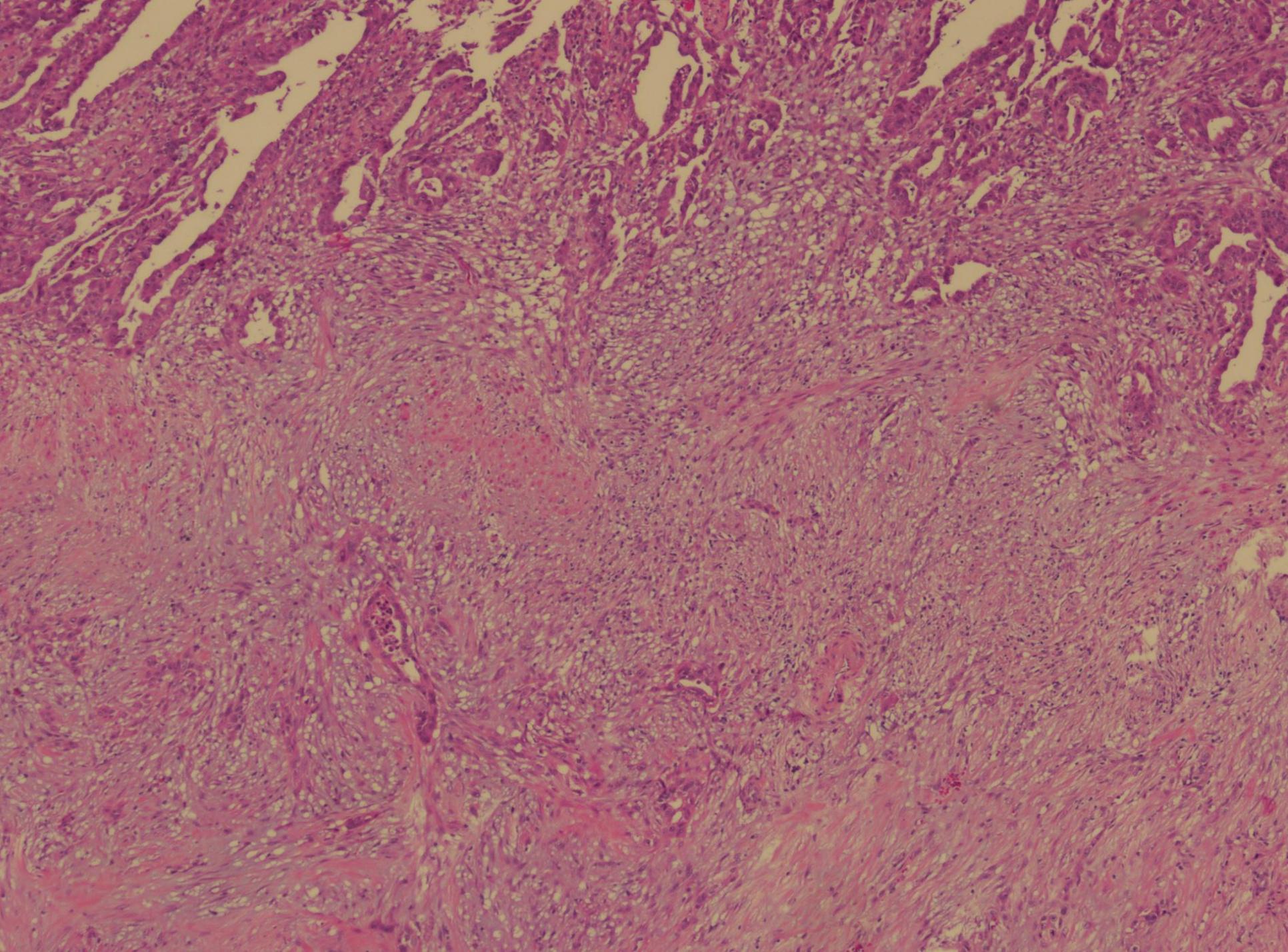


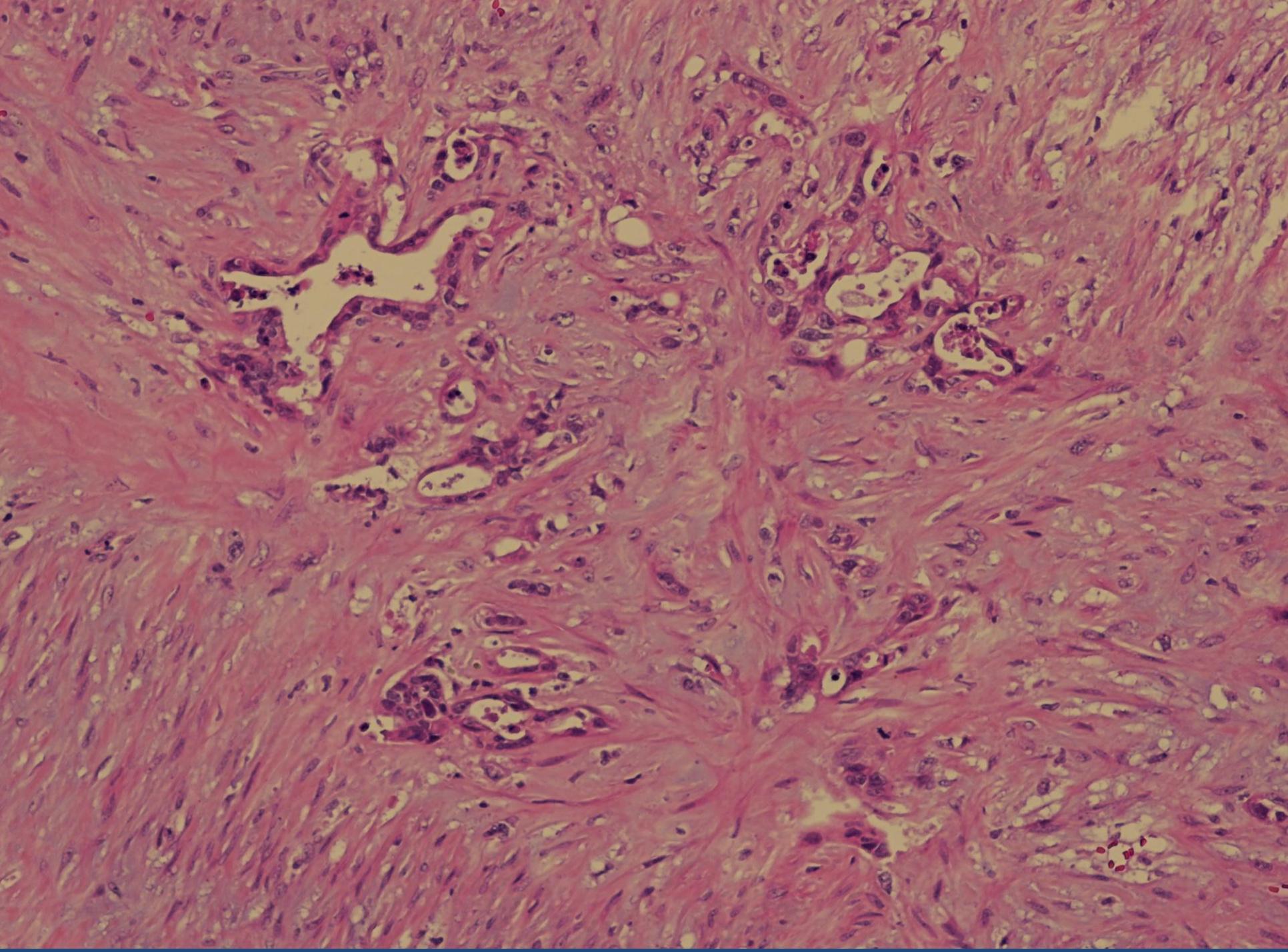
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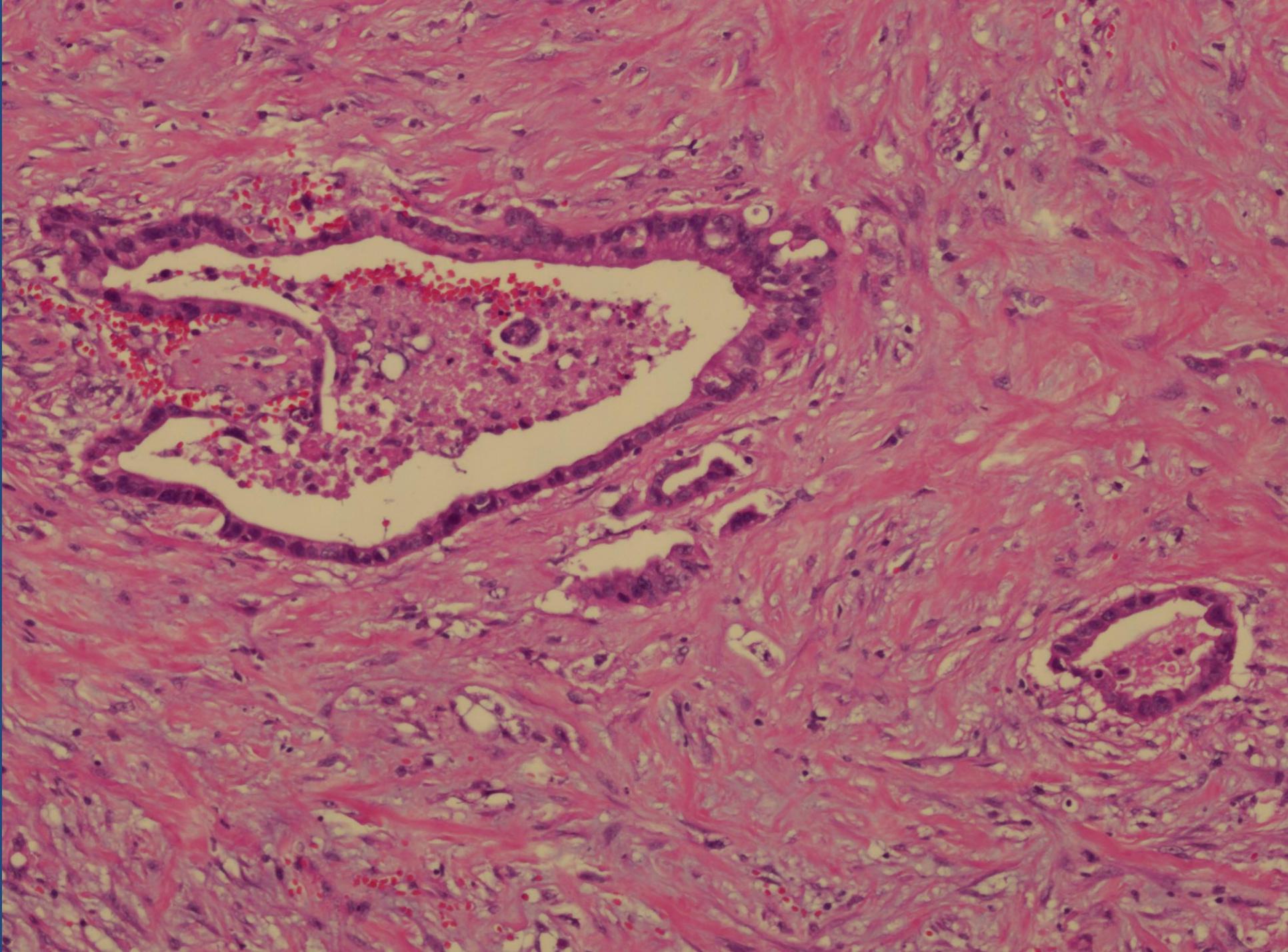
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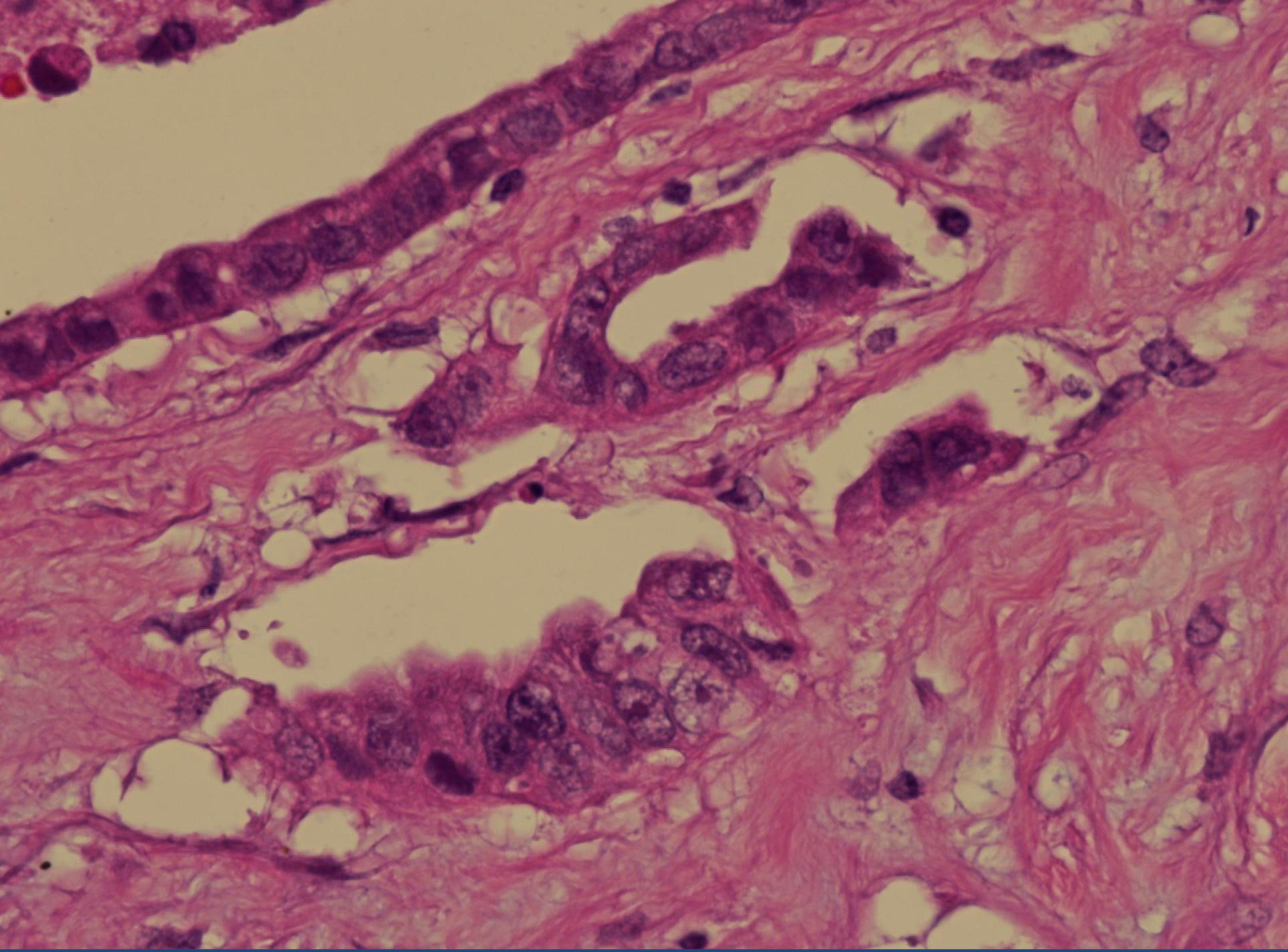


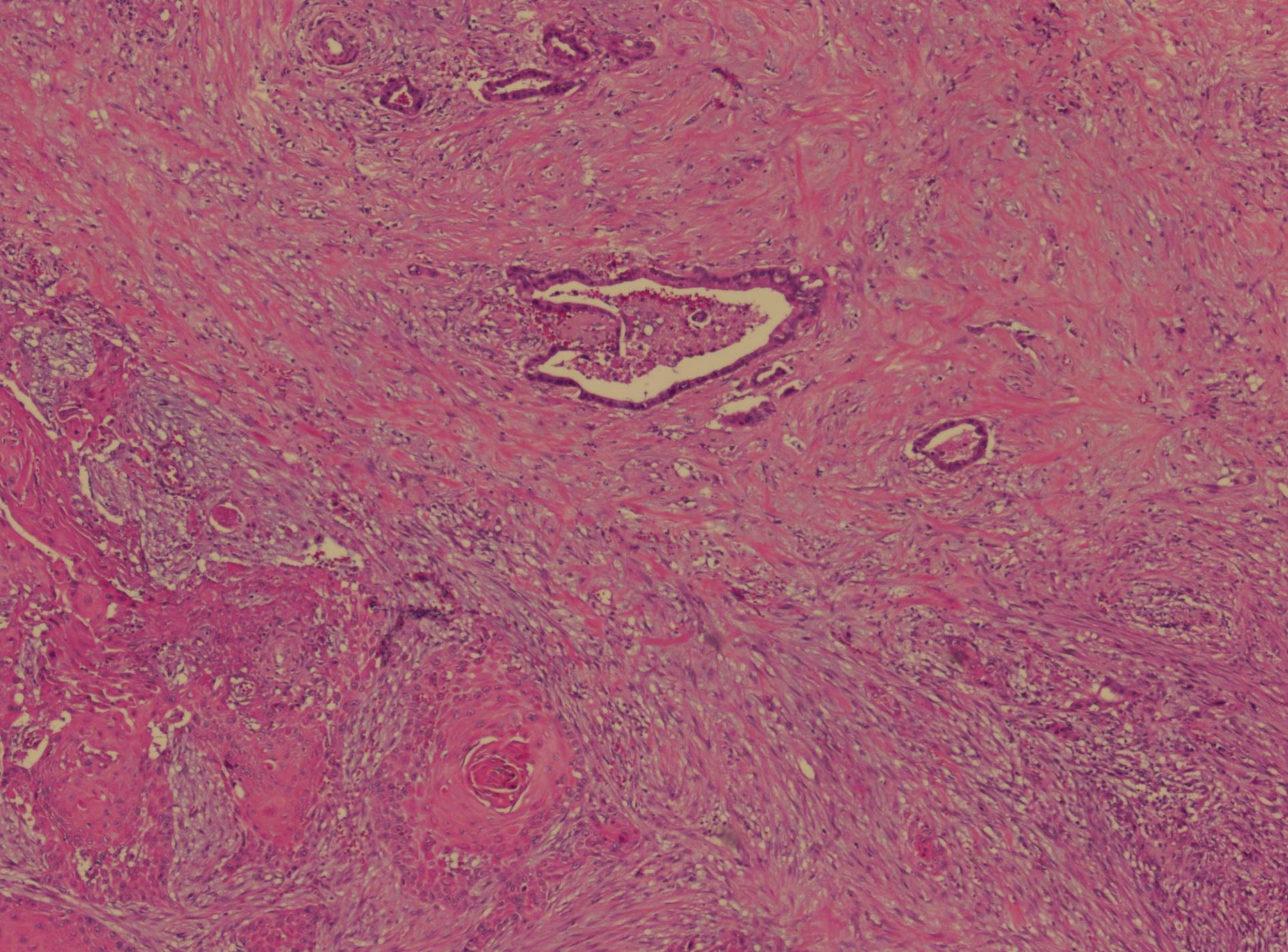
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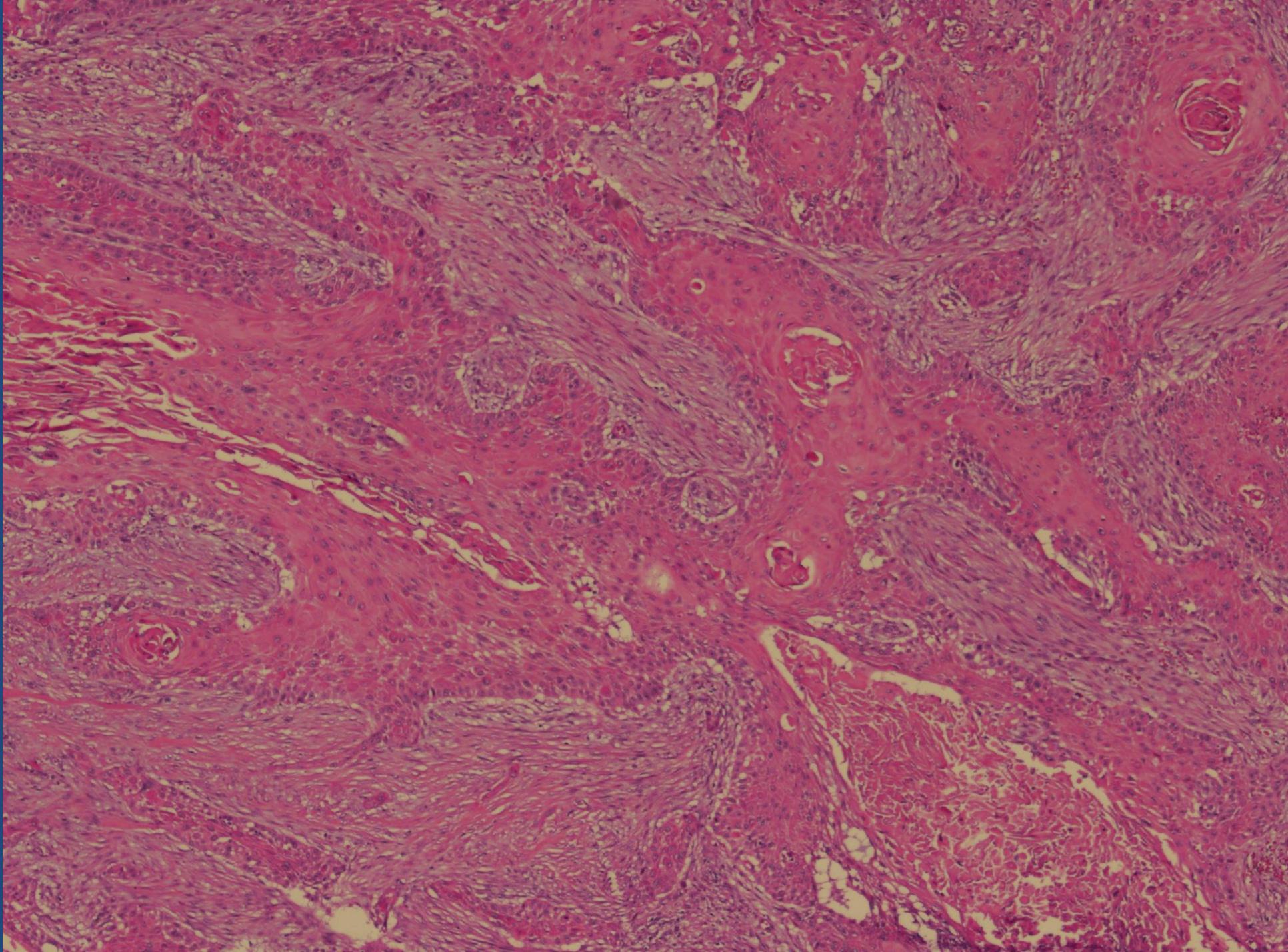


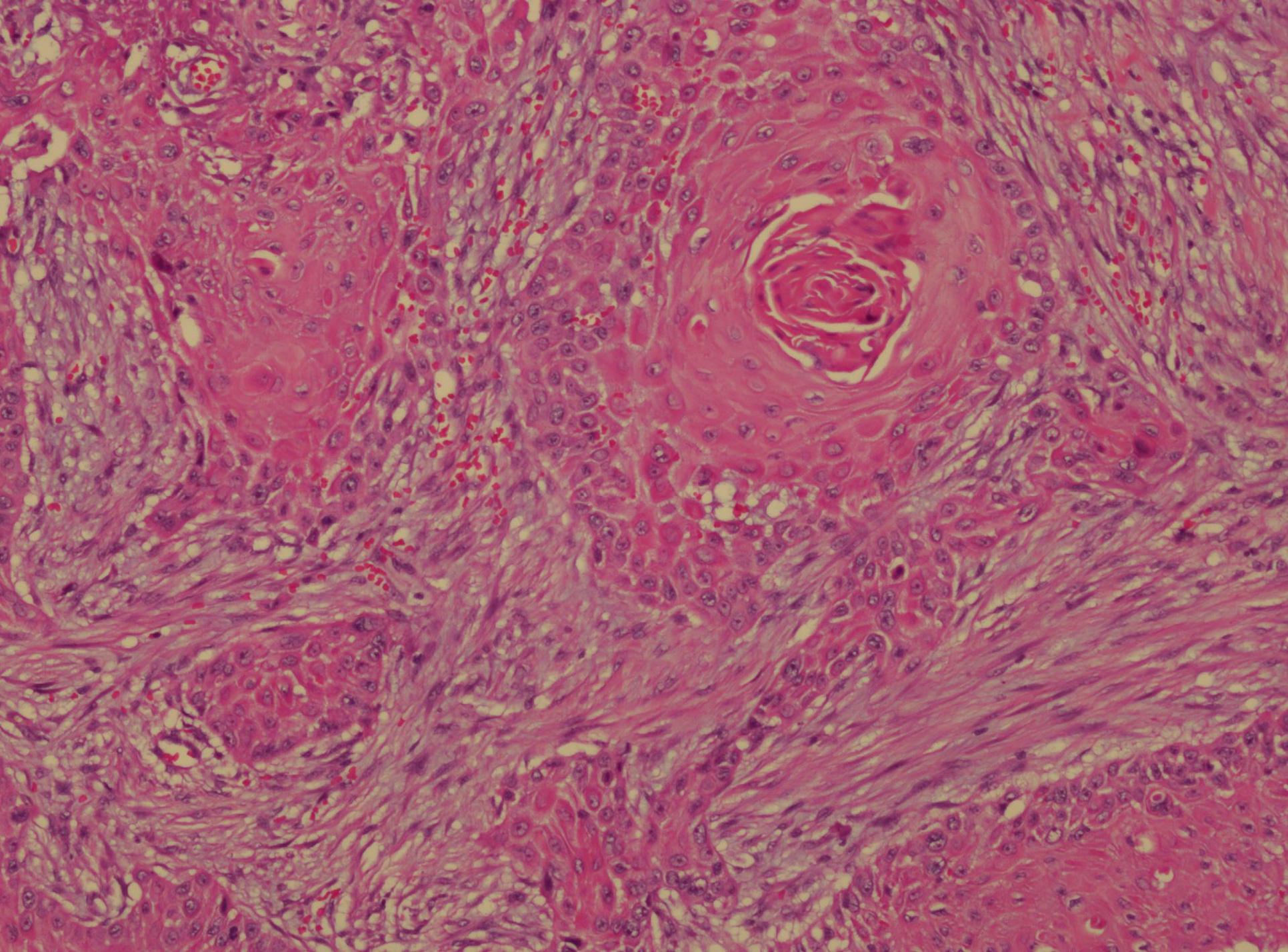


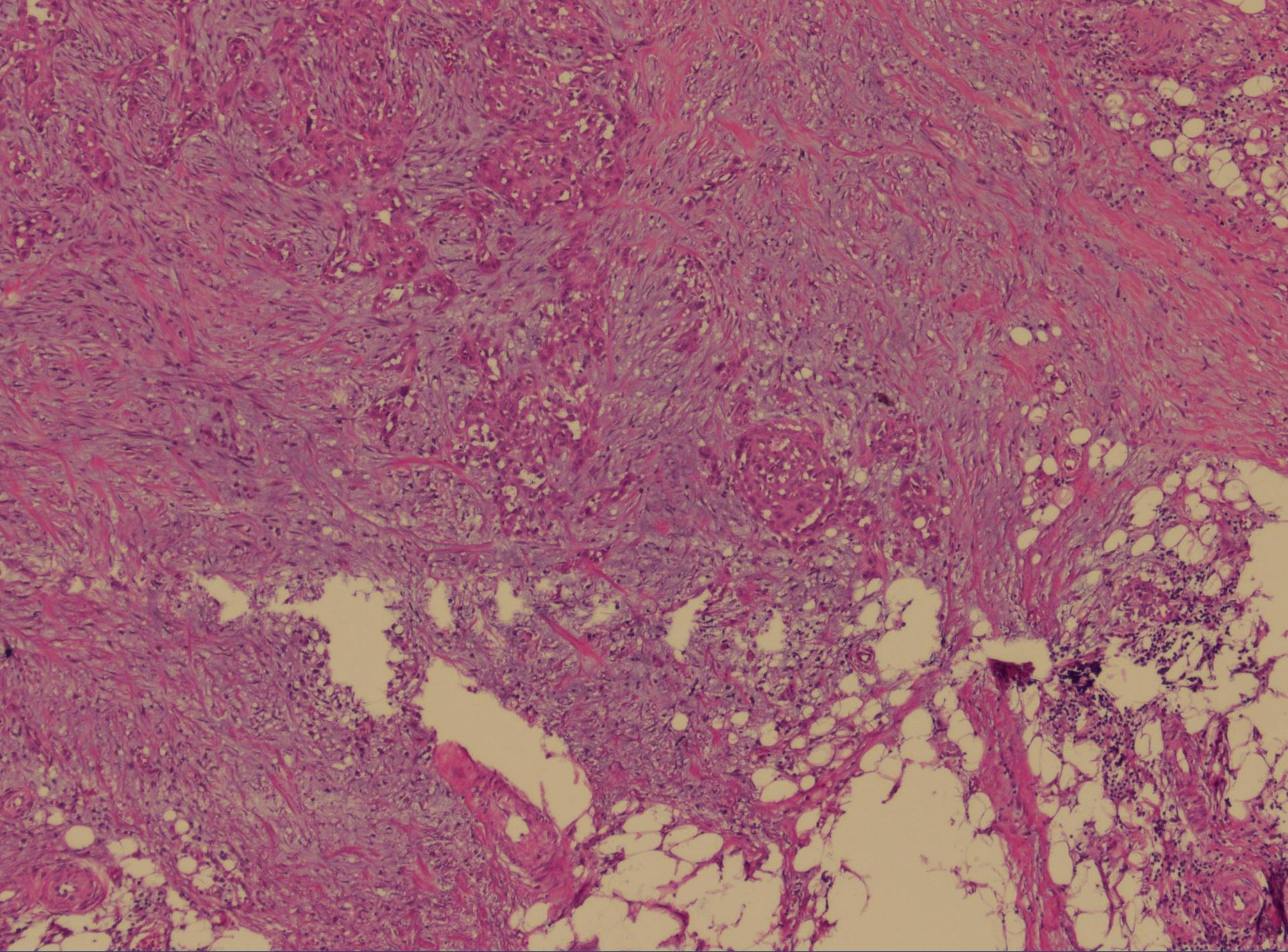


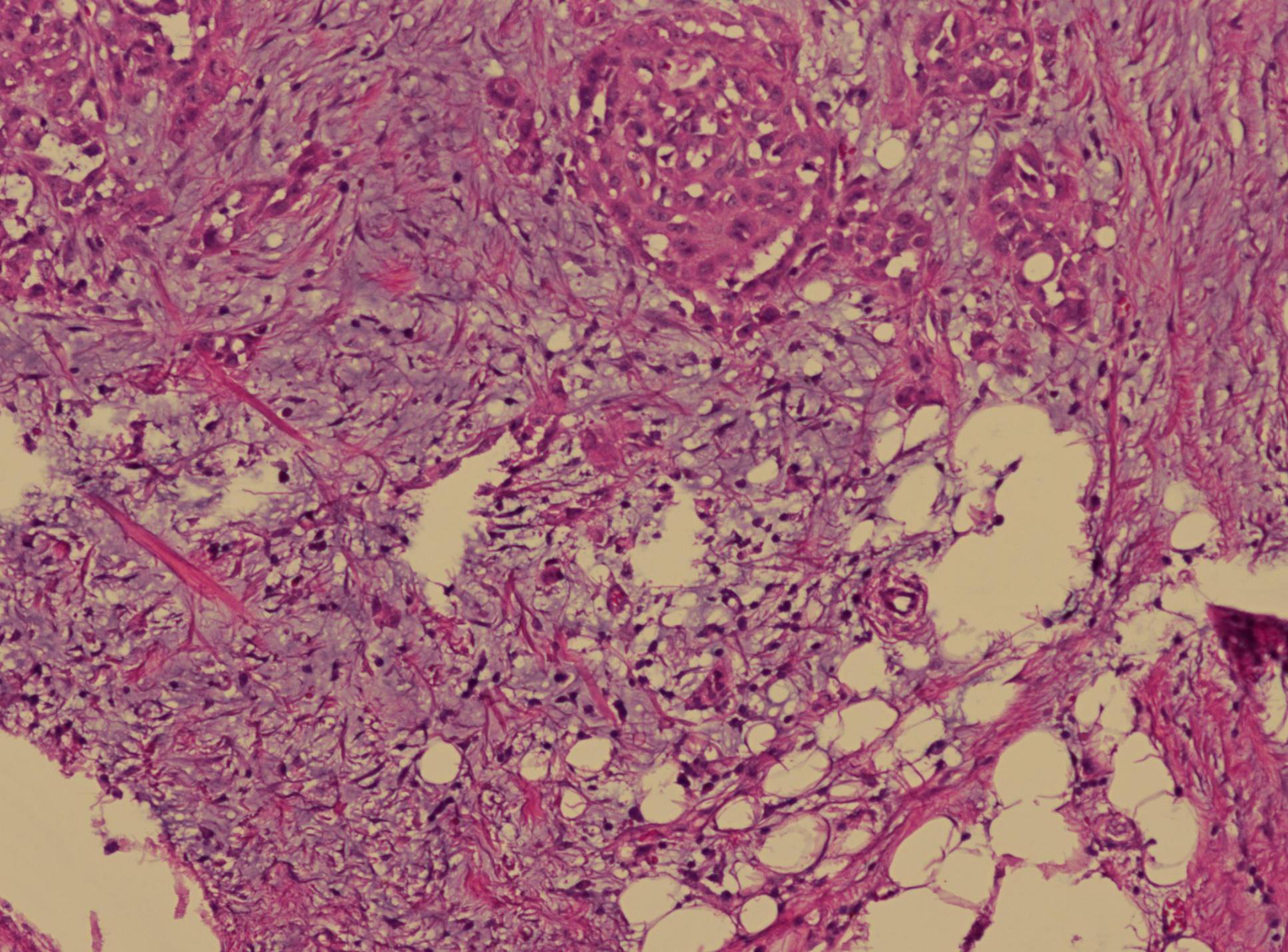


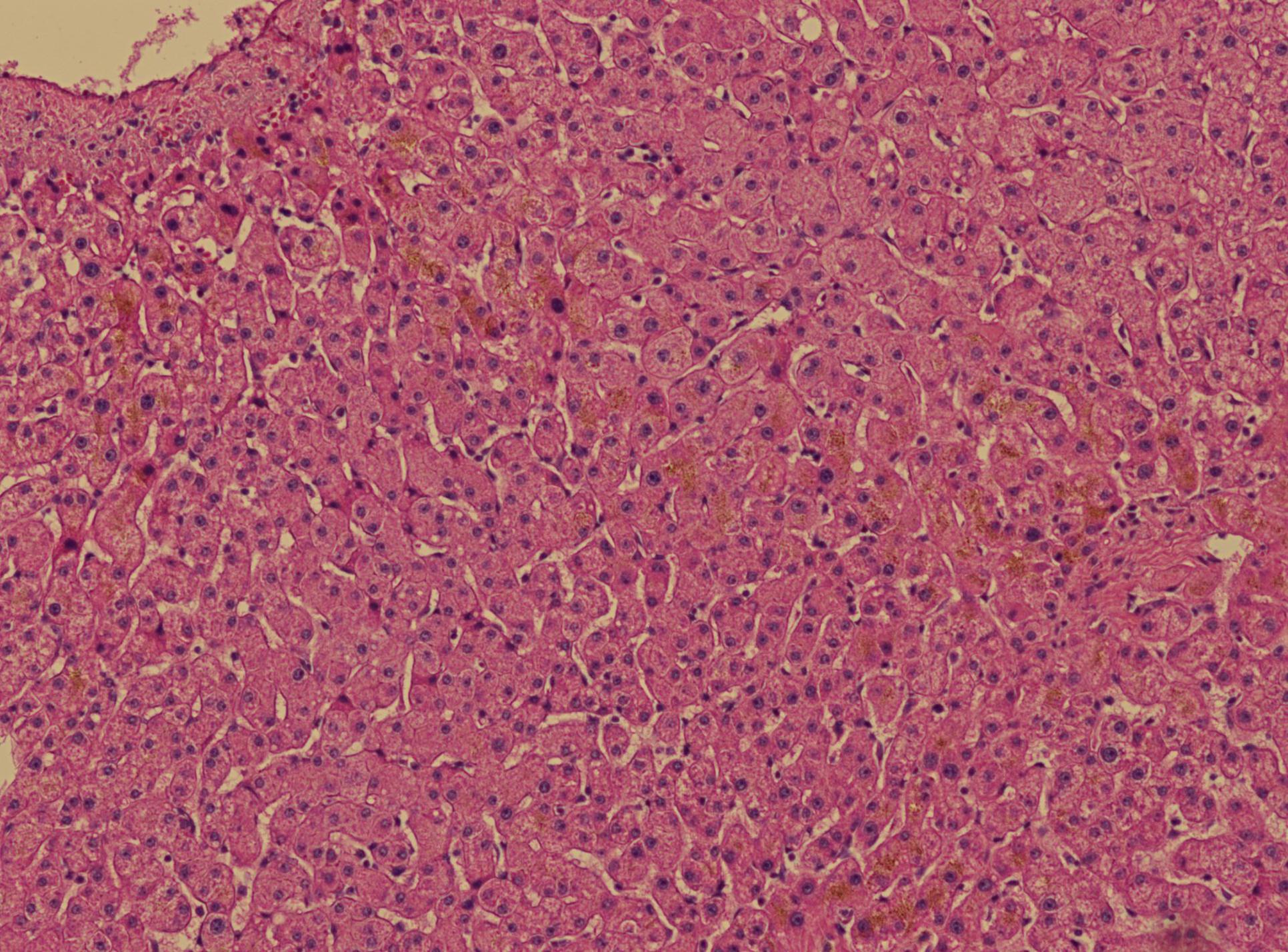


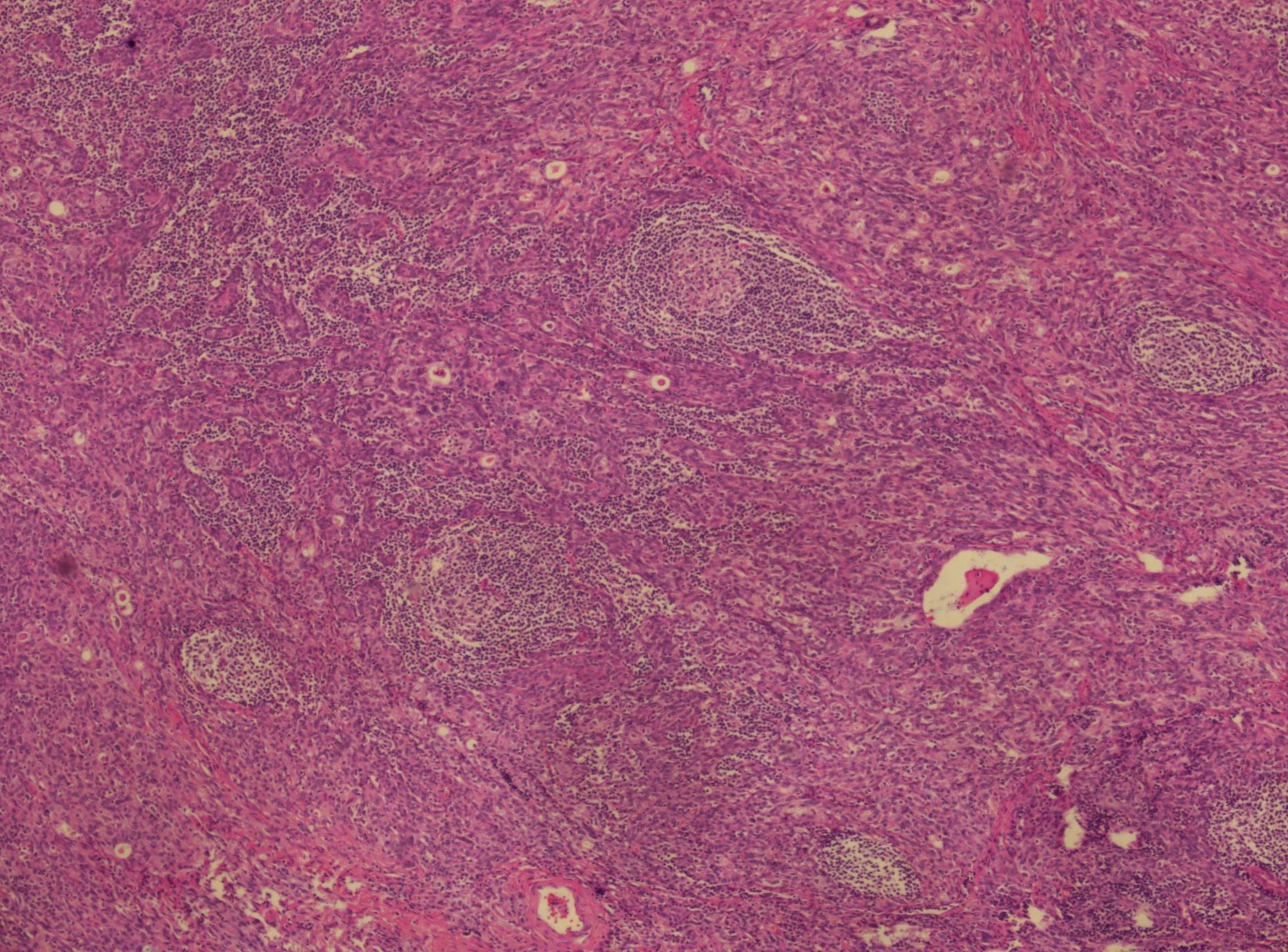


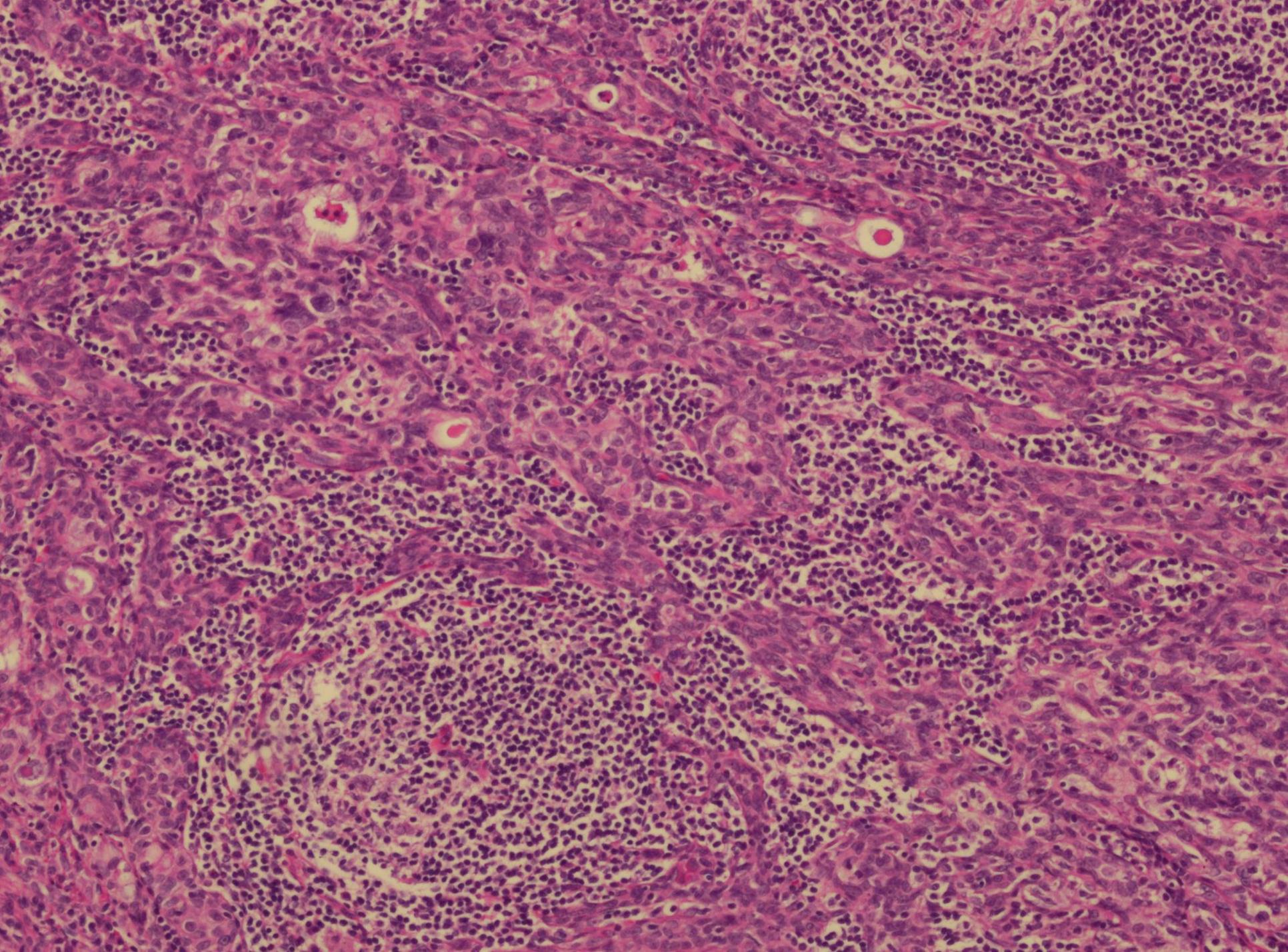


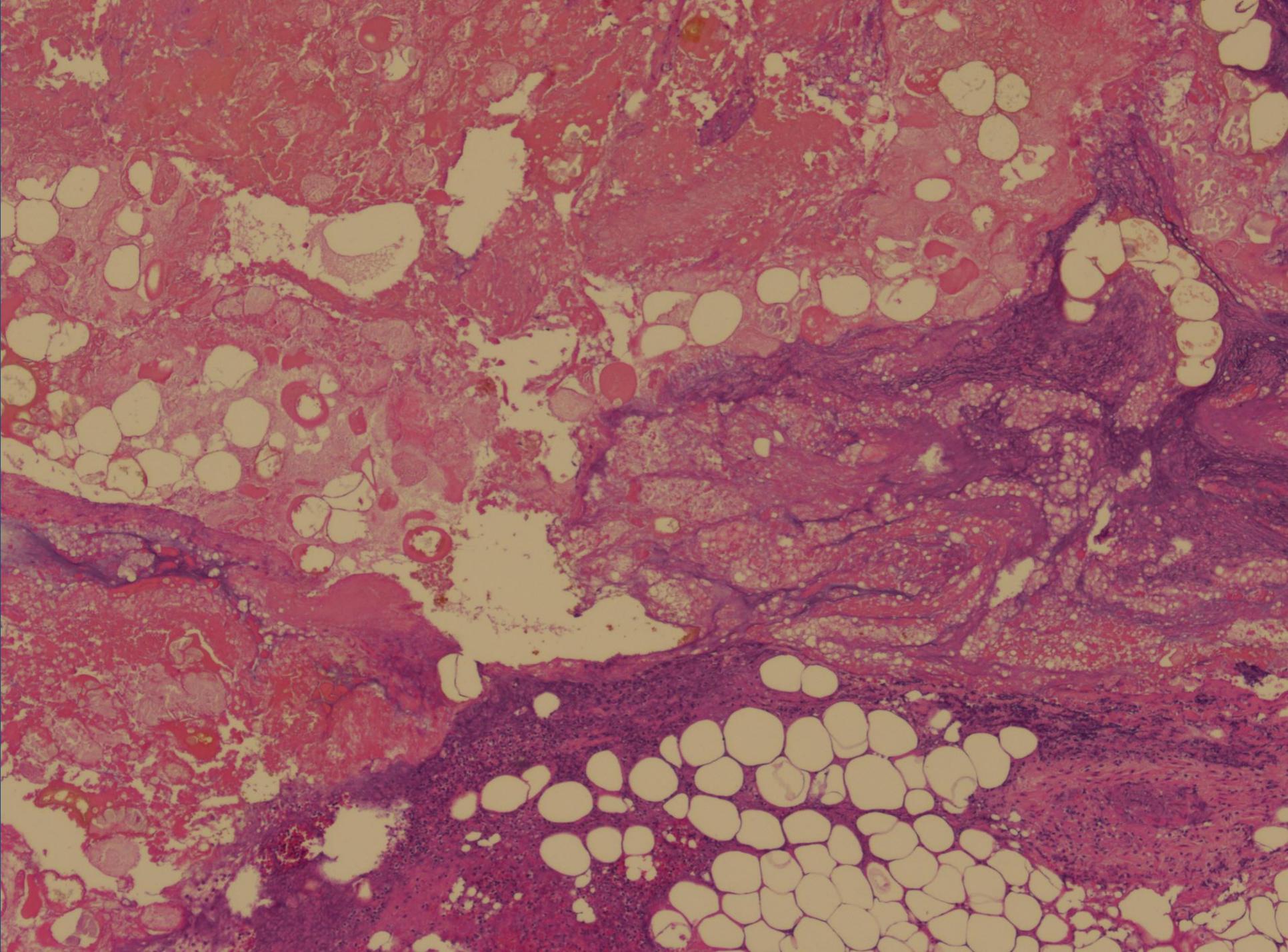


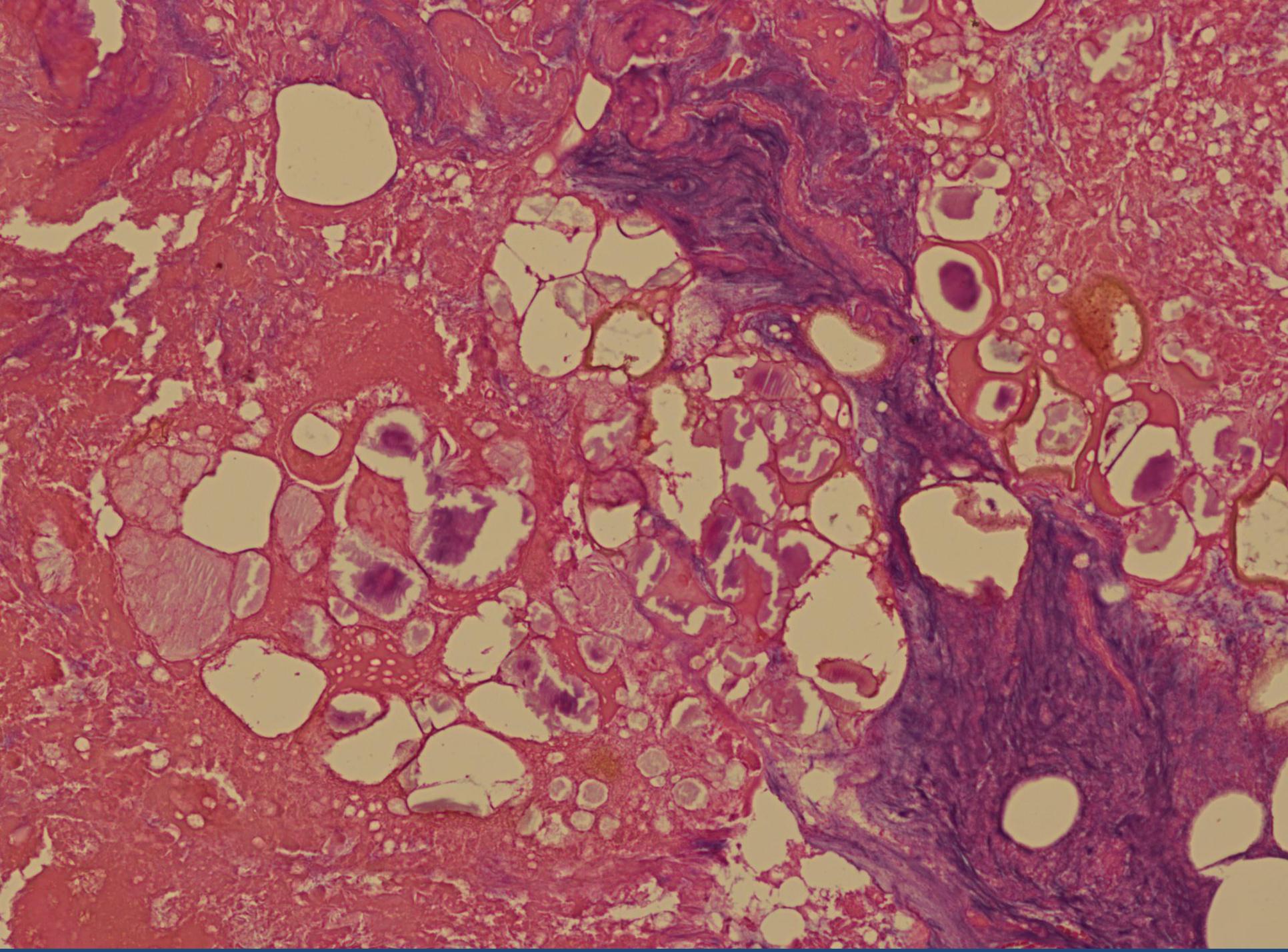












Adenosquamous Carcinoma of Gallbladder

Gallbladder Carcinoma

- Relatively uncommon
- Age 60+ years (mean 72 years), usually not resectable
- 90% are adenocarcinoma, 5% squamous cell or adenosquamous, 5% undifferentiated

Epidemiology

- 2.5 per 100,000 population
- F : M = 1.77: 1
- Lower incidence in Asia, where pyogenic and parasitic disease of biliary tree are more common
- More common in American Indians and Hispanics; very rare in blacks
- Approximately 5000 new cases are discovered each year in US

Etiology

- Gallstones
- Diffuse calcification of the GB wall (porcelain gallbladder)
- Abnormal junction between CBD and pancreatic duct (abnormal choledochopancreatic junction)
- Familial adenomatous polyposis (FAP)
- Ulcerative colitis
- Primary sclerosing cholangitis

- Squamous metaplasia of GB mucosa

Clinical features

- Currently, close to 50% of GB carcinoma are diagnosed incidentally in cholecystectomy specimens from pts with s/s attributed only to the presence of gallstones.
- Unfortunately, gallbladder carcinoma usually presents at a late stage, even when found incidentally
- Abdominal pain, jaundice, weight loss

Gross

- Infiltrating grey-white mass
- Most arise in GB fundus (60%) as nodular masses or diffusely involve the GB
- Body (30%), neck (10%)
- The GB may be distended by the tumor, or collapsed owing to obstruction of the neck or cystic duct

Micro of Adenosquamous Ca

- The extent of differentiation of the two malignant components, glandular and squamous, varies, but in general they tend to be moderately-differentiated
- Keratin pearls are often present in the squamous component
- Mucin is usually demonstrable in the neoplastic glands.

D.D.

- Cholecystitis with reactive atypia
- Carcinoma in situ colonizing Rokitansky-Aschoff sinuses (nonrounded, irregularly placed gland contours, single cells, and the presence of desmoplasia favor invasive ca)

Prognosis

- GB ca tend to present at an advanced stage; 70% involve liver at diagnosis; 50% involve regional lymph nodes
- Overall prognosis is poor, with 5% to 10% survival at 5 years
- **5 year survival:**
 - Overall 1% ([J Surg Oncol 2008;98:485](#))
 - 85-100% for T1, 30-40% for T2
 - Median survival 6 months

Prognostic factors

- Favorable: papillary histology, low stage
- Unfavorable: small cell or undifferentiated types, angiolymphatic invasion, poorly differentiated, high stage, tumor budding and dedifferentiation

Prognostic factors

Molecular markers, poor prognosis:

- Overexpression of MCM2 or loss of expression of Tat-interacting protein 30 ([Hum Pathol 2011;42:1676](#)), overexpression of PEG10 and TSG101 ([Pathol Oncol Res 2011;17:859](#)), reduced expression of Raf-1 kinase inhibitory protein ([Hum Pathol 2010;41:1609](#))
- L1 adhesion molecule ([Hum Pathol 2011;42:1476](#))
- Strong cytoplasmic expression of COX2 at invasive fronts ([J Clin Pathol 2010;63:1048](#))

TX	Primary tumor cannot be assessed
T0	No evidence of primary tumor
Tis	Carcinoma in situ
T1a	Tumor invades lamina propria
T1b	Tumor invades muscular layer
T2	Tumor invades perimuscular connective tissue
T3	Tumor perforates serosa or directly invades the liver and/or one other adjacent organ
T4	Tumor invades main portal vein or hepatic artery or invades multiple extrahepatic organs
NX	Regional nodes cannot be assessed
N0	No regional nodal metastasis
N1	Metastasis to nodes along the cystic duct, common bile duct, hepatic artery and/or portal vein
N2	Metastasis to periaortic, pericaval, superior mesenteric artery, and/or celiac artery lymph nodes*
M0	No distant metastasis
M1	Distant metastasis

Stage 0	Tis	N0	M0
Stage I	T1	N0	M0
Stage II	T2	N0	M0
Stage IIIA	T3	N0	M0
Stage IIIB	T1-3	N1	M0
Stage IVA	T4	N0-1	M0
Stage IVB	Any T	Any N	M0
			M1

* Denotes changes from 6th edition classification.