BILE DUCT STRICTURES

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and Hepatology

BILE DUCT STRICTURES

Benign

- Post-operative
- Primary sclerosing cholangitis
- Post-transplant
- Chronic pancreatitis
- Malignant
 - Pancreatic/Ampullary cancer
 - Cholangiocarcinoma
 - Metastatic cancers
 - Lymphoma



BILIARY IMAGING

- Transcutaneous US or CT:
 - •Often the first studies obtained in a patient with jaundice
 - May demonstrate dilation with or w/o a stricture or mass

MRCP

- Offers more detail
- Features of malignancy: stricture length >10 mm, irregular margins, shouldering
- Cholangiocarcinoma diagnosis: 77-86% sensitive; 63-98% specific



BILIARY IMAGING

Endoscopic ultrasound •Used when other imaging is inconclusive EUS-FNA for extra-hepatic cholangiocarcinoma •77% sensitive, 53-89% specific FNA avoided at many centers if patient is a transplant candidate due to fear of seeding



WHAT IS THE ROLE OF ERCP?

Tissue Confirmation with Brush and Biopsy

- Disappointing sensitivity: 18-60%
- Allows Cholangioscopy
 - Directed biopsy
 - Visual appearance may suggest malignancy; although prior stenting may alter this
- Allows Intra-ductal US
 - Does not help with tissue acquisition
- Allows palliative stenting in non-surgical candidates



CHOLANGIOCARCINOMA BISMUTH CLASSIFICATION

TYPE I	TYPE II	TYPE IIIa	TYPE IIIb	TYPE IV
NVL	N N	Y L	N/	North Contraction

Bismuth 1 in good surgical candidate: Consider surgery Bismuth 2-5 in good surgical candidate: Consider transplant



ERCP FOR STENTING IN POOR SURGICAL CANDIDATES

- MRCP prior to ERCP helps determine best areas to stent: "volumetric liver assessment"
- Avoid injection of contrast into areas that will not be stented to avoid cholangitis
- SEMS provide better drainage and require less procedures than plastic stents
- Open mesh uncovered metal stents best for hilar strictures
 - Bilateral stents with either side by side or stent within a stent
 - Also allows future percutaneous approach if needed



INDETERMINATE BILIARY STRICTURES: TISSUE ACQUISITION METHODS

- Brush cytology alone: 30% sensitivity
- FISH: increases cytology sensitivity by 20%*
- Multimodal tissue (brush + biopsy): 60-70%
 - Standard biopsy forceps after ES
- Methionly-tRNA synthetase 1 immunofluorescence (MARS1 IF): Accuracy=94.5% in study of 240 pts**
- DNA methylation markers: Accuracy 91%***
- **EUS-FNA: 77% S, NPV: 29%**

Contra-indicated due to seeding in extra-pancreatic strictures



*Fritcher Gastro 2009 **Jang GIE 2021 ***Prachayakul GIE 2022

WHAT IS THE VALUE OF SPYGLASS CHOLANGIOSCOPY?

Pros:

- Single operator, Two dials, wire guided
- Channel for both water irrigation and device
- Reported accuracy of visualization: up to 89%*
- Allows directed biopsy

Cons:

- Small biopsy forceps, less sensitive than cytology plus FISH in one study**
- Increases risk of cholangitis***



INDETERMINATE BILIARY STRICTURES: IMAGING METHODS

- Cholangioscopic appearance
 - Blinded viewer accuracy: 45%*
 - Mendoza criteria: diagnostic accuracy 77%**
 - ^{2nd} Blinded viewer study: 74% sensitive, 46% specific***
- Intra-ductal US non-blinded accuracy
 - **•**71% in PSC; 90% in non-PSC

Problem: no tissue

*Sethi GIE 2012 **Kahaleh GIE 2021 ***Stassen GIE 2021



CONFOCAL LASER ENDOMICROSCOPY

- Cholangioflex probes: real time microscopic images
- More accurate than ERCP tissue sampling in one study*
- Multicenter study in PSC dominant strictures:**
 - Sensitivity for cancer: 85.7%; specificity: 73%
 - Did not perform well in CBD or left hepatic duct locations
- Problems with pCLE:
 - Underpowered studies
 - Only fair to poor inter-observer agreement
 - Tissue diagnosis still required to start therapy



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*Meining GIE 2011 **Han GIE 2021

INTRA-HEPATIC BILIARY STRICTURES

- Serum CA 19-9: Sensitivity 42%, Specificity 90%
- Segmental Right lobe strictures: consider post-cholecystectomy
- Benign vs. Malignant:
 - Mainstay is MRCP Imaging
 - Refer to Liver cancer center
 - Serial MRCPs in Indeterminate Strictures: q 3, 6 or 12 months
- If Unreachable by ERCP in non-operative candidate: EUS FNA or transcutaneous biopsy options



ENDOSCOPIC RX FOR BENIGN BILIARY STRICTURES: RANK ORDER OF SUCCESS

1. Primary sclerosing cholangitis

- -Stricture Rx delays transplantation; No effect mortality
- -Balloon dilation; usually no stent

2. Post-transplant -Anastomosis, donor duct, or SOD

- -Stent-free patency: 73-90%
- 3. Post-operative

4. Chronic pancreatitis

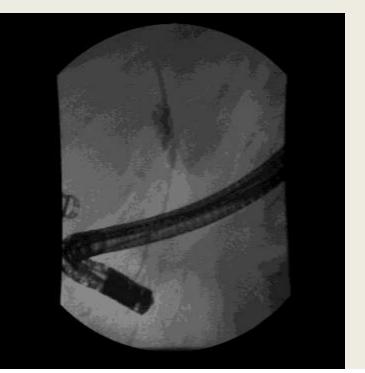


RARE CAUSES OF BENIGN BILIARY STRICTURES

- Autoimmune cholangiopathy / pancreatitis: type I (66-85% + IgG4) & type II (rare + IgG4)
- Choledochol cysts
- Previous stone disease
- Sphincter of Oddi stenosis not high-grade obstruction



LONG PSC STRICTURE





3 MONTH FOLLOW-UP





POST-TRANSPLANT STRICTURE



Dilation, then multiple 10F stents: 87% long term patency*
FCSEMs useful in recalcitrant cases



*Morelli GIE 2008

POST-CHOLECYSTECTOMY STRICTURE

- Cholecystectomy often complicated
- Symptoms within 2 yrs of surgery
- Short stricture in CHD typical
- Post-op stricture treatment:
 - Roux-hepaticojejunostomy
 - Morbidity: 10-20%; mortality: 2-3%
 - Stricture recurrence: 20-25%
- Endoscopic: dilation & stenting





BILE DUCT STRICTURE DUE TO CHRONIC PANCREATITIS

Treatment indications: duct >12mm or alk phos >3 X normal

Dilation, then MPS or FCSEMs

Better without pancreatic calcifications

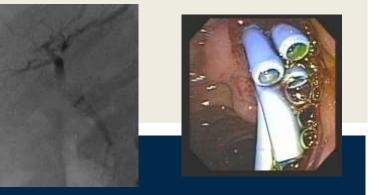
RCT of MPS vs FCSEMs:*

Similar resolution at 24 months: 76%

Less ERCPs (2.6 vs. 3.9) in FCSEMs

Similar Adverse Event rate







*Ramchandani Gastro 2021

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BENIGN STRICTURE CONCLUSIONS

- PSC + post-OLT: Endo Rx successful
- Recalcitrant strictures (post-op + CP):
 - Added value to multiple plastic stents
 - Requires 4-5 ERCPs
 - Morbidity: stent occlusion / infection
 - Biliary bypass still reasonable alternative for CP
- **FCSEMs** excellent alternative for recalcitrant strictures



WHEN TO CHOOSE A FCSEM?

Benign biliary strictures

- Simpler to place and requires fewer ERCPs
- Faster stricture resolution
- More stent migration (25% vs. 16% with plastics)
 - Less migration risk with flanged FCSEM or with double pigtail within
- Avoid in small ducts (<6 mm)</p>
- Probably no need to avoid cystic take off
- Treat post ES bleed / perforation, treat intra-ductal bleed
- When cancer diagnosis is unclear or stricture is close to hilum
- More costly and more migration than bare metal SEMS for inoperable panc cancer with low strictures



Cote JAMA 2016, Park AJG 2011

MALIGNANT BILIARY OBSTRUCTION

- Presentation: Painless jaundice & wt. loss
- Diagnoses
 - Pancreatic cancer
 - Bile duct cancer
 - Ampullary carcinoma
 - Gallbladder cancer
 - Metastatic lymph nodes



SUSPECTED MALIGNANT OBSTRUCTION EVALUATION

- Pancreatic protocol CT for diagnosis / operability decision
- EUS / FNA: tissue confirmation if neoadjuvant Rx planned

*Van der Gaag NEJM 2010

- No need for pre-op biliary drainage*
 - Studied for t. bili <14
- Neoadjuvant Rx planned pre-op:
 - Short SEMs due to surgery delay
- Suspected hilar cancer: MRCP before ERCP



MANAGEMENT OF MALIGNANT HILAR OBSTRUCTION

- 5-year survival <5%</p>
- Determine resectability or OLT candidancy: 73% are unresectable
- Most patients require biliary drainage for symptoms & to preserve liver function
- SEMSs is superior to Plastic stents unless operability not yet determined
- Bilateral SEMS (either stent in stent or side by side) better than unilateral
- Avoid Percutaneous drainage in potentially operative patients
- Inoperative patients:
 - PTBD likely higher success rate and lower adverse events
 - However, endoscopic drainage strongly preferred by patients



Qumeya ASGE Guideline; GIE 2021

CONCLUSIONS

 Determine benign vs. malignant & resectability
 Multiple imaging modalities: MRCP, CT, EUS, cholangioscopy, IDUS, probe confocal microscopy
 Role of ERCP

- Obtain tissue diagnosis
- Palliative stenting in malignancy
- Endoscopic treatment of benign strictures

