In vivo appearances of gallbladder carcinoma under magnifying endoscopy and probe-based confocal laser endomicroscopy after endosonographic gallbladder drainage

Fig. 1 a,b Computed tomography showing the lumen-apposing stent drainage of the gallbladder to the first part of the duodenum.

Fig. 2 Gallbladder adenocarcinoma in an 87-year-old man, seen at follow-up endoscopy 3 months after endosonographic gallbladder drainage with a lumen-apposing stent: **a** white light endoscopic view; **b** magnified narrow band imaging (NBI) appearance.

An 87-year-old man with multiple medical co-morbidities was admitted for right upper quadrant pain and fever. Computed tomography showed a distended gallbladder with gallstones and pericholecystic fluid compatible with a diagnosis of acute cholecystitis. There was no gallbladder mass. Since the patient had sepsis and was unfit for surgery, endoscopic ultrasound (EUS) drainage of the gallbladder was performed with a lumen-apposing stent (10×15 mm, AXIOS; Xlumena, Mountain View, California, USA) as an alternative to percutaneous cholecystostomy (Fig. 1 a,b). The patient had an uneventful recovery.

Follow-up cholecystoscopy performed 3 months later showed clearance of all stones. However, a 2-cm polypoid lesion was noted at the fundus of the gallbladder. Magnifying narrow band imaging (NBI) showed enlarged and irregular mucosal glands with dilated and corkscrew-appearance microvasculature that was suspicious for malignancy (Fig. 2). Probe-based confocal laser endomicroscopy (CLE) (GastroFlex; Mauna Kea Technologies, France) showed darkened and irregular columnar cells with loss of villous architecture (Fig. 3 a,b). Miniprobe EUS examination (UM-DP12–25R; Olympus, Tokyo, Japan) showed suspicion of tumor

involvement of the gallbladder muscularis propria (Fig. 4). Final histological findings confirmed the presence of a gallbladder adenocarcinoma (Fig. 5). The patient was then treated conservatively as he was too frail to undergo any major surgery.

To our knowledge, this is the first description of a gallbladder adenocarcinoma discovered endoscopically. This was made possible through the use of endosonographic drainage that allowed endoscopic assessment of the gallbladder [1,2]. Magnifying NBI endoscopy has revolutionized the diagnosis of early gastrointestinal neoplasms and is pivotal to performance of endoscopic mucosal resection or submucosal dissection [3]. CLE allows in vivo assessment of cellular architecture and has been shown to be associated with high sensitivity and specificity for diagnosis of Barrett's metaplasia and biliary malignancy [4]. EUS-guided gallbladder drainage in this patient opened the way for application of the above instruments in aiding diagnosis that was not suspected on CT.

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Fig. 3 **a** Confocal laser endomicroscopy (CLE) image of the normal-looking gallbladder mucosa, showing villous architecture with columnar epithelium. **b** CLE image of the gallbladder adenocarcinoma, displaying marked darkened, variably sized glands.

Fig. 4 Miniprobe examination with endoscopic ultrasound (EUS) showed tumor involvement of the muscularis propria of the gallbladder.

Fig. 5 A complex cribriform malignant gland is present in the stroma supporting the diagnosis of adenocarcinoma (hematoxylin and eosin [H&E], original magnification $\times 400$).

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