Choledochal malformations (CMs) may be characterised as an abnormal dilatation of the biliary tract in the absence of acute obstruction. Most appear to be of congenital origin, probably related to distal bile duct stenosis, and almost 15% can now be detected antenatally.

Excision and biliary reconstruction using a Roux loop as an open operation is still the standard to compare against, although laparoscopic reconstruction is increasingly reported. This article discusses recent advances in the understanding of choledochal malformation aetiology and classification, together with the role of newer modalities of surgical treatment such as laparoscopic excision and biliary reconstruction. Although these are definitely feasible, care should be taken before dispensing with standard open techniques that have minimal complications and proven long-term benefit.


**Classification of choledochal malformations**

One early point of controversy is what to call this entity (International Classification of Diseases, 9th revision, clinical modification (ICD-9-CM) diagnosis code 751.69). The older term, choledochal cyst, describes a predominantly spherical or globular biliary dilatation, and is actually only one type of CM. Modern authors have generally adopted a less prescriptive phrase, for example congenital CM (or dilatation).

The Japanese authors Todani et al. developed their own classification during the 1970s and 1980s, defining three variants of type 1, two variants of type 4 (A & B) and adding isolated intrahepatic dilatation as type 5. The classification is complex and many authors continue to misquote the original, especially confusing cystic and fusiform.

The King's College Hospital classification (Fig. 1) simplifies the Todani classification into types 1C and 1F (depending on the predominant appearance as cystic or fusiform) and limits type 4 to the combination of intra- and extrahepatic dilatation. This classification has been the basis of our attempts to define pathophysiological characteristics to each type. Other workers have emulated this simpler approach.

**Epidemiology**

CMs can present at any point in the life cycle from an antenatal scan to the postmortem table, which makes the true incidence hard to define.

If biliary atresia is used as a guide for a condition where the incidence is known (1 in 17,000) and the ratio of the two conditions presenting in infancy is taken from a specialist hepatobiliary unit in the UK, then an approximate figure of about 1 in 100,000 live births may be suggested. However, the incidence is much higher in Asian populations, and a female predominance of around 4:1 is consistently found in most series.

There is no obvious hereditary element in most CMs, though a recent review identified 10 pairs of familial occurrence (siblings and offspring) in the literature. Conversely, six pairs of monozygotic twins have been described in which a distinct CM appeared in only one pair. Aside from Caroli’s syndrome (see below), the condition is usually isolated. Fusiform biliary dilatation occasionally follows the repair of duodenal atresia, although proving this as an acquired aetiology is often difficult.
Type 4 CM is found in about 20% of large surgical series of CM and it is arguable whether this is simply the natural history of untreated types 1C and 1F. Some authors have distinguished cystic intrahepatic dilatation from a more fusiform intrahepatic dilatation, implying that the latter diminishes considerably following effective surgery. Our own follow-up study certainly demonstrated considerable resolution in intrahepatic duct dilatation in the first year post surgery.[8]

Whether the choledochocele (type 3 CM) should be included at all in any classification was the subject of a recent North American series.[9] The authors argued that there were so many significant differences when compared with types 1, 2 and 4, particularly age at presentation (invariably as an adult) and gender (predominantly male) that it barely merited the term ‘congenital’. Certainly, the distinction between this and a long-standing dilated common channel, coupled with a degree of ampullary stenosis, is difficult.

Isolated type 5 intrahepatic dilatation does occur and is distinct from what we recognise as Caroli’s syndrome. This latter entity struggles to fit into any classification of CM because it is so distinctly different. There is an obvious genetic abnormality, leading to a fundamental probable basement membrane defect in small bile ducts. The syndrome is associated with similar fundamental pathology affecting the kidney, manifesting as renal cystic disease or renal...

**Fig. 2.** Choledochal pressure in relation to type of CM (n = 47) (A); choledochal pressure in relation to ELMS (n = 47) (B); and bile amylase concentration in relation to ELMS (n = 73) (C). Overall P = 0.002 (Kruskal-Wallis ANOVA).

Figures reproduced with permission from Turoski et al.[13] (CM = choledochal malformation; ELMS = epithelial lining mural score.)

**Fig. 3.** A and B. Type 4 CM: Two-year-old boy presenting with recurrent jaundice. There is fusiform extra- and intrahepatic dilatation on magnetic resonance cholangiopancreatogram (A); at surgery the cholangiogram also shows a dilated common channel, partially obstructed by debris (white arrows) (B) with obvious reflux into the pancreatic duct (black arrows). The bile amylase level in this case was 20 000 IU/L. (CM = choledochal malformation.)
fibrosis. Furthermore, there is often associated parenchymal liver fibrosis in the absence of any particular bile duct obstruction.

There is no doubt that part of the CM complex (at least in types 1C, 1F and 4) is a degree of pancreaticobiliary malunion or 'long' common channel, with the junction lying outside of the wall of the duodenum (and therefore not under sphincteric control). Whether it is simply a reflection of disordered bile duct development during fetal life, and hence incidental, or is actually aetiological, as the Babbitt hypothesis implies (see below), remains to be argued.

Aetiology

Two hypotheses have been put forward to explain the appearance and configuration of the common types of CM. The most fashionable has been the so-called Babbitt hypothesis, which suggests that there is reflux of activated pancreatic enzymes via the common channel into the biliary tract. Donald Babbitt was an American radiologist who first observed this during cholangiography. The hypothesis suggests that this results in enzymatic damage to the local biliary epithelium and weakening of the bile duct wall, leading to dilatation and loss of bile duct structural integrity. An older hypothesis posits that there is a narrow stenotic portion of common bile duct, proximal to the common channel, that generates high intraductal pressures upstream. Prolonged exposure to elevated ductal pressure dilates the proximal lumen, similar to the dilatation seen in the bowel in congenital bowel atresia. Ligation of the duct in fetal lambs by the South African Lewis Spitz certainly produced cystic change in the delivered animals. We sought to disentangle these competing hypotheses in a series of reports based on clinical observations at the time of the laparotomy. Initially, we identified an inverse relationship between bile amylase (a surrogate of pancreatic juice reflux) and measured choledochal pressure. We also determined that there was a stepwise increase in pressures observed from type 1F through type 1C to type 4 (Fig. 2A), accompanied by a reduction in bile amylases. We then related these variables to a semiquantitative histological score, reflecting the damage done to the biliary epithelium, and showed that the most abnormal histology was in bile ducts with the highest pressures (and therefore the lowest amylase levels) (Fig. 2B and 2C). These observations in clinical practice seemed to negate the central aetiological mechanism underlying the Babbitt hypothesis.

Clinical features

The classic presentation with abdominal pain, jaundice and a mass in the right upper quadrant is in fact rarely seen in large paediatric series. Infants tend to present with obstructive jaundice, and in later childhood and adult life, patients present with recurrent abdominal pain, with or without features of acute pancreatitis (Fig. 3). Recurrent pancreatitis should be seriously considered as a cause for recurrent abdominal pain in childhood and actively excluded by the measurement of serum amylase or lipase. A value of >3 times the upper normal limit, in association with typical abdominal pain and/or features of pancreatitis on cross-sectional imaging (the Atlanta criteria), confirms the diagnosis. Once a diagnosis of pancreatitis has been made, imaging (if not already completed) should at minimum consist of ultrasonography with a low threshold and continue on to a magnetic resonance cholangiopancreatogram and, if necessary, an endoscopic retrograde cholangiopancreatogram. Historically, untreated choledochal cysts caused cirrhosis and death in many instances (mortality rates approached 100% in the 1940s). Prompt surgical resection after diagnosis has rendered this scenario rare in recent times.

Surgical intervention

Early attempts at surgical treatment of choledochal cysts usually involved internal drainage to adjacent intestinal loops or to the duodenum. This approach, leaving the cyst intact, persisted until at least the 1970s. The Chicago surgeon G McWhorter is credited with the first successful resection and hepaticoduodenostomy (HD) in 1924.

Surgery is the only effective treatment for CMs, although the timing can be negotiated. With antenatal detection possible, some obvious type 1C CMs are picked up in the presymptom phase (Fig. 4). Some surgeons advocate immediate surgery during the neonatal period to forestall the development of silent hepatic fibrosis. However, deferment to about 3 - 6 months of age seems more pragmatic as liver biochemistry demonstrably returns to normal after the early neonatal period.
Farello et al. first described the laparoscopic resection of a choledochal cyst in a 6-year-old child in 1995. Since then, increasing numbers of specialist centres worldwide have adopted this challenging approach using minimally invasive techniques. In comparison with open surgery, operative times are longer and, certainly in early series, there was a higher incidence of complications such as anastomotic leaks without any real decrease in hospital stay (at least in children). More recently, huge single-institution series of more than 200 cases have been published from Beijing and Hanoi with much lower complication rates. This alludes to the increased centralisation of expertise in those countries and the effect of high-volume surgery.

Minimally invasive resection is an advanced technique requiring considerable dexterity. Other techniques have been used to try to improve the quality of the biliary anastomosis – the Achilles heel of the entire operation. Thus, robot-assisted surgery has been reported to facilitate manual techniques, although the results from small series show a long learning curve and considerable complications.[24] Robots are a prohibitively expensive technology for many healthcare systems, and surgeons in large Asian series do not seem to require them. Extracorporeal jejunojunostomy through a widened umbilical incision may also reduce operating times. Even single-incision surgery is possible, though apart from marginal improvement in cosmesis, there are no other advantages.

What has been the most controversial recent technical development is a swing back to HD as the reconstruction device. Although the long (40-50 cm) Roux loop and hepatojejunostomy have been the mainstay of biliary reconstructive surgery for approximately the past 50 years, it is difficult to do laparoscopically. An easier alternative is a simple HD. The transected hepatic duct and the mobilised first and second parts of the duodenum can be in close proximity. However, there is a fundamental problem with this technique, which manifested some years ago by surgeons who once preferred it as an open technique.

Todani was a long-time advocate of HD, but also showed that about 10-20% of patients have symptomatic bile gastritis due to reflux across the pylorus and a 10% incidence of cholangitis due to reflux of non-sterile duodenal content into the biliary tree, which is often dilated. The final straw, from his perspective, was the reflux of non-sterile duodenal content into the biliary tree, which is often dilated. The final straw, from his perspective, was the development of a bile duct carcinoma at the portal hilum 19 years after primary cyst excision and HD performed on a 13-month-old patient. He then switched to hepatojejunostomy for the rest of his career. Other authors in comparative trials have also come to the same conclusion.[27] Therefore, what is wrong with hepatojejunostomy-en-Roux? In our opinion, not much. It is non-physiological and takes a significant amount of absorptive jejunal surface area out of the intestinal sequence. Furthermore, there is constant bile drainage and no intermixing with pancreatic juice until the jejunal union. In a recent study of growth indices in our series of over 100 operated adults with CM, we found a significant incidence of failure to thrive and lower preoperative weights when compared with the age-corrected normal population. However, after surgery, all patients showed satisfactory catch-up weight gain at 1 year and sustained weight gain thereafter (unpublished observations).

Malignant transformation and cancer risk

The long-term risk of malignant transformation in CM has been well described and is evident in many published series of adults with CM. This risk is probably due to ongoing chronic inflammation in the epithelium over many years, exacerbated by episodic cholangitis and stone formation. The youngest case appears to be an 11-year-old boy, reported by Tanaka et al.[26] in 2006, who developed adenocarcinoma in a type 4 CM. Tanaka et al. also identified ten more cases from the literature in patients aged <20 years, the majority also arising in long-standing type 4 CMs. What is not known with any certainty is the risk of carcinoma in the residual biliary epithelium in the liver or in the head of the pancreas following CM resection in the modern world. We simply have not been doing this kind of surgery long enough to know what happens years later during adulthood. At the very least, however, it should be axiomatic that treated children and adolescents are followed up – ideally by serial ultrasonography. Persistence of intrahepatic dilatation and the development of stones and debris should encourage more invasive investigation.

Discussion of CM and cancer inevitably overlaps with the problems of what to do in patients with an isolated common channel (usually defined as >6 mm long and also referred to in the literature as pancreatocholedochal maljunction). Almost paradoxically, there seems to be a greater malignancy risk with common channels unassociated with CM. In one Japanese survey of hepatobiliary surgical practice, biliary cancer occurred in 22% of adult patients with biliary dilatation and 42% of those without biliary dilatation. Gallbladder cancer is the usual malignancy in the latter group.[27]

Conclusion

Treatment outcome in CM has improved dramatically over the last 60 years, led by earlier diagnosis, prompt surgical excision and effective biliary drainage. There are still unresolved issues surrounding the nature and style of surgical reconstruction, but long-term success with normal liver function is unequivocal.

References