A study of the anatomic variations of the pancreatico-biliary system in Palestine: a national study

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INTRODUCTION

In recent years in Palestine, hepatobiliary surgeries, such as cholecystectomies, liver resections, and partial liver transplantsations are increasing in frequency and complexity. This fact calls for a better understanding of biliary anatomy and potential biliary variations. In this article, we present an unprecedented national study that deserves attention in both clinical and surgical settings.

There is no absolute anatomy of the hepto-biliary tract, as only about 60% of humans present with the proposed typical anatomy.1-3 The anatomy of the hepatobiliary system is complex and there are several anatomic variations, some of them are common and others being less frequent.4 Although in the majority of cases the anatomic variations can be asymptomatic these variants, if unrecognized, can increase the probability of unintentional iatrogenic injury during surgical procedures, as cholecystectomy due to inadvertent ligation or transection.5 These injuries consecutively can lead to significant morbidity and mortality.6 Therefore, familiarity and knowledge of these anatomic variations are highly important for the hepatobiliary surgeons.7 A common pattern of verities exists and it is the surgeon's responsibility to recognize these variations when present.1,8 In addition, knowing the precise anatomy of the biliary duct is very important in some procedures like...
the selection of donors in liver transplantation. For example, one of the contraindications to a safe right hepatectomy is the presence of "triple confluence" and the existence of "right posterior duct" draining into "the left hepatic duct is a contraindication to both right and left hepatectomy."

To identify the anatomic variations, it is absolutely necessary to appreciate the basic knowledge of the embryological development and normal anatomy of the biliary system. The biliary development starts in the fourth week of embryonic life in which a hepatic diverticulum gives rise to two buds: the cranial bud that later grows into the liver and the extrahepatic biliary tree and the caudal bud which develops into superior and inferior ducts which become the gallbladder and cystic duct, and the right and left ventral pancreas, respectively. The intrahepatic development is not fully known with different theories were suggested by different studies.

A variety of imaging modalities have been emerged that may be used to visualize the anatomy of the biliary system such as intravenous cholangiography, Endoscopic retrograde cholangiopancreatography (ERCP), Intraoperative cholangiography and Magnetic resonance cholangiopancreatography (MRCP). MRCP is a recent and constantly evolving imaging technique that studies the bile and pancreatic duct, with an accuracy reaches 90-95%. It has been preferred over ERCP and other imaging techniques, and have been considered the ideal modality to study the pancreaticobiliary tract anatomy, as it is noninvasive, ionizing radiation free, does not require anesthesia. For example the common and potentially severe complications of ERCP and their high risk of mortality and morbidity as pancreatitis, hemorrhage, bowel perforation, and infection are not encountered using MRCP. Due to these reasons in this study we relied on MRCP images to demonstrate the anatomic variations and anomalies of the biliary and pancreatic system.

The aim of this study is to detect and document the prevalence of anatomical variations of the biliary system in the Palestinian population, to assess their frequencies and to compare our findings to anatomical variations described elsewhere in the literature as no data are available on this important subject locally.

METHODS

During a period of three years, from March 2016 to January 2019, a total of 401 consecutive magnetic resonance cholangiopancreatographies was performed at three different Palestinian institutions- NNUH, Patients Friend Society Center, and Ramallah hospital. These centers are the main ones in Palestine and they receive patients from all over the region. Sampling method was convenient, including all patients referred for MRCP suspected of having inflammatory, lithiasic, or neoplastic disease of the hepatobiliary and pancreatic system.

Two professional radiologists, each with over ten years of experience, prospectively assessed the MRCP’s and were blinded to the patients’ identities. Results obtained from both radiologists were compared to assess the degree of inter observer agreement. Any discrepancy between the reviewers was resolved by a consensus. Structures mainly examined and evaluated were the gallbladder, cystic duct, intrahepatic bile ducts, pancreas, pancreatic duct, major papilla and common bile duct. A detailed analysis was made in regards to their location, shape, length and course. MRCP in these patients was carried out for the purpose of detecting stones causing biliary obstruction, identifying hepatobiliary and pancreatic tumors, staging tumors, assessing iatrogenic biliary injuries, and evaluating congenital biliary anomalies.

Regardless of the underlying cause, the MRCP’s were chosen to be included in the study only if opacification of the biliary and pancreatic ducts was adequate, and if the anatomy was clearly visualized. In some images, part of the hepatobiliary system was obscured, yet these MRCP’s were still included in the study as other biliary features were clearly visualized and variations were detected. The feature that was obscured was excluded and the study was made accordingly. In view of this, fifty five images were totally excluded from the study, because of unclear and obscured biliary anatomy or because the MR examination did not image the entire biliary tree as required.

The anatomy of intrahepatic bile ducts was labeled normal when the following conditions applied as shown in Figure 1a. First the right posterior duct runs horizontally posterior to the vertically coursed right anterior hepatic duct. Second the posterior duct joins the
anterior in the medial aspect. Third the union of the two ducts forms a common duct, the right hepatic duct. Fourth the right duct integrates the left hepatic duct to form the common hepatic duct (CHD). Any deviation from the above description was considered as anatomic variation. Figure 1 shows the normal intrahepatic bile ducts anatomy (a) and its common anatomical variations.

Normal gallbladder was defined as an elliptical organ lies in the fossa between the right and left lobes of the liver. It was studied for the presence of different variations in its shape and location. Variations include failure in its formation (agenesis), abnormal positioning (ectopic), duplication, bilobing and the presence of a fold or septum between the body and fundus (Phrygian cap), multiseptation and diverticulum.

Cystic duct was defined as the portion that connects the gallbladder to the CHD. It was considered normal if it inserts at the middle third of the CHD from the lateral aspect and ranges 2-4 cm in length. It was evaluated for different anatomical variations regarding its insertion, length, and course. Studied anatomical variations include absence of cystic ducts, duplicated cystic duct, cystic ducts with medial, anterior and posterior aspect of insertion, higher and lower level of insertion than the normal consideration, shorter and longer cystic ducts than the mentioned length range and a cystic duct with a parallel course to the common hepatic duct extending for 2 cm or more.

A common bile duct (CBD) was defined as the duct which is formed at the junction of the CHD and the cystic duct and joins the pancreatic duct at the ampulla of vater. It was studied for the presence of any dilation or enlargement represented as choledochal cyst. The absence of a common channel of pancreatic duct and CBD was considered as a variant.

A normal anatomy of pancreatic duct was considered when the main pancreatic duct is divided into two ducts, the duct of Wirsung, which is posterior and drains the ventral pancreas in the major papilla and the accessory duct of Santorini, which is anterior and drains the dorsal pancreas in minor papilla. Failure of fusion of the two duct (pancreatic divisum) Figure 2 (A-C), switching of the site of drainage (ansa pancreatica) Figure 2 (D), bifiding, duplication and cystic dilation of the pancreatic duct were all considered as variations. We also studied different courses of the pancreatic duct Figure 3.

A normal pancreas was defined as a J shaped organ lies transversely in the posterior abdomen. It was studied for the presence of different anatomical variations including agenesis, ectopic pancreas, annular pancreas, accessory pancreatic lobe, hypoplasia of the dorsal pancreas and pancreatic cysts.

**MRCP technique**

All MRCPs were performed by using a 1.5T scanner (Philips Ingenia). MR examinations were made including axial thick slice turbo spin echo (TSE) with parameters of TR 400, TE 80, voxel size of slice thickness 6 mm x 1.4 mm x 1.6 mm. Flip angle is 90 degree.

Dimensions of frequency and phase encodings and number of slices are patient dependent, which should cover the abdomen AP and RL and FH. NSA is 1 with no fat suppression. The other sequence is called MRCP 3D HR designed to show the biliary system. TR is 1024 TE600 with voxel size 1x1x0.9 slice thickness. NSA with fat suppression. Both sequences done with respiratory
trigger for respiratory compensation. Date of images is processed to reconstruct maximum intensity projection (MIP) images and multiplanar reformatted images (MPR).

**Statistical analysis**

SPSS statistical software was used for data entry and analysis. Frequency and percentages for categorical variables was computed and presented in tables and figures. Chi square test was used to compare the anatomic variation between male and female groups. A p-value of less than 0.05 was considered significant.

The study was ethically approved and permissions were taken from the different hospitals.

**RESULTS**

A total of 346 MRCP images were assessed for hepatobiliary system anatomical variations, 189 were females (54.1%) and 157 were males (45.3%), with a male to female ratio of 0.83:1. Their ages ranged from 5 to 89 years, with a mean of 51.8 years. Among the studied images, 43.35% had the proposed normal anatomy.

**Intrahepatic ducts variations**

Intrahepatic ducts were observed in 342 images out of the 364 images included in the study. Of them 266 (77.8%) images had normal anatomy. Anatomical variations of the intrahepatic duct were documented as right posterior duct joining the right anterior duct by its lateral side in 43 images (12.6%), Right posterior duct draining into the left hepatic duct (Figure 4) in 2 images (0.6%), triple confluence in 29 images (8.5%), right posterior duct draining directly into the cystic duct in 1 image (0.3%) and right posterior duct draining directly into the common hepatic duct in 1 image (Figure 5).

**Gallbladder variations**

Regarding the gallbladder, it was visualized in 242 images out of the whole studied images (346 images). It was normal in 216 images (89.3%), bilobed in 1 image (0.4%), ectopic in 9 images (3.7%), with Phrygian cap (Figure 6A) in 9 images (3.7%), diverticular in 3 images (1.2%), and multi-septate (Figure 6B) in 10 images (4.1%) (Figure 7).

**Figure 5:** Distribution of intrahepatic ducts normal anatomy and variations.

**Figure 6:** MRCP images show some variations in the anatomy of gallbladder, (A) Phrygian cap and (B) multi-septate gallbladder.

**Figure 7:** Distribution of gallbladder normal anatomy and variations.
We found a significant association \((p=0.04)\) between the existence of varied anatomy and pathological gallbladders in males. The number of males who have gallbladder anatomical variations and pathology at the same time was 15, while in females the number was 11. Pathology includes stones, sludge, wall thickening, acute/chronic inflammation, distention, tumor or cyst.

**Cystic duct variations**

The cystic duct was evaluated for its level of insertion, site of insertion, length and parallelism. The MRCP was adequate for evaluation 302 images. Of them, in general 228 (75.5%) images showed normal anatomy and the remaining 74 (24.5%) images had anatomical variations (Figure 8).

![Figure 8: Distribution of cystic duct normal anatomy and variations.](image)

For the cystic duct insertion it was found to be normal (middle) in 257 images (85.1%), low in 29 images (9.6%) and high in 16 images (5.3%). Regarding its site of insertion, it was lateral (normal) in 270 images (89.4%), medial (Figure 9) in 20 images (6.6%), anterior in 5 images (1.7%) and posterior in 7 images (2.3%). The cystic duct was found to be parallel in 8 (2.3%), and not parallel (normal) in 294 (85%). As for the length, it was normal in 274 images (90.7%), short in 7 (2.3%), and long in 21 (7%).

**CBD and major duodenal papilla variations**

According to common bile duct, it was normal in 336 out of 340 (98.8%), and a choledochal cyst was found in 4 images (1.2%). The MRCP successfully depicted the major duodenal papilla in 345 images. Of these 345 images, it was normal in 335 images (97.1%) while 10 images (2.9%) had no common channel.

**Pancreatic duct variations**

Variations of the pancreatic ducts were evident in 13 (3.9%) of the 341 images in which the pancreatic duct was visualized. In particular the MRCP showed divisum classical in 4 (1.2%), divisum absent in 5 (1.5%), divisum incomplete in 2 (0.6%) and ansa pancreatica in 2 images (0.6%) (Figure 10).

![Figure 10: Distribution of pancreatic duct normal anatomy and variations.](image)

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The course of pancreatic duct was descending in 258 (75.4%), sigmoid in 65 (18.7%), vertical in 15 (4.4%) and loop in 4 (1.2%) (Figure 11). We were able to visualize the pancreas in 343 images and it was normal in all of them.

Table 1: Hepatobiliary system anatomical variations in relation to gender.

<table>
<thead>
<tr>
<th>Hepatobiliary system</th>
<th>Males frequency (%)</th>
<th>Females frequency (%)</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Intra-hepatic bile ducts</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>121 (35.3)</td>
<td>145 (42.4)</td>
<td>0.749</td>
</tr>
<tr>
<td>Abnormal</td>
<td>33 (9.6)</td>
<td>43 (12.5)</td>
<td></td>
</tr>
<tr>
<td><strong>Gallbladder</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>99 (40.9)</td>
<td>117 (48.1)</td>
<td>0.252</td>
</tr>
<tr>
<td>Abnormal</td>
<td>15 (6.19)</td>
<td>11 (5.5)</td>
<td></td>
</tr>
<tr>
<td><strong>Cystic duct</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>101 (33.4)</td>
<td>127 (42.0)</td>
<td>0.514</td>
</tr>
<tr>
<td>Abnormal</td>
<td>36 (11.9)</td>
<td>38 (12.5)</td>
<td></td>
</tr>
<tr>
<td><strong>Cystic duct site of insertion</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>120 (39.7)</td>
<td>150 (49.6)</td>
<td>0.647</td>
</tr>
<tr>
<td>Abnormal</td>
<td>17 (5.6)</td>
<td>15 (4.96)</td>
<td></td>
</tr>
<tr>
<td><strong>Cystic duct level insertion</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>116 (38.4)</td>
<td>141 (46.6)</td>
<td>0.982</td>
</tr>
<tr>
<td>Abnormal</td>
<td>21 (6.95)</td>
<td>24 (7.94)</td>
<td></td>
</tr>
<tr>
<td><strong>Cystic duct length</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>123 (40.7)</td>
<td>151 (50)</td>
<td>0.875</td>
</tr>
<tr>
<td>Abnormal</td>
<td>14 (4.63)</td>
<td>14 (4.63)</td>
<td></td>
</tr>
<tr>
<td><strong>Cystic duct parallelism</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>132 (43.7)</td>
<td>162 (53.64)</td>
<td>0.615</td>
</tr>
<tr>
<td>Abnormal</td>
<td>5 (1.65)</td>
<td>3 (0.99)</td>
<td></td>
</tr>
<tr>
<td><strong>Pancreatic duct</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>148 (43.4)</td>
<td>180 (52.7)</td>
<td>0.942</td>
</tr>
<tr>
<td>Abnormal</td>
<td>6 (1.76)</td>
<td>7 (2.02)</td>
<td></td>
</tr>
<tr>
<td><strong>Common bile duct</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Normal</td>
<td>151 (44.4)</td>
<td>185 (54.4)</td>
<td>0.425</td>
</tr>
<tr>
<td>Abnormal</td>
<td>1 (0.29)</td>
<td>3 (0.88)</td>
<td></td>
</tr>
</tbody>
</table>

Table 2: The distribution of the pancreatic duct course types.

<table>
<thead>
<tr>
<th></th>
<th>Number of variations</th>
<th>Female</th>
<th>Male</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>N (%)</td>
<td>N (%)</td>
<td>N (%)</td>
</tr>
<tr>
<td><strong>Descending type</strong></td>
<td>259 (75.7)</td>
<td>142 (75.5)</td>
<td>117 (75.9)</td>
</tr>
<tr>
<td><strong>Sigmoid type</strong></td>
<td>64 (18.7)</td>
<td>39 (20.7)</td>
<td>25 (16.2)</td>
</tr>
<tr>
<td><strong>Vertical type</strong></td>
<td>15 (4.38)</td>
<td>7 (3.7)</td>
<td>8 (5.19)</td>
</tr>
<tr>
<td><strong>Loop type</strong></td>
<td>4 (1.16)</td>
<td>0 (0)</td>
<td>4 (2.59)</td>
</tr>
<tr>
<td><strong>total</strong></td>
<td>342</td>
<td>188</td>
<td>154</td>
</tr>
</tbody>
</table>

The anatomical variations of hepatobiliary system was studied in relation to gender (Table 1). No significant difference was found between gender and the variables.

We found a significant difference in loop pancreatic duct between males and females (p=0.026). 4 males were documented to have a loop course with no documented cases in females. No significant difference was found with other pancreatic duct courses (Table 2).

DISCUSSION

A thorough knowledge and complete understanding of the normal hepatobiliary tract is of fundamental importance for hepatobiliary surgeons and should not be overlooked. A surgeon can encounter different anomalies intraoperatively, in the form of absence, aplasia, and abnormal drainage of ducts, which he/she should be aware of. Failure to recognize these anomalies can have deleterious consequences for the patient. For instance,
complete transection of the CBD occurs when it is mistaken for the cystic duct, and it is one of the dreaded complications of laparoscopic and open cholecystectomy.\textsuperscript{18} Accurate preoperative delineation and assessment of the biliary anatomy is thus necessary to ensure safe and successful percutaneous, endoscopic, and surgical procedures. Such surgical procedures can range from the more commonly and routinely performed cholecystectomies, to the more complex surgeries as tumor resection, partial hepatectomy, and liver transplantation.

MRCP is an always evolving imaging technique that is used to optimally image the biliary and pancreatic ductal anatomy.\textsuperscript{5} It is non-invasive, ionizing radiation free, and does not require any anesthesia.\textsuperscript{4} Studies have revealed that preoperative MRCP provides substantial information regarding cystic duct anatomy and has a key safeguarding effect on laparoscopic cholecystectomy.\textsuperscript{3,4} In addition to the fact that it helps avoid iatrogenic injuries, preoperative identification of bile duct anatomy may also help in medicolegal purposes.\textsuperscript{18}

In the present study, MRCP demonstrated biliary and pancreatic duct anatomical variations in 43.35\% of patients. Compared to other regions of the world, this frequency is similar, as according to the literature, the incidence of the normal pattern of the biliary system has been reported to be 57\%–72\%.\textsuperscript{19} This highlights the importance of our study and stresses its implications for hepatobiliary surgery.

A wide variability is noted in the course of the cystic duct and its junction with the extrahepatic bile duct. Classical anatomy of the cystic duct is joining the common hepatic duct at its middle third from the lateral aspect; this is seen in 58\%–75\% of cases.\textsuperscript{20} In our study, we have seen this anatomy in 85.1\% of our cases. In the study of Sarawagi et al, they had this classical anatomy in 51.5\% of their cases.\textsuperscript{18}

Low insertion of the cystic duct was observed in 9 to 11\% of cases in previous studies.\textsuperscript{1,4,21} The study of Sarawagi et al reported 9\% of its cases as having low insertion of the cystic duct.\textsuperscript{18} Similarly, our study revealed LICD in 9.6\% of our cases. A study conducted in Italy by De Filippo et al reported low insertion of the cystic duct in 4.5\% of its cases.\textsuperscript{5}

Medial insertion of the cystic duct (MICD) was reported in 10–18\% of cases in previous studies.\textsuperscript{22} This variant is important during surgical procedures, as dissection of the medial cystic duct up to its end is considered unsafe and it is recommended to leave a long remnant of the cystic duct.\textsuperscript{10} In our study, we have seen a MICD in 6.6\% of our cases.

Parallel course of the cystic duct was reported in 7.5\% cases of a study conducted in Pakistan.\textsuperscript{18} It was seen to be found in 4\% of cases in an Indian study.\textsuperscript{23} Our study has seen it in 2.3\% of its cases.

Bilobed gallbladder was reported in 0.4\% of its cases. A study conducted by Nadeem in UAE reported bilobed gallbladder in 2\% of its cases.\textsuperscript{24} Septate gallbladder was noted in 4.1\% of our cases, a finding that is in concordance with Blumberg, Spech et al, where the incidence of septate gallbladder was also 4\%.\textsuperscript{25,26}

A study conducted in Turkey by Zehra and her colleagues reported variations in the course of the pancreatic duct. Results showed 62.5\% for descending versus 75.4\% at our present study, 30\% for sigmoid versus 18.7\% at our present study, 5.5\% for vertical versus 4.4\% at our present study and 2\% for loop versus 1.2\% at our present study.\textsuperscript{27}

In Korea, a study was carried out to detect anomalies of the pancreaticobiliary ducts of Koreans. Choledocal cyst was identified in 0.32\% of its cases versus 1.2\% in the current study. The clinical significance of this anomaly lies in the remarkably high incidence of malignant transformation in the wall of the cyst.\textsuperscript{28}

A study conducted in Taiwan, has shown that drainage of the right posterior segment into the left hepatic duct before its confluence with the right anterior segmental duct was the most frequent anatomical variation of the intrahepatic biliary system with a prevalence of 13.0\%.\textsuperscript{19} This is contrary to findings in our present study, which revealed this anatomic variant in only 0.6\% of its cases. Triple confluence is characterized by emptying of the right anterior segmental duct, right posterior segmental duct, and left hepatic duct into the common hepatic duct simultaneously. The study of Sinyi et al in China revealed triple confluence in 9.1\% of its cases, as compared to 8.5\% in our study.\textsuperscript{19} Another study conducted in Europe-France by Denis Castign reported triple confluence in 12\% of its cases.\textsuperscript{29} Similarly, Koenrad et al from the United States reported the prevalence of triple confluence as 11\% in a pictorial essay.\textsuperscript{7} In the Middle East three studies were conducted in Iran, Saudi Arabia and turkey and found the prevalence of this variant to be 21.5\%, 10.7\%, and 8.01\%, respectively.\textsuperscript{30,32} These results do not seem to follow a certain geographic pattern.

In the current study, no correlation has been noted between anatomical variations and gender, except in the pancreatic duct course (loop); a statistically significant association has been found between gender and looped pancreatic duct course with a p value of 0.026. This variant was present only in males in the present study. In comparison to this, Zehra et al, concluded in their study a statistically significant association between gender and pancreatic duct vertical course (p=0.004), with a higher frequency in females than males, and no significant association was noted with other pancreatic duct course.\textsuperscript{27}
CONCLUSION

Anomalies and variations of the pancreatico-biliary system are relatively common, in which the standard, so-called normal anatomy was found in 56.64% of patients. This accentuates the significance of recognizing these anomalies prior to and during surgeries, to avoid unnecessary and preventable iatrogenic injuries and complications.

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Ethical approval: The study was approved by the Institutional Ethics Committee

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