Investigation of Biliary Canal Variations as a Cause of Stone Formation in the Choledochal Canal

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Abstract: The present study aimed to investigate if there is an association between the diameter of the choledochal duct and choledochal duct stone formation. The present study consisted of 79 patients who had endoscopic interventions and MRCP procedure with surgery history. Some followed due to disorders of the liver, gall bladder, and biliary tract and some of whom presented hepatobiliary complaints between 2017 and 2019. The choledochal duct diameter measured from MRCP images and choledochal duct stone had examined; the type classified according to Huang classification. Among the cases classified, 29 patients, was Huang Type A1, 27 patients were Huang Type A2, 16 patients were Huang Type A3, and seven patients were Huang Type A4. There was not any statistically significant association in terms of choledochal duct diameter regarding the types. Choledochal duct diameter was statistically higher in female patients than male patients (p<0.05). According to the age group, a statistically significant difference detected for choledochal duct stone formation; individuals over 45 years of age present an increase for choledochal duct stone (p<0.05). The choledochal duct diameter was found higher in female patients compared with male patients; stone formation has found increased in both gender over 45 years of age. It should consider before surgical procedures and radiological tests.

Keywords: Biliary tract variations; choledochal diameter; choledochal stone.

INTRODUCTION

Biliary stone disease is a common gastrointestinal problem. Prolongation of the average life period increases the older population; however, an increase in prevalence had also detected due to the changes in nutritional habits¹. Gallbladder stone is one of the most common causes of elective abdominal procedures in general surgery. Although mortality is lower in gall bladder procedures, financial and health effects should consider due to higher morbidity rates. Despite advances in techniques in liver surgeries, biliary complications are a significant cause of morbidity and mortality. It is exceedingly essential to know the anatomical structures very well to reduce mortality and conduct the treatment correctly². Risks in terms of the gallbladder and liver surgery and transplantation surgery, radiologists interpreted
percutaneous biliary tract drainage anomalies as opaque substance distribution or difficulties in their drainage. If the aim is to increase success and decrease liver operations and transplantation complications, we need to know the anatomy of intrahepatic and extrahepatic bile ducts\(^3,4,5\). Previous studies investigated the relationship between the diameter of the ductus cysticus and the angle between the ductus cysticus and the gallbladder with stone formation. Also, studies investigated the anatomical structures and variational developments\(^6,7,8\). Despite these studies' availability, there are no adequate studies in the literature in recent decades regarding the relationships between biliary canal variations and stone formation. Under the light of these data, we aimed to classify biliary tract formations according to Huang variation types and investigate any possible association between common bile duct (CBD) diameter, CBD stone, and gender as well as age.

**MATERIALS AND METHODS**

After approval of the Ethical Committee of Kocaeli University with GOKAEK-2018/18. 2018/86 dated and numbered decision, the present study included 79 patients who had endoscopic interventions and MRCP procedure with surgery history. Some of whom followed due to disorders of the liver, gall bladder, and biliary tract and some of whom presented hepatobiliary complaints in Derince Training and Research Hospital within Health Sciences University between 2017 and 2019. After reviewing age, gender, clinical presentation, history of any previous surgery has obtained in 79 patients, CBD diameter and cholelithiasis reviewed on MRCP images in each case. The measurements on the images performed by the same radiologist and taking the same points for measurement considered for each case. CBD diameter had measured from the middle point of the distance from the junction point of the cystic duct (CD) and common hepatic duct (CHD) to ampulla of Vater.

The biliary tract has divided into two-part included intrahepatic and extrahepatic biliary tracts. As identified by Couinaud, the liver consists of eight individual segments with individual portal circulation and venous blood flow\(^1\). It also contains the anatomically segmental structure of the liver. Huang classification has used to classify anatomic variations of the biliary tract. This method is one of the classification methods used for radiological measurements and examinations. Variational states of the biliary tracts were identified and classified differently by the researchers. Couinaud, Champetier, Onkubu, Choi, and Huang made type classifications. Variation classification of the biliary tract is usually done depending on the junction status of the right posterior hepatic duct in general\(^9\). We used Huang classification in our study. In addition to descriptive statistics, multivariate analysis (binary logistic regression) has used to determine the analytical parameters. P-value<0.05 in multivariate analysis was considered to a significant association.

**RESULTS AND DISCUSSION**

The participants included 38 females (53.04±19.56 years) and 41 males (54.7±20.75 years). The youngest female patient was 22 years old, and the eldest was 87 years old—the median age value of 56 years. The youngest male case was 19 years old, and the most adult male was 91 years old—the median age value of 58 years.
Classification of anatomical variation of the biliary tract revealed the following; Female patients, 11 (27.5%) patients in Huang A1 type, 20 (50%) patients in Huang A2 type, 5 (17.5%) patients in Huang A3 type, and 2 (5%) patients in Huang A4 type. In the male patients, 18 (43.9%) patients were Huang A1 type, 7 (17.07%) patients were Huang A2 type, 11 (26.8%) patients were Huang A3 type, 5 (12.1%) patients were Huang A4 type.

Accordingly, 29 (36.70%) patients were Huang A1 type, 27 (34.2%) patients were Huang A2 type, 16 (20.3%) patients were Huang A3 type, 7 (8.64%) patients were Huang A4 type.

Table 1. Investigation of the Association Between Variation Type and Common Bile Duct Stone

<table>
<thead>
<tr>
<th>Variation Type</th>
<th>A1</th>
<th>A2</th>
<th>A3+A4</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Common Bile Duct Stone</td>
<td>No</td>
<td>20</td>
<td>16</td>
<td>16</td>
</tr>
<tr>
<td>Yes</td>
<td>9</td>
<td>11</td>
<td>7</td>
<td>0.678</td>
</tr>
<tr>
<td>Total</td>
<td>29</td>
<td>27</td>
<td>23</td>
<td></td>
</tr>
</tbody>
</table>

There was no difference in common bile duct stone according to the various types (p>0.05).

Table 2. The Difference Between CBD Diameter and CHD-CD Angle Investigated According to the Age Above and Below 45 Years

<table>
<thead>
<tr>
<th>Group</th>
<th>Age</th>
<th>N</th>
<th>Mean</th>
<th>Std. Deviation</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>CHD Diameter</td>
<td>45≥</td>
<td>24</td>
<td>5.25</td>
<td>3.86</td>
<td>0.001</td>
</tr>
<tr>
<td></td>
<td>45&lt;</td>
<td>55</td>
<td>8.68</td>
<td>4.71</td>
<td></td>
</tr>
<tr>
<td>CHD-CD Angle</td>
<td>45≥</td>
<td>17</td>
<td>45.17</td>
<td>30.08</td>
<td>0.510</td>
</tr>
<tr>
<td></td>
<td>45&lt;</td>
<td>38</td>
<td>39.34</td>
<td>29.87</td>
<td></td>
</tr>
</tbody>
</table>

A significant increase detected in CHD diameter over 45 years of age (p<0.05). According to the age group, there was no difference detected in the CHD-CD angle (p>0.05).

Table 3. Investigation of the Difference for CHD Stone Formation According to the Age Above and Below 45 Years

<table>
<thead>
<tr>
<th>CHD Stone Present</th>
<th>Age_Group (years)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>45&gt; years</td>
<td>45&lt; years</td>
</tr>
<tr>
<td>No</td>
<td>20</td>
<td>32</td>
</tr>
<tr>
<td>Yes</td>
<td>4</td>
<td>23</td>
</tr>
</tbody>
</table>

A statistically significant difference detected in CHD stone formation according to the age group (p<0.05). CHD stone incidence increases in patients over 45 years of age.
Table 4. Analysis of the Association Between CHD-CD Angle and Age as well as CHD Diameter

<table>
<thead>
<tr>
<th></th>
<th>Age</th>
<th>CHD Diameter</th>
<th>CHD-CD Angle</th>
<th>CHD Stone Dimension</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age CHD</td>
<td>0.00</td>
<td>0.43</td>
<td>0.154</td>
<td></td>
</tr>
<tr>
<td>CHD Diameter</td>
<td>0.000</td>
<td>0.106</td>
<td>0.003</td>
<td></td>
</tr>
<tr>
<td>CHD-CD Angle</td>
<td>0.434</td>
<td>0.106</td>
<td>0.752</td>
<td></td>
</tr>
<tr>
<td>CHD Stone Dimension</td>
<td>0.154</td>
<td>0.003</td>
<td>0.752</td>
<td></td>
</tr>
</tbody>
</table>

A positive correlation was detected between the age and CHD diameter and between CHD stone dimension and CHD diameter (p<0.05). There was no significant correlation between CHD-CD angle and age and between CHD-CD angle and CHD stone dimension (p>0.05).

Table 5. Comparison of CHD Diameter (mm) According to the Gender

<table>
<thead>
<tr>
<th>Gender</th>
<th>N</th>
<th>Mean (mm)</th>
<th>Std. Deviation</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>CHD Diameter</td>
<td>Male</td>
<td>41</td>
<td>6.28</td>
<td>4.15</td>
</tr>
<tr>
<td></td>
<td>Female</td>
<td>38</td>
<td>9.10</td>
<td>4.91</td>
</tr>
</tbody>
</table>

Mean CHD diameter was measured 9.1±4.91 mm in female patients and 6.28±4.15 mm in male patients. The mean CHD diameter measured at 7.67 mm.

Table 6. Comparison of Biliary Vesicle Stone Depending on the Gender

<table>
<thead>
<tr>
<th>Biliary Vesicle Stone Formation</th>
<th>Gender</th>
<th>Male</th>
<th>Female</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>No</td>
<td></td>
<td>23</td>
<td>8</td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td></td>
<td>10</td>
<td>18</td>
<td>0.005</td>
</tr>
<tr>
<td>Operated</td>
<td></td>
<td>8</td>
<td>12</td>
<td></td>
</tr>
</tbody>
</table>

A significant difference found between female and male groups for biliary vesicle stone depending on gender (p<0.05). The biliary vesicle stone formation was significantly higher in female patients (p<0.05).

The biliary tract may present different anatomic variations both in intrahepatic and extrahepatic locations beyond its normal structure. Biliary tract junction forms may also present different variations; however, the normal anatomic structure has detected in approximately 58% of the global population\textsuperscript{10,11}. Beyond identified anatomic pattern, the biliary tract may present intrahepatic or extrahepatic developmental variations\textsuperscript{12}; since such variations may potentially cause problems in surgical procedures in the patients, they may increase the risk of iatrogenic injury during surgical procedures such as open and laparoscopic gall bladder procedures, liver resection or liver transplantation from an alive donor. Different anatomic variations demonstrated to be associated with stone formation, recurrent
pancreatitis, cholangitis, and malignancies\textsuperscript{11,13}. Recognition of all gallbladder variations and possible association of such variations with cholelithiasis is significant to reduce the risks for malpractice and reduce mortality and complication rates.

The present study investigated any possible association between variations of the extrahepatic biliary tract and cholelithiasis. There was no statistically significant association between Huang variation types and cholelithiasis (Table 1). In the present study, any possible correlation between CHD diameter, CHD-CD angle, and CHD stone dimension; consequently, a statistically significant correlation has detected. Another statistically significant output of such correlation was the increase of stone formation in individuals over 45 years of age. Many studies had conducted to search the association between CHD diameter and age, body weight, height as well as BMI. Kaude measured the diameter of the common bile duct on 600 individuals by ultrasound; CHD diameter was 2.8 mm in the individuals below 20 years of age, whereas CHD diameter was measured 4.1 mm in the individuals over 71 years age\textsuperscript{14}. Recent studies emphasize the effect of aging on CHD diameter. Bachar et al. detected an age-dependent change in CHD diameter by an annual dilatation of 0.04 mm through ultrasound\textsuperscript{15}. Daradkeh et al. reported that the factors affecting common bile duct diameter were age, cholecystectomy, and BMI\textsuperscript{16}.

Tom et al. conducted a study on 187 individuals and measured a significant CHD diameter increase by aging\textsuperscript{17}. Wu et al., Kaim et al., Niederau et al., and Bowie et al. carried out studies about CHD diameter increase by aging and their possible causes\textsuperscript{18-21}. Kaim et al. stated that age-dependent increase in CHD diameter might appear due to age-dependent increase of the loss in the reticulo-elastic web and the destruction of longitudinally extending myositis bands and destruction on intermediate connective tissue\textsuperscript{11}. Adibi and Givechian examined the association between CHD diameter and age, BMI, portal vein diameter, and drug addiction; they observed that CHD diameter increases along with wide portal vein in drug addicts in the advanced age\textsuperscript{22}(Table2). In the present study, stone formation increased in individuals over 45 years of age (Table 3). Many studies of the literature showed that cholelithiasis prevalence increased by age. Attili et al. reported a linear increase in cholelithiasis and cholecystectomy in both genders by aging\textsuperscript{23}.

Barbara et al. stated that cholelithiasis' prevalence increases by aging between 18 and 65 years of age\textsuperscript{24}. This is probably because stone formation becomes visible and manifest in the elder ages because of being a gradual and time-taking process. In our study, it measured that individuals with cholelithiasis have increased DCH diameter (Table 4). A statistically significant association of stone formation has observed with CHD diameter increase. This has been explained by the presence of the stone to produce a physical dilatation of the DCH. Boys et al. divided CHD diameter into three groups included 6 mm, 6 to 9.9 mm, and over 10 mm; they measured the diameter smaller in the individuals without CHD stone. In that study, the stone formation rate was 14% in the first and second groups and 39% in the third group\textsuperscript{25}.

The previous studies show differences in anatomic formations where the stone has developed in male and female patients. A significant difference has found in the CHD diameter in MRCP images between female and male patients within this context. The normal range for CHD diameter is 4 to 6 mm; CHD diameter below 5 mm has accepted average whereas a diameter at and above 8 mm is dilated. Mean CHD diameter was measured 9.10±4.91 mm in female patients and 6.28±4.15 mm in male patients. Poralla et al. measured CHD diameter and the pressure inside the
canal in their study in both genders; both CHD diameter and the pressure inside the canal were detected higher in male patients than female patients\textsuperscript{26}. Matcuk et al. reported that CHD diameter is higher in females, and it increases in advanced age and in those who had cholecystectomy in both genders\textsuperscript{27}. The researchers state that cholelithiasis incidence is more in female patients with differences in biliary tract\textsuperscript{28}. This may be expressed with a full CHD diameter in females compared with males (Table 5).

In this study, cholelithiasis rate was statistically higher in female patients than male patients of the present study; the incidence is 2.3-fold higher in female patients (Table 6). Many studies focused on possible risk factors or underlying causes for cholelithiasis among female and male patients. Palermo et al. conducted a research of 1,875 volunteers to investigate the association between cholelithiasis, age, gender, BMI, family history, sedentary life; cholelithiasis incidence was 2.6-fold higher in female volunteers\textsuperscript{29}. Another previous study reported that age, BMI, history of cholic pain, family history, smoking, and hepatosteatosis were all effects on cholelithiasis and radiological anatomy of the biliary vesicle and biliary duct\textsuperscript{30,31}. Yıldırım in 2008 conducted a study under the title of "Incidence of cholelithiasis in adults in the province of Tokat and possible risk factors" on 1,095 individuals in Tokat province of our country; he investigated the effect of gender, age, glucose, ALT, AST, total cholesterol, triglyceride, BMI and anthropometric measurements on the frequency of cholelithiasis; he also evaluated anatomic formations and clinical findings through ultrasound. An association has detected between cholelithiasis and female gender, age, fasting glucose, total cholesterol, BMI, triglycerides, and waist circumference\textsuperscript{32}. Some researches reported that stone prevalence in women is 1.7 to 4-fold more than men\textsuperscript{33}. In our country, Sezer, in 2016, carried out a study with 865 volunteers and detected biliary vesicle stone in 52 volunteers included 41 female individuals\textsuperscript{34}.

Studies indicate that higher cholelithiasis incidence in women may be associated with some factors such as fertility and sex hormones, an increase of estrogen hormone increases cholesterol secretion and causes cholesterol supersaturation\textsuperscript{35}. Some studies suggest that the risk of cholelithiasis increases in women receiving hormone replacement therapy\textsuperscript{36,37}. Some studies demonstrated the association between oral contraceptive use and an increase in the risk of cholelithiasis\textsuperscript{38,39}. A previous study showed that cholesterol and calcium bilirubin crystals in the bile sludge increase by more than 30\%\textsuperscript{40}. According to the outcomes stated above, gender should consider in the patients presenting colic pain without symptoms of cholelithiasis; furthermore, MRCP images show that CHD diameter may be more comprehensive in women when compared with men, which may increase the rate of small stone accumulation and stone formation in the biliary vesicle. Consideration of possible dilatation in CHD diameter by aging reveals that age may be a determinant factor in CHD diameter and stone formation in MRCP or radiological images.

This study is a retrospective study and had done with the measurements made on the images previously taken in the hospital. Limitations in this study have been that the available figures are not of the desired quality, and the number of patients is low. The lack of previous studies on this subject is one of the limitations of our research.
CONCLUSION

Review of the findings of the present study and literature information, a full CHD diameter may not necessarily indicate a stone; further studies will need to investigate the effect of CHD stone existence on gender, and the impact of age on diameter increase.

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CONFLICT OF INTEREST

The author report no conflicts of interest and no funding resources in this study.

REFERENCES


