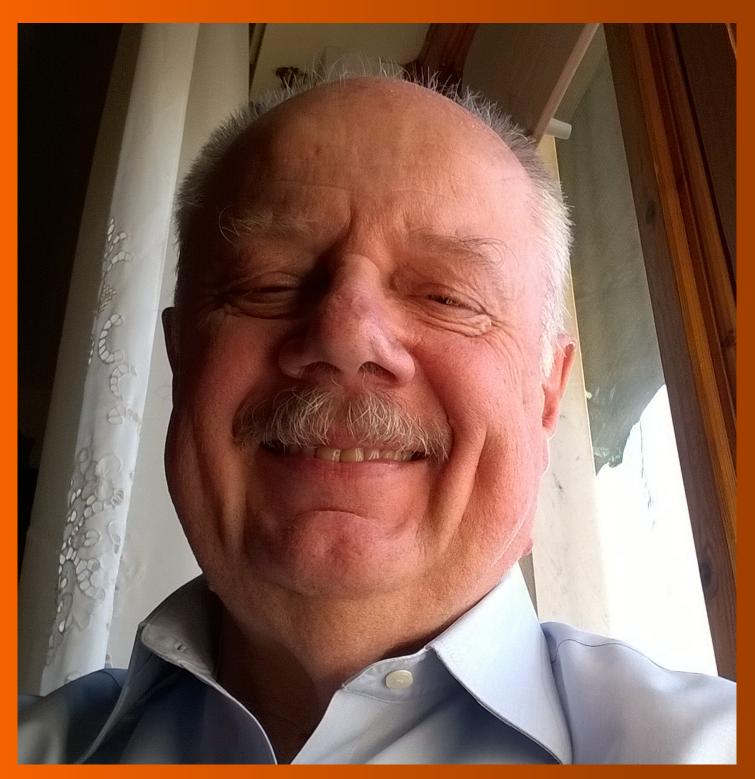
# World Journal of *Gastroenterology*

World J Gastroenterol 2017 April 7; 23(13): 2269-2452





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# United Sector of World Journal of Gastroenterology

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AIMS AND SCOPE	ISSN 2219-2840, DOI: 10.3748) is a peer-re- lished on October 1, 1995. It is published we The <i>WJG</i> Editorial Board consists of 1375 from 68 countries. The primary task of <i>WJG</i> is to rapidly p and commentaries in the fields of gastroen copy, gastrointestinal surgery, hepatobiliary testinal radiation oncology, gastrointestinal apy, gastrointestinal infectious diseases, gas pathophysiology, gastrointestinal laborato ogy, gastrointestinal immunology, gastrointestinal translational medicine, gast therapeutics. <i>WJG</i> is dedicated to become	<i>roenterol, WJG</i> , print ISSN 1007-9327, online viewed open access journal. <i>WJG</i> was estab- ekly on the 7 <sup>th</sup> , 14 <sup>th</sup> , 21 <sup>st</sup> , and 28 <sup>th</sup> each month. experts in gastroenterology and hepatology publish high-quality original articles, reviews, terology, hepatology, gastrointestinal endos- surgery, gastrointestinal oncology, gastroin- imaging, gastrointestinal oncology, gastroin- strointestinal pharmacology, gastrointestinal , evidence-based medicine in gastroenterol- ry medicine, gastrointestinal molecular biol- stinal microbiology, gastrointestinal genetics, rointestinal diagnostics, and gastrointestinal an influential and prestigious journal in gas- he development of above disciplines, and to and expertise of clinicians.	
INDEXING/ABSTRACTING	Science Citation Index Expanded (also known Medicus, MEDLINE, PubMed, PubMed Cen Open Access Journals. The 2015 edition of Jo Reuters (ISI) cites the 2015 impact factor for W	dexed in Current Contents <sup>®</sup> /Clinical Medicine, as SciSearch <sup>®</sup> ), Journal Citation Reports <sup>®</sup> , Index tral, Digital Object Identifier, and Directory of ournal Citation Reports <sup>®</sup> released by Thomson JG as 2.787 (5-year impact factor: 2.848), rank- rology and hepatology (quartile in category Q2).	
FLYLEAF I-IX	Editorial Board		
EDITORS FOR Respon		le Science Editor: Ze-Mao Gong Editorial Office Director: Jin-Lei Wang	
NAME OF JOURNAL	CA 90822, United States	http://www.wjgnet.com	
World Journal of Gastroenterology ISSN ISSN 1007-9327 (print) ISSN 2219-2840 (online)	EDITORIAL BOARD MEMBERS All editorial board members resources online at http:// www.wignet.com/1007-9327/editorialboard.htm	PUBLICATION DATE April 7, 2017 COPYRIGHT	
LAUNCH DATE October 1, 1995	EDITORIAL OFFICE Jin-Lei Wang, Director Yuan Qi, Vice Director	© 2017 Baishideng Publishing Group Inc. Articles pub- lished by this Open-Access journal are distributed under the terms of the Creative Commons Attribution Non-	
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World J Gastroenterol 2017 April 7; 23(13): 2424-2434

ISSN 1007-9327 (print) ISSN 2219-2840 (online)

DOI: 10.3748/wjg.v23.i13.2424

ORIGINAL ARTICLE

### Prospective Study Therapeutic experience of 289 elderly patients with biliary diseases

Zong-Ming Zhang, Zhuo Liu, Li-Min Liu, Chong Zhang, Hong-Wei Yu, Bai-Jiang Wan, Hai Deng, Ming-Wen Zhu, Zi-Xu Liu, Wen-Ping Wei, Meng-Meng Song, Yue Zhao

Zong-Ming Zhang, Zhuo Liu, Li-Min Liu, Chong Zhang, Hong-Wei Yu, Bai-Jiang Wan, Hai Deng, Ming-Wen Zhu, Zi-Xu Liu, Wen-Ping Wei, Meng-Meng Song, Yue Zhao, Department of General Surgery, Beijing Electric Power Hospital, State Grid Corporation of China, Capital Medical University, Beijing 100073, China

Author contributions: Zhang ZM, Liu Z, Liu LM, Zhang C, Yu HW, Wan BJ, Deng H, Zhu MW, Liu ZX, Wei WP, Song MM and Zhao Y performed the study; Liu Z collected and analyzed the data; and Zhang ZM provided the funding, designed the study and wrote the manuscript.

Supported by the Science and Technology Project of State Grid Corporation of China, No. SGHB0000AJJS1400182.

**Institutional review board statement:** This study was reviewed and approved by the Ethics Committee of Beijing Electric Power Hospital, State Grid Corporation of China, Capital Medical University, Beijing, China.

Clinical trial registration statement: The clinical trial is registered with the Research Registry under the Research Registry Unique Identifying Number: researchregistry 2032. Details can be found at http://www.researchregistry. com /browse-the-registry.html#user-researchregistry/ ?view\_26\_search=researchregistry2032&view\_26\_page=1.

Informed consent statement: All study participants, or their legal guardian, provided written consent prior to study enrollment.

**Conflict-of-interest statement:** The authors declare that there are no conflicts of interest related to this study.

Data sharing statement: No additional data are available.

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Received: November 13, 2016 Peer-review started: November 13, 2016 First decision: December 29, 2016 Revised: January 14, 2017 Accepted: February 17, 2017 Article in press: February 17, 2017 Published online: April 7, 2017

#### Abstract

#### AIM

To present clinical characteristics, diagnosis and treatment strategies in elderly patients with biliary diseases.

#### **METHODS**

A total of 289 elderly patients with biliary diseases were enrolled in this study. The clinical data relating to these patients were collected in our hospital from June 2013 to May 2016. Patient age, disease type, coexisting diseases, laboratory examinations, surgical methods, postoperative complications and therapeutic outcomes were analyzed.

#### RESULTS

The average age of the 289 patients with biliary diseases



was  $73.9 \pm 8.5$  years (range, 60-102 years). One hundred and thirty-one patients (45.3%) had one of 10 different biliary diseases, such as gallbladder stones, common bile duct stones, and cholangiocarcinoma. The remaining patients (54.7%) had two types of biliary diseases. One hundred and seventy-nine patients underwent 9 different surgical treatments, including pancreaticoduodenectomy, radical resection of hilar cholangiocarcinoma and laparoscopic cholecystectomy. Ten postoperative complications occurred with an incidence of 39.3% (68/173), and hypopotassemia showed the highest incidence (33.8%, 23/68). One hundred and sixteen patients underwent non-surgical treatments, including anti-infection, symptomatic and supportive treatments. The cure rate was 97.1% (168/173) in the surgical group and 87.1% (101/116) in the non-surgical group. The difference between these two groups was statistically significant ( $\chi^2$  = 17.227, *P* < 0.05).

#### CONCLUSION

Active treatment of coexisting diseases, management of indications and surgical opportunities, appropriate selection of surgical procedures, improvements in perioperative therapy, and timely management of postoperative complications are key factors in enhancing therapeutic efficacy in elderly patients with biliary diseases.

**Key words:** Biliary diseases; Elderly patients; Surgical treatment; Non-operative treatment

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**Core tip:** With global prolongation of life expectancy, biliary diseases in elderly patients appear to be common and multifaceted. The purpose of this study was to determine the clinical characteristics and treatment strategies in elderly patients with biliary diseases in order to improve diagnosis and treatment. Active treatment of coexisting diseases, management of indications and surgical opportunities, appropriate selection of surgical procedures, improvements in perioperative therapy, and timely management of postoperative complications are key factors in enhancing therapeutic efficacy in elderly patients with biliary diseases.

Zhang ZM, Liu Z, Liu LM, Zhang C, Yu HW, Wan BJ, Deng H, Zhu MW, Liu ZX, Wei WP, Song MM, Zhao Y. Therapeutic experience of 289 elderly patients with biliary diseases. *World J Gastroenterol* 2017; 23(13): 2424-2434 Available from: URL: http://www.wjgnet.com/1007-9327/full/v23/i13/2424.htm DOI: http://dx.doi.org/10.3748/wjg.v23.i13.2424

#### INTRODUCTION

With global prolongation of life expectancy, the

incidence of diseases in the aging population has increased. Since 1999, biliary diseases in elderly patients have become common and multifaceted in China, with a morbidity rate of 8%-11%<sup>[1]</sup>.

Due to lowered stress response, an increase in coexisting diseases, rapid progression, and poor surgical tolerance, the risk of surgery for biliary diseases in elderly patients has increased<sup>[2,3]</sup>. In particular, a subset of elderly patients may be at higher risk of developing postoperative complications and increased mortality due to the loss of physiological function and physical reserves<sup>[4,5]</sup>. Therefore, there is an unmet clinical need to characterize biliary diseases in elderly patients, in order to improve diagnosis and treatment for these diseases<sup>[6-10]</sup>.

In the present study, the clinical data of 289 elderly patients with biliary diseases admitted to our hospital over the past 3 years were summarized and analyzed.

#### MATERIALS AND METHODS

#### Clinical data

A prospective study of 289 elderly patients with biliary diseases, aged 60-102 years, including 129 males (44.6%) and 160 females (55.4%), was conducted in the Department of General Surgery, Beijing Electric Power Hospital, State Grid Corporation of China, Capital Medical University from June 2013 to May 2016.

The inclusion criterion was a diagnosis of biliary disease in elderly patients. According to surgical indications or contraindications, patients were randomly assigned to the surgical group and the non-surgical group.

The exclusion criterion was patients who were voluntarily discharged from hospital due to refusal to cooperate with the assigned group therapy.

The study was approved by the Ethics Committee of Beijing Electric Power Hospital, State Grid Corporation of China, Capital Medical University, Beijing, China.

#### Laboratory examinations

Following admission, blood samples were obtained from all patients for laboratory examinations, including routine blood test, high sensitivity C-reactive protein (hs-CRP) level<sup>[11,12]</sup>, liver function, tumor markers, and blood clotting function.

#### Imaging examination

Prior to treatment, all patients were examined by B ultrasound, computed tomography (CT) and magnetic resonance cholangiopancreatography (MRCP) to determine the presence of gallbladder or bile duct stones, inflammation or tumor. Double-layer structure of gallbladder wall revealed by B ultrasound suggested the existence of acute suppurative and gangrenous cholecystitis. Stones, inflammation, tumors of the gallbladder and bile duct, and thickness of the gallbladder wall can be determined by CT scanning<sup>[11]</sup>.



Table 1 Comparison of age distribution in surgical and non- surgical groups, $n$ (%)				
Age (yr)	Surgical group	Non-surgical group	χ²	P value
60-69	75 (43.4)	25 (21.6)	23.376	0.000
70-79	64 (37.0)	42 (36.2)		
80-89	30 (17.3)	45 (38.8)		
90-99	4 (2.3)	3 (2.6)		
$\ge 100$	0 (0.0)	1 (0.9)		
Total	173 (100.0)	116 (100.0)		

# Table 2 Distribution of disease type in surgical and non-surgical groups, n (%)

Type of disease	Surgical group	Non-surgical group	χ²	<i>P</i> value
Gallbladder stones	133 (36.8)	66 (32.7)	0.308	0.016
Common bile duct stones	41 (11.4)	23 (11.4)		
Hepatolithiasis	4 (1.1)	7 (3.5)		
Chronic cholecystitis	36 (10.0)	14 (6.9)		
Acute cholecystitis	92 (25.5)	67 (33.2)		
Gallbladder polyps	8 (2.2)	0 (0.0)		
Acute cholangitis	16 (4.4)	16 (7.9)		
Biliary pancreatitis	15 (4.2)	3 (1.5)		
Biliary tract tumors	12 (3.3)	3 (1.5)		
Gallbladder carcinoma	4 (1.1)	3 (1.5)		

A single patient may have 1-2 types of biliary diseases.

Table 3 Proportion of types of diseases in surgical and non- surgical groups, $n$ (%)					
Group	п	One type of disease	Two types of diseases	$\chi^2$	<i>P</i> value
Surgical Non-surgical	173 60 (51.7)	71 (41.0) 56 (48.3)	102 (59.0)	3.198	0.074

MRCP helps to identify stones and obstruction of intrahepatic and extrahepatic bile ducts<sup>[13]</sup>.

Based on the patient's condition, electrocardiography (ECG), dynamic ECG, echocardiography, neck vascular ultrasound, head magnetic resonance angiography, and head and neck CT angiography were also performed<sup>[14,15]</sup>.

#### Therapeutic methods

According to the individual patient's status, appropriate surgical procedures were selected, including pancreaticoduodenectomy, radical resection of hilar cholangiocarcinoma, and laparoscopic cholecystectomy (LC).

Non-surgical treatment strategies included, but were not limited to, anti-infection, symptomatic and supportive treatments.

#### Statistical analysis

SPSS 18.0 was used for statistical analysis<sup>[16]</sup>. The data from laboratory tests were expressed as mean  $\pm$  SD, using the Student's *t*-test for comparisons between the surgical and non-surgical groups. The

rate (%) or constituent ratio was used for count data related to the distribution of age, disease types and coexisting diseases, surgical procedures, postoperative complications and therapeutic efficacy, and the  $\chi^2$  test was used for comparisons between two groups. A *P* value < 0.05 was considered statistically significant.

#### RESULTS

#### Age distribution

The age of the 289 patients with biliary diseases ranged from 60 to 102 years, with an average age of  $73.9 \pm 8.5$  years.

In the surgical group, the average age of patients was  $72.1 \pm 8.2$  years, ranging from 60 to 95 years. The maximum age of patients undergoing open surgery was 95 years, and was 86 years for patients undergoing LC.

In the non-surgical group, the age of the patients ranged from 60 to 102 years, with an average age of  $76.5 \pm 8.2$  years.

There was a significant difference in the age distribution between the two groups (P < 0.05), as shown in Table 1.

#### Types of biliary diseases

The 289 patients had 1 or 2 types of 10 different biliary diseases, including gallbladder stones, common bile duct stones, and cholangiocarcinoma (Table 2).

One hundred and thirty-one (45.3%) patients had one type of biliary disease, and 158 (54.7%) patients had two types of biliary diseases. There was a significant difference in the distribution of types of disease (P < 0.05) in the surgical group compared with the non-surgical group (Table 2). However, there was no significant difference in the proportion of disease types (P > 0.05) between the two groups (Table 3).

#### **Coexisting diseases**

Among the 289 patients, 223 (77.2%) had 1-6 types of 13 various coexisting diseases, such as hypertension, coronary heart disease (CHD), and diabetes. Of these patients, 128 (57.4%) were in the surgical group, and 95 (42.6%) were in the non-surgical group. There was no significant difference between the two groups (P > 0.05; Table 4).

#### Laboratory examinations

Using the *t*-test, it was found that the white blood cell count, percentage of neutrophils, prothrombin time and international normalized ratio at admission were significantly decreased (P < 0.05), whereas hemoglobin, albumin and total serum bilirubin were significantly increased in the surgical group compared with the non-surgical group (P < 0.05). There were no differences in the other test parameters between the two groups (P > 0.05; Table 5).

#### Surgical procedures

One hundred and seventy-three patients underwent



Table 4 Distribution of coexisting diseases in surgical andnon-surgical groups, n (%)

Type of coexisting disease	Surgical group	Non-surgical group	$\chi^2$	<i>P</i> value
Coronary heart disease	40 (14.7)	28 (13.5)	17.526	0.131
Atrial fibrillation	8 (2.9)	1 (0.5)		
Hypertension	84 (30.8)	55 (26.4)		
Diabetes	33 (12.1)	22 (10.6)		
Renal insufficiency	1 (0.4)	5 (2.4)		
Chronic bronchitis with	5 (1.8)	9 (4.3)		
emphysema				
Portal hypertension	3 (1.1)	0 (0.0)		
Pulmonary infection	7 (2.6)	10 (4.8)		
Hypoalbuminemia	9 (3.3)	9 (4.3)		
Anemia	8 (2.9)	5 (2.4)		
Electrolyte disorder	21 (7.7)	21 (10.1)		
Cerebral infarction	30 (11.0)	28 (13.5)		
Hepatic insufficiency	24 (8.8)	15 (7.2)		

A single patient may have 1-6 types of coexisting diseases.

surgical treatment with 1 of 9 different surgical procedures (Table 6), including pancreaticoduodenectomy (Figure 1), radical resection of hilar cholangiocarcinoma, and LC (Figure 2). Of these patients, 105 underwent simple LC, 7 of which were LC plus endoscopic sphincterotomy (EST) or endoscopic papillary balloon dilation (EPBD)<sup>[17-19]</sup>. The success rate was 100%, and none of the patients required conversion to laparotomy.

#### Postoperative complications

Ten types of postoperative complications were recorded, including pulmonary infection, hypopotassemia, incision fat liquefaction and others, with an incidence of 39.3% (68/173). Of these complications, hypopotassemia had the highest incidence (33.8%, 23/68), followed by hypoalbuminemia (22.1%), and pulmonary infection (19.1%) (Figure 3).

#### Therapeutic outcomes

Among these 289 elderly patients, 269 (93.1%) patients were cured, 11 (3.8%) patients improved, and 9 patients (3.1%) died.

In the surgical group, 168 patients were cured, with a cure rate of 97.1% (168/173). Five patients died at 4, 6, 18, 31 and 100 d after surgery, with 3 who died due to acute myocardial infarction (AMI), 1 due to pulmonary embolism, and 1 due to multiple organ failure. The mortality rate was 2.9% in the surgical group (5/173).

In the non-surgical group, 101 patients were cured, with a cure rate of 87.1% (101/116), and 11 patients improved, with a rate of 9.5% (11/116). Four patients died due to multiple organ failure at 5, 6, 6 and 10 d after admission, resulting in a mortality rate of 3.4% (4/116).

A significant difference in the therapeutic effects between the surgical group and the non-surgical group was observed using the  $\chi^2$  test ( $\chi^2 = 17.227$ , P < 0.05;

Figure 4).

#### DISCUSSION

#### Clinical characteristics of biliary disease in elderly patients

It is known that elderly people have a reduced response to stress, can be insensitive to pain, and have reduced abdominal muscle strength. In elderly patients with acute cholecystitis, acute suppurative cholangitis or other biliary diseases, clinical manifestations and pathological changes are often poorly correlated. These patients typically show mild symptoms, but have severe pathological alterations<sup>[20]</sup>.

Due to decreased defense in the body, lower tension in the gallbladder wall, and difficulty in removing biliary stones, patients can have stones lodged in the neck of the gallbladder under stimulating conditions, *e.g.*, a fatty diet. This can lead to an increase in internal pressure in the gallbladder, causing secondary infection and acute suppurative cholecystitis. In addition, small artery sclerosis in the gallbladder wall can result in an insufficient blood supply which can cause necrotic perforation of the gallbladder, biliary peritonitis, and even toxic shock. Therefore, elderly patients have the characteristics of acute onset and rapid progression of diseases.

Due to reduced immune function, increased incidence of biliary tract tumors and difficult early diagnosis in the elderly, the occurrence of jaundice may cause severe damage to vital organs and the immune system. Thus, elderly patients have a low surgical resection rate, an increase in postoperative complications and high mortality.

In this study, we found that 289 elderly patients had 1-2 types of 10 different biliary diseases. Of these patients, 45.3% had one type of biliary disease, and 54.7% had two types of biliary diseases. These findings indicated that many elderly patients have multiple biliary tract diseases.

#### Active management of coexisting diseases

It has been reported that morbidity due to coexisting diseases in elderly patients with biliary diseases is as high as 65%-90%<sup>[21]</sup>. From our current study, we found that 77.2% of patients had 1-6 types of 13 coexisting diseases, such as hypertension, CHD and diabetes. The incidence of coexisting diseases in the surgical group was 57.4%, and was 42.6% in the non-surgical group. No significant difference in coexisting diseases was observed between the two groups. Our study indicated that by actively managing coexisting diseases, surgery in elderly patients with biliary diseases is safe and feasible<sup>[22]</sup>.

If hypertension coexists with biliary diseases<sup>[23-25]</sup>, perioperative hypertension can enhance bleeding during surgery, induce myocardial ischemia, AMI, stroke, renal failure and other complications. It was recommended by the Anesthesiology Society of the

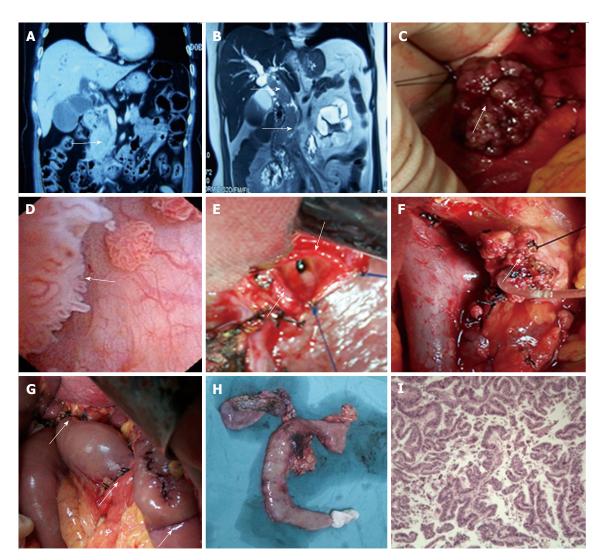


Figure 1 Pancreatoduodenectomy in a 73-year old female patient with bile duct carcinoma. A: Three-dimensional computed tomography reconstruction found an extrahepatic bile duct obstruction (arrow); B: MRI coronal image confirmed entire extrahepatic bile duct obstruction (arrows); C: Tumor (arrow) seen at the incision of the common bile duct; D: Tumor (arrow) seen in the common bile duct by intraoperative choledochoscopy; E: Upper cut end of the hepatic common duct (arrows); F: The cut end of the pancreas (arrow), and the portal vein; G: Child anastomosis (arrows); H: Resected specimen following pancreatoduodenectomy; I: Papillary adenocarcinoma of extrahepatic bile duct, including the hepatic common duct, common bile duct and Vater ampulla. Hematoxylin and eosin staining. Objective magnification, × 40.

Chinese Medical Association in 2014 that preoperative blood pressure should be maintained below 140/90 mmHg. Some researchers have also suggested that preoperative blood pressure should be below 135/85 mmHg, and remain stable for 1-2 wk. It is noteworthy that in the case of non-emergency surgery, patients should cease to use reserpine and anti-hypertensive drug No. 0 and switch to other anti-hypertensive drugs 7 d prior to surgery, as it is possible that use of these drugs can cause catecholamine depletion in the peripheral sympathetic nerve endings, which makes it difficult to improve and maintain blood pressure when low blood pressure occurs during surgery.

If patients with biliary diseases also have CHD<sup>[26-28]</sup>, preoperative ECG, dynamic ECG and echocardiography should be performed before surgery. If cardiac dys-function or arrhythmia can be controlled by drugs, and the ejection fraction is more than 60% with better

cardiac compensatory function, surgical treatment can be considered. If patients have a history of cardiac dysfunction, surgery can only be performed when the illness has been stable for 1 mo. In patients with a history of AMI, surgery should only be performed when the illness has been stable for at least 6 mo. In the case of AMI, the blood levels of cardiac Troponin I (cTnI), brain-type natriuretic peptide (BNP) and N-terminal proBNP (NT-proBNP)<sup>[29-31]</sup> should be evaluated to determine the severity of myocardial injury and heart failure. If the patient has a history of bradycardia, a cardiac pacemaker should be implanted before surgery.

If patients with biliary diseases also have an old cerebral infarction<sup>[32-34]</sup>, neck vascular ultrasound, head magnetic vascular imaging, or head and neck CT angiography should be performed. Carotid artery stenosis should be less than 50%, and pre-, intraand post-operative blood pressure should be kept

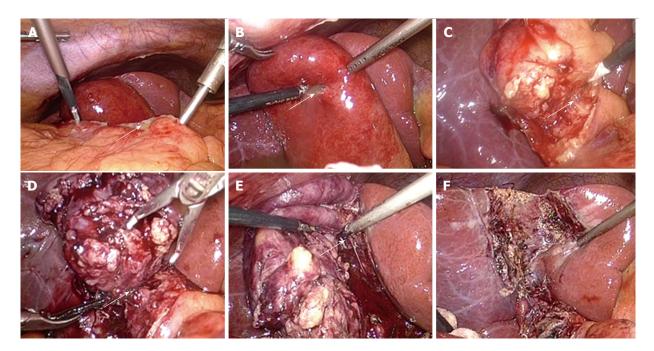


Figure 2 Laparoscopic cholecystectomy in a 78-year-old male patient with acute calculous cholecystitis. A: Pus (arrow) surrounding the gallbladder; B: Pus (arrow) in the lumen of the gallbladder; C: Congestion and edema of the gallbladder wall (arrow); D: Ordinary silk thread to ligate cystic duct (arrow); E: Inflammatory adhesion (arrow) of gallbladder bed; F: Wound after cholecystectomy.

ltem	Surgical group, n = 173	Non-surgical group, n = 116	t	<i>P</i> value
Routine blood				
WBC (× 10 <sup>9</sup> /L)	$8.54 \pm 4.77$	$10.41 \pm 5.58$	-3.054	$0.002^{a}$
N%	$71.94 \pm 14.81$	79.45 ± 13.55	-4.450	$0.000^{a}$
Hb (g/L)	$127.41 \pm 20.44$	$121.00 \pm 20.06$	2.631	0.009 <sup>a</sup>
hs-CRP (mg/L)	$50.02 \pm 76.60$	$52.48 \pm 58.76$	-0.288	0.774
Liver function				
ALT (U/L)	$70.17 \pm 103.96$	$94.13 \pm 139.50$	-1.573	0.117
AST (U/L)	$71.32 \pm 115.81$	$105.00 \pm 190.88$	-1.696	0.092
TP (g/L)	$62.65 \pm 8.33$	$62.06 \pm 8.94$	-0.553	0.581
Alb (g/L)	$36.59 \pm 5.92$	$35.16 \pm 5.75$	2.030	0.043 <sup>a</sup>
TB (μmol/L)	$51.74 \pm 103.87$	$34.33 \pm 37.06$	2.021	$0.044^{a}$
DB (µmol/L)	$21.99 \pm 39.50$	$15.85 \pm 23.37$	1.659	0.098
Tumor marker				
CA19-9 (U/mL)	$340.19 \pm 1317.16$	896.95 ± 6346.89	-0.880	0.380
CEA (ng/mL)	$2.76 \pm 4.13$	$142.66 \pm 1283.35$	-1.005	0.318
Blood clotting function				
APTT (s)	$32.49 \pm 5.20$	$32.59 \pm 7.56$	-0.124	0.901
PT (s)	$11.61 \pm 1.38$	$12.12 \pm 1.54$	-2.739	$0.007^{a}$
INR	$1.05 \pm 0.13$	$1.09 \pm 0.14$	-2.318	0.021 <sup>a</sup>
FIB (g/L)	$3.63 \pm 1.21$	$3.69 \pm 0.90$	-0.449	0.654

<sup>a</sup>*P* < 0.05, surgical group *vs* non-surgical group. Alb: Albumin; ALT: Alanine aminotransferase; APTT: Activated partial thromboplastin time; AST: Aspartate aminotransferase; CA19-9: Carbohydrate antigen 19-9; CEA: Carcinoembryonic antigen; DB: Direct bilirubin; FIB: Fibrin; Hb: Hemoglobin; hs-CRP: High sensitivity C-reactive protein; INR: International normalized ratio; N%: Neutrophil percentage; PT: Prothrombin time; TB: Total bilirubin; TP: Total protein; WBC: White blood cell count.

constant to avoid large fluctuations. This preventive measurement can effectively reduce the incidence of perioperative cerebral infarction and mortality.

In patients with coexisting diabetes<sup>[35,36]</sup>, surgical stress can cause disorders of glucose metabolism and insulin resistance, and even produce ketone bodies or ketoacidosis. Therefore, the use of oral hypoglycemic

agents and intermediate- or long-acting insulin should be stopped and switched to regular insulin. The expert consensus for the perioperative management of blood glucose, proposed by the Anesthesiology Society of the Chinese Medical Association in 2014, suggested that preoperative blood glucose should be below 10 mmol/L<sup>[35]</sup>. In addition, sustained high glucose

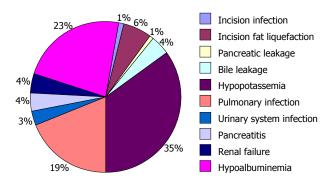


Figure 3 Post-operative complications occurred in 68 elderly patients with biliary diseases.

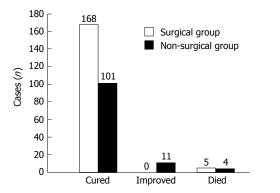


Figure 4 Therapeutic outcomes in the surgical and non-surgical groups.

blood levels can benefit the growth of bacteria, lower immune function, and result in abnormal neutrophil function with reduced phagocytosis. Antibiotics should be administered to prevent infection prior to surgery.

#### Management of indications and surgical opportunities

In elderly patients with biliary diseases combined with coexisting diseases, it is essential to manage indications and surgical opportunities.

In elderly patients with cholecystolithiasis, surgery can be determined based on the size of stones, gallbladder function, history of biliary colic, and whether the disease is complicated by cholangitis or pancreatitis. Surgery should be performed as soon as possible to avoid increased surgical risk due to increased patient age, disease progression and other coexisting diseases.

For patients with choledocholithiasis, surgery can be determined based on the size and number of stones and diameter of the common bile duct. Surgery should be carried out as soon as possible<sup>[37]</sup>.

In patients with biliary tract tumors, surgery should be based on the site of obstruction, the degree of invasion and presence of distant metastasis. When favorable, surgery should be performed as early as possible, in order to improve quality of life and extend life expectancy<sup>[38]</sup>.

In patients with acute cholangitis, conservative treatment should be actively considered in order to change emergency surgery to scheduled surgery as

Table 6 Surgical procedures in elderly patients with biliary diseases Surgical procedure n (%) 6 (3.5) Pancreaticoduodenectomy Radical resection of hilar cholangiocarcinoma 6 (3.5) Radical resection of gallbladder carcinoma 4 (2.3) 4 (2.3) Hepatectomy Open cholecystectomy (OC) 6 (3.5) OC + common bile duct exploration + T-tube 21 (12 1) OC + transcystic common bile duct exploration 14 (8.1) Laparoscopic cholecystectomy (LC) 105 (60.7) LC + endoscopic sphincterotomy or endoscopic 7 (4.0) papillary balloon dilation Total 173 (100.0)

early as possible<sup>[39]</sup>. However, in elderly patients with acute obstructive suppurative cholangitis or acute gangrenous cholecystitis, emergency surgery should be performed as soon as possible to prevent the likely complications of sepsis and even toxic shock<sup>[40-42]</sup>.

Our laboratory tests showed that the white blood cell count, percentage of neutrophils, prothrombin time and international normalized ratio at admission were significantly decreased (P < 0.05), whereas the levels of hemoglobin, albumin and total serum bilirubin were significantly increased in the surgical group compared with the non-surgical group (P < 0.05). These results suggested that the management of indications and surgical opportunities can significantly reduce surgical risk, and avoid unnecessary surgery, especially emergency surgery. Although the results of serum hs-CRP was not significantly different between the two groups (P > 0.05), dynamic monitoring of changes in hs-CRP may result in important reference values<sup>[11,12]</sup> for assessing the progression of acute cholecystitis and in choosing a suitable time window for surgery.

#### Appropriate selection of surgical procedures

The choice of surgical procedure for biliary diseases in the elderly is crucial. It is important to avoid a long disease process or even a second operation following conservative surgery. On the other hand, the increased surgical risk due to aggressive surgery should also be prevented. For example, conservative surgery of acute gangrenous cholecystitis, such as percutaneous cholecystostomy<sup>[43,44]</sup>, an alternative treatment for resolving acute inflammation, is theoretically suitable for patients in poor general condition, with a serious illness, unstable blood pressure during surgery, and other special circumstances. However, percutaneous cholecystostomy should be avoided for the reasons mentioned above. In our treatment group, none of the patients underwent percutaneous cholecystostomy.

Aggressive surgery, such as LC, should be selected with caution in patients with poor heart and pulmonary function before surgery, especially in the presence of emphysema. If LC must be performed, intraoperative pneumoperitoneum pressure should be

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controlled to below 10 mmHg to reduce the impact of pneumoperitoneum on heart and pulmonary function. There is no definitive view on the upper age limit for patients undergoing LC. The oldest patients were a 96-year-old reported by Lee *et al*<sup>(10)</sup> and a 102-year-old reported in China<sup>(45)</sup>. In our study, 112 patients underwent LC (including 7 patients who underwent LC + EST/EPBD)<sup>[46,47]</sup>, with a success rate of 100%, and none of the patients were converted to laparotomy. The key to success is management of indications and the opportunity to perform LC.

It is important to skillfully separate inflammatory edema and adhesions in the neck of the gallbladder using a suction device, and to ligate or suture the enlarged, thickened and merged sheet-like cystic duct with silk thread during surgery. If necessary, removing the subtotal gallbladder wall and cauterizing the residual gallbladder mucosa can help to avoid bleeding in the gallbladder bed and increased surgery time.

#### Improving perioperative therapy

**Strengthen the function of the heart, lungs, kidneys and other important organs:** In elderly patients with biliary diseases, it is important to frequently monitor ECG, central venous pressure, and urine volume. When necessary, the amount of urine per h should be recorded, and used as a guideline for the infusion volume and speed to prevent heart failure and pulmonary edema caused by excessive and fast infusion. Patients should be encouraged to take a deep breath, expectorate and change position, to maintain an unobstructed respiratory tract and to prevent respiratory tract infection. These measures can help to attenuate postoperative pulmonary atelectasis and pneumonia<sup>[48,49]</sup>.

In patients with coexisting CHD, the serum levels of BNP, cTnI, myoglobin and creatine kinase isoenzyme MB should be monitored to identify heart failure and/ or AMI. In patients with acute heart failure, besides the selective administration of digitalis and other positive inotropic drugs, appropriate use of diuretics can promote the discharge of absorbed interstitial fluid. In addition, the administration of a small dose of dopamine (0.5-2.0 µg/kg per min) can expand the renal and mesenteric vessels, increase renal blood flow and glomerular filtration rate, and then increase urinary output mainly by acting on dopamine receptors; the administration of a medium dose of dopamine (2.0-10.0  $\mu$ g/kg per min) can directly act on  $\beta$ 1 receptors, leading to an increase in myocardial contraction and cardiac output.

In patients with coexisting hypertension<sup>[50]</sup>, besides the use of nitroglycerin and other antihypertensive drugs, increased blood pressure induced by pain<sup>[51]</sup> should be carefully monitored and, when necessary, analgesics should be administered or an analgesic pump used.

In patients with coexisting diabetes, blood glucose

levels may fluctuate shortly after surgery due to the stress response, vasoactive drugs and other factors. Therefore, maintaining postoperative blood glucose at 6.0-10.0 mmol/L may reduce the incidence of postoperative complications<sup>[35]</sup>.

#### Strengthen postoperative nutritional support:

In elderly patients with biliary diseases, multiple coexisting diseases and poor physical condition, supplementation with glucose and vitamins, milk fat, amino acids, albumin and other nutritional support should be given after surgery. When necessary, total parenteral nutrition should be administered. During postoperative fasting, the necessary basic calorie count should be maintained at 25-30 kcal/kg per day. In patients undergoing major surgery, the calorie count should be increased by 30%-40%, and the ratio of calories to nitrogen should be kept at 120-150 kcal:1 g.

### Maintain water, electrolyte and acid-base balance:

In elderly patients with biliary diseases, an electrolyte imbalance can occur due to preoperative and postoperative fasting, the stress response caused by surgery, postoperative gastrointestinal decompression and biliary drainage. Therefore, various electrolytes should be supplied to maintain water, electrolyte and acid-base balance.

In addition, elderly patients often have a long history of aspirin administration. Due to its role in anti-platelet aggregation, the use of aspirin may interfere with normal coagulation function during the perioperative period. However, discontinuation of aspirin before surgery remains controversial<sup>[52,53]</sup>. We recommend discontinuation of aspirin 7-10 d prior to elective surgery in order to avoid the increased risk of surgery-related bleeding, given that the lifespan of platelets is reported to be 7 to 10 d<sup>[54]</sup>. If necessary, such as after coronary artery bypass surgery, low molecular weight heparin can be used as a substitute. In the case of emergency surgery, fibrinogen, prothrombin complex and plasma should be used to antagonize possible intraoperative and postoperative coagulopathy.

#### Timely management of postoperative complications

In elderly patients with biliary diseases, postoperative complications are common due to decreased defense capability, reduced immune function and coexisting diseases.

We identified 10 different postoperative complications in our study cohort, including pulmonary infection, hypopotassemia, incision fat liquefaction and others, with an incidence rate of 39.3% (68/173). Hypopotassemia (33.8%) showed the highest incidence, followed by hypoalbuminemia (22.1%) and pulmonary infection (19.1%). Hypopotassemia should be actively treated, and if necessary, potassium can be pumped into the blood vessels using a micro-infusion pump.

In order to prevent and treat postoperative pulmonary infection, patients should be asked to routinely inhale an aerosol, ensure early ambulation, maintain oral health care, and be tested by regular sputum culture to identify bacterial or fungal infections. If shortness of breath and difficult expectoration occur, the patient should be ventilated as soon as possible and respiratory secretions removed to maintain adequate ventilation. One should never wait until the emergence of respiratory failure before considering the use of a ventilator. In addition, it is necessary to prevent inhalation-induced pulmonary infection due to vomiting and aspiration caused by hypotonia of the lower esophageal sphincter.

Surgical treatment in elderly patients with biliary disease is safe and feasible<sup>[55-59]</sup>. According to the patient's condition, active treatment of coexisting diseases, management of indications and surgical opportunities, appropriate selection of surgical procedures, improvements in perioperative therapy and timely management of postoperative complications are key factors in improving therapeutic efficacy in elderly patients with biliary diseases<sup>[60-62]</sup>.

#### ACKNOWLEDGMENTS

We thank Professor Yu-Min Zhang (Department of Anatomy, Physiology and Genetics, Uniformed Services University of the Health Sciences, United States) for critical reading of the manuscript.

#### COMMENTS

#### Background

With global prolongation of life expectancy, biliary diseases in elderly patients are common and multifaceted.

#### **Research frontiers**

The purpose of this study was to determine the clinical characteristics and different treatment strategies for biliary diseases in elderly patients, in order to improve the diagnosis and treatment of biliary diseases in these patients.

#### Innovations and breakthroughs

Active treatment of coexisting diseases, management of indications and surgical opportunities, appropriate selection of surgical procedures, improved perioperative therapy and timely management of postoperative complications are key factors in enhancing therapeutic efficacy in elderly patients with biliary diseases.

#### Applications

Surgical treatment in elderly patients with biliary diseases is safe and feasible.

#### Terminology

Biliary diseases in the elderly: Biliary diseases in elderly patients aged over 60 years.

#### Peer-review

The article studied an adequate sample size of elderly patients with biliary disease. A positive point of the study is that all patients underwent the same protocol and conducts. Currently, there is a more aggressive trend towards the surgical treatment of these patients based on data from the recent literature,

using better imaging and endoscopic methods for diagnosis and surgical techniques.

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P- Reviewer: Sipahi AM S- Editor: Ma YJ L- Editor: Filipodia E- Editor: Zhang FF







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