# IMPROVEMENT OF THE ALGORITHM FOR DIAGNOSTICS OF POSTCHOLECYSTECTOMY SYNDROME TAKING INTO ACCOUNT FUNCTIONAL AND ORGANIC CHANGES IN THE AREA OF THE LARGE DUODENAL NIPPLE

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#### Abstract

The aim. Improving the algorithm for diagnosing patients with postcholecystectomy syndrome (PCS), taking into account functional and organic changes in the major duodenal papilla (MDP).

**Materials and methods.** 208 patients with PCS were examined. Laparoscopic cholecystectomy (CE) was performed in 172 patients in history, open CE was performed in 36 patients. 47 patients had obstructive jaundice, 84 had signs of biliary hypertension without an increase in bilirubin, 77 had no signs of biliary hypertension and an increase in bilirubin.

The following instrumental research methods were used: abdominal organs ultrasound examination (AO USE), duodenoscopy, endoscopic retrograde cholangiopancreatography (ERCP), computed tomography (CT). In the main group, to study the morphofunctional changes in MDP, duodenoscopy with parietal impedancemetry, ultrasound of the hepatobiliary zone with a choleretic load were used.

**Results.** In the main group, 34 patients underwent ultrasound of the hepatobiliary zone and Vater's nipple area with a choleretic breakfast according to Boyden according to the method of Grigoriev P.Ya. in our modification, 21 patients had an expansion of the choledochus by 2 mm. At the same time, taking meverin led to a decrease in the diameter of the choledochus, which indicated functional changes in the MDP. In 13 patients, the expansion of the choledochus, observed when taking a choleretic breakfast, persisted with the use of antispasmodics, which indicated the presence of an organic pathology of MDP. Duodenoscopy with impedancemetry was performed in 41 patients: 20 patients had 760–820 Ohm, which indicated the presence of functional changes, 12 patients had inflammatory changes in the MDP and 703–760 Ohm were detected, 9 patients with cicatricial fibrous changes had indicators impedance measurement 640–703 Ohm. The use of ultrasound with a choleretic breakfast and duodenoscopy with parietal impedancemetry in patients of the main group made it possible to identify functional and organic (inflammatory, fibrous) changes in the MDP, which made it possible to develop a therapeutic algorithm using both conservative therapy and the use of various options for transpapillary endoscopic interventions.

**Conclusions.** Conducting an in-depth diagnosis of morpho-functional changes in MDP using parietal impedancemetry of MDP and ultrasound of the hepatobiliary zone with choleretic load allows us to identify both functional and morphological changes in MDP, which makes it possible to develop a therapeutic algorithm using drug therapy and transpapillary endoscopic techniques.

Keywords: postcholecystectomy syndrome, impedancemetry, functional and organic changes in the major duodenal papilla.

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#### 1. Introduction

Postcholecystectomy syndrome (PCS) is a symptom complex that contains functional and organic disorders of the major duodenal papilla (MDP) [1, 2]. The most common causes of PCS, both after cholecystectomy (CE) performed by the «open» method, and after laparoscopic CE, are:

stones forgotten or moved during intervention in the choledochus from the gallbladder or cystic duct [3], recurrent choledocholithiasis (mainly after preliminary endoscopic manipulations with stone extraction attempts) [4, 5], MDP restenosis after endoscopic papillosphincterotomy (EPST) [6], blind pouch syndrome, long vesical stump syndrome, biliary fistula, tubular stenosis of the distal choledochus due to chronic indurative pancreatitis, reflux cholangitis due to total EPST and wide choledochoduodenoanastomosis, electrical injury of the biliary tract [7, 8]. An important role may be played by the lack of proper compliance in patients in the postoperative period regarding preventive measures aimed at preventing PCS [9].

Examination of patients who have undergone CE is relevant, both for the purpose of choosing further treatment tactics depending on the identified pathology, and for the purpose of developing measures that are aimed at preventing PCS [10, 11].

**The aim of the research:** improvement of the algorithm for diagnosing patients with PCS, taking into account functional and organic changes in the MDP area.

### 2. Materials and methods

The study was conducted on the basis of the KCC «City Clinical Hospital for Emergency and Emergency Medical Care named after prof. A. I. Meshchaninov» from 2018 to 2021.

208 (100 %) patients with PCS were examined: in the main group -94 (45.2 %) patients, in the comparison group -114 (54.8 %) patients. The comparison group consisted of patients who were hospitalized from 2018 to 2020 in surgical departments and who underwent standard examinations - laboratory, ultrasound examination of the abdominal organs, endoscopic fibrogastro-duodenoscopy, endoscopic retrograde cholangiopancreatography, multidetector computed tomography or magnetic resonance imaging. The main group included patients who were hospitalized in surgical departments from 2020 to 2021, who, in addition to standard examination methods, used additional methods – parietal impedancemetry MDP and ultrasound of the hepatobiliary zone with choleretic load in order to assess morphofunctional changes in MDP.

Laparoscopic CE was performed in 172 (82.7 %) patients, and open CE was performed in 36 (17.3 %) patients. 47 (22.6 %) patients had obstructive jaundice, 84 (40.4 %) had signs of biliary hypertension without an increase in bilirubin, 77 (37 %) had no signs of biliary hypertension and an increase in bilirubin.

An informed agreement was obtained from all study participants. The work was carried out in accordance with the Code of Ethics of the World Medical Association (Declaration of Helsinki). The Committee of the Commission of the Kharkiv Medical Academy of Postgraduate Education on Bioethics approved and approved Protocol No. 3 dated 10/12/2021.

The following instrumental research methods were used: abdominal organs ultrasound examination (AO USE), duodenoscopy, endoscopic retrograde cholangiopancreatography (ERCP), computed tomography (CT). In the main group, to study the morphological and functional changes in MDP, duodenoscopy with parietal impedancemetry, ultrasound with a choleretic breakfast were used.

To perform impedancemetry, a device for ph-impedance monitoring (pH-Z) was used, to which a probe with 6 sensors was connected, of which 4 impedance sensors and 2 ph sensors (**Fig. 1**). After cannulation of the MDP, the impedance spectrum was determined at frequencies from 1 to 10 kHz.



Fig. 1. Ph- and impedance probe sensors for ph-impedance monitoring MDP

Ultrasound of the hepatobiliary zone and the region of Vater's nipple with a choleretic breakfast according to Boyden according to the method of Grigoriev P.Ya. in our modification (author's certificate No. 106257 «Method for diagnosing morphological changes in the sphincter of Oddi») was carried out in patients in order to assess the function of the sphincter of Oddi (SO) and in the absence of data for organic obstruction of the MDP zone at previous stages of examination. First, a thorough study of the hepatobiliary zone, including the MDP area, was performed, measuring the initial width of the CBD. After a food load in the amount of two raw chicken yolks, the measurement of the common bile duct was carried out every 15 minutes for 1 hour. Its expansion by 2 mm or more, in comparison with the initial one, made it possible to suspect incomplete obstruction of the choledochus, the cause of which could be SO dysfunction or organic changes in the MDP area. A decrease in the width of the common bile duct while taking an antispasmodic (meverin) indicated functional disorders of the SO (spasm).

Statistical processing was performed on a personal computer using the standard Microsoft Office Excel 2016 application package and the Statistica 6.0 program. To determine the statistical significance of differences between the study groups, a non-parametric  $\chi^2$  method (Pearson's test) was used.

# 3. Results

In the main group, 34 (36.2 %) patients underwent ultrasound of the hepatobiliary zone and Vater's nipple area with a choleretic breakfast according to Boyden according to the method of Grigoriev P. Ya. in our modification. When performing this study, 21 (22.3 %) patients showed an expansion of the choledochus by 2 mm. At the same time, taking meverin led to a decrease in the diameter of the choledochus, which indicated functional changes in MDP. In 13 (13.8 %) patients, the expansion of the choledochus, observed when taking a choleretic breakfast, persisted with the use of antispasmodics, which indicated the presence of an organic pathology of MDP.

Duodenoscopy with impedancemetry (**Fig. 2**) was performed in 41 (43.6 %) patients: in 20 (21.3 %) patients, indicators of 760–820 Ohm were noted, which indicated the presence of functional changes, in 12 (12.8 %) Inflammatory changes in the MDP were noted in patients and indicators of 703–760 Ohm were detected, in 9 (9.6 %) patients with cicatricial fibrous changes, impedancemetry values of 640–703 Ohm were noted.



Fig. 2. Cannulation of the MDP with a ph-impedance probe during duodenoscopy

When comparing the data obtained by ultrasound with a choleretic breakfast and the data of parietal impedancemetry, a strong direct correlation was revealed (**Table 1**).

That is, there is a significant relationship between the frequency of detection of choledocholithiasis and the type of changes in the MDP that are detected by impedancemetry. The use of ultrasound with a choleretic breakfast and duodenoscopy with parietal impedancemetry in patients of the main group made it possible to identify functional and organic (inflammatory, fibrous) changes in the MDP, which made it possible to develop a therapeutic algorithm using both conservative therapy and the use of various options for transpapillary endoscopic interventions.

## Table 1

Morphofunctional changes in the MDP area

The nature of the changes in the MDP	With choledocholiasis		Without choledocholiasis		Total
	Ν	%	Ν	%	Total
MDP functional changes	14	47 %	16	53 %	30
Inflammatory changes (stenosing) of MDP	21	88 %	3	13 %	24
Scar-fibrous changes of MDP	20	77 %	6	23 %	26

*Note: differences between groups are statistically significant*  $X_{calc}^2 = 11.5$ ,  $X_{crit}^2 = 9.2$ , p < 0.01

In 16 (17 %) patients with functional changes in MDP and absence of choledocholithiasis, conservative therapy was applied with the use of duspatalin, ursodeoxycholic acid preparations.

In 14 (14.9 %) patients with functional changes with choledocholithiasis (**Fig. 3, 4**), the following transpapillary endoscopic interventions were used: in 6 (6.4 %) patients with choledocholithiasis (calculi size 3-7 mm), partial endoscopic papillosphincterotomy, choledocholithoextraction (**Fig. 5**), all 8 (8.5 %) patients with biliary sludge and microcholedocholithiasis underwent balloon dilatation and sanitation of hepaticocholedochus with good immediate and long-term results. In this regard, we note that the use of balloon dilatation according to indications in the treatment of microcholedocholithiasis and biliary sludge in patients with functional changes in MDP is the most optimal surgical intervention.



Fig. 3. Ultrasonography. Choledocholithiasis



Fig. 4. ERCP. Choledocholithiasis



Fig. 5. Endoscopic choledocholithoextraction

In 24 (25.5 %) patients, inflammatory changes in the MDP (stenosing processes) were detected. In 3 (3.2 %) patients with MDP inflammatory processes and the absence of choledocholithiasis, conservative therapy was applied using duspatalin, ursodeoxycholic acid preparations, in 21 (22.3 %) patients with MDP inflammatory stenosing processes and choledocholithiasis, transpapillary endoscopic interventions were used: in 12 (12.8 %) patients with choledocholithiasis (calculi size 3–7 mm) underwent partial papillosphykterotomy, choledocholithoextraction, 9 (9.6 %) patients with large calculi (8–15 mm) underwent subtotal papillosphincterotomy.

In 26 (27.7 %) patients, cicatricial-fibrotic changes in MDP were detected, while 20 (21.3 %) patients had choledocholithiasis. This group underwent transpapillary endoscopic interventions. 13 (13.8 %) patients underwent partial EPST, 8 (8.5 %) patients underwent subtotal EPST, and 5 (5.3 %) patients underwent total EPST.

# 4. Discussion

PCS is a condition that quite often complicates (25-40 %) the postoperative course of cholelithiasis [12, 13]. The presence of symptoms of the disease indicates a deterioration in the quality of life of patients, however, with the diagnostic confirmation of PCS, it is not always possible to identify morphological or functional changes [14]. The use of ultrasound of the hepatobiliary region and the zone of Vater's nipple with a choleretic breakfast according to Boyden according to the method of P.Ya. Grigoriev in our modification made it possible in the study group in 22.3 % of patients to reveal the expansion of the choledochus by 2 mm, the width of which decreased after taking meverin, which was regarded as functional changes in MDP. In 13.8 %, the expansion of the choledochus, after taking breakfast with a choleretic load, persisted against the background of the use of meverin, which indicated an organic pathology in the MDP area. The method of parietal impedancemetry MDP, during duodenoscopy, in the study group in 21.3 % of patients showed resistance indicators of 760-820 Ohm, which indicated functional changes in this area [15]. In 12.8 % of patients, resistance values of 703-760 Ohm were detected in the presence of visual signs of MDP inflammation (edema, hyperemia), and in 9.6 % of patients with visual cicatricial fibrous changes, resistance indicators were 640-703 Ohm. The methods used can be used for the purpose of differential diagnosis between functional and organic pathology of MDP [16, 17].

**Study limitations.** Ultrasound examination of the hepatobiliary region with choleretic load is a non-invasive method for the differential diagnosis of organic and functional pathology in the MDP region, however, it has its limitations, namely, proven choledocholithiasis, previously performed by EPST, which may affect the function of the sphincter of Oddi [18, 19]. MDP

impedancemetry is performed during duodenoscopy, and cannulation of the MDP may be difficult in the presence of MDP diverticula, wedged calculi in Vater's nipple, etc. [20].

**Prospects for further research.** Despite the possible limitations of these research methods, they remain promising for the differential diagnosis of morphological and functional changes in the MDP area, since the method of impedancemetry can be used routinely during duodenoscopy, and the method of ultrasound with a choleretic load is non-invasive for diagnosing disorders of the sphincter of Oddi in patients with PCS.

## 5. Conclusions

Conducting an in-depth diagnosis of morphofunctional changes in MDP makes it possible to identify both functional and morphological changes in MDP, which makes it possible to develop a therapeutic algorithm using drug therapy and transpapillary endoscopic techniques, including balloon dilatation, endosanation of the biliary tract, endoscopic papillosphincterotomy, endoscopic stenting of the common bile duct and others.

When detecting inflammatory and cicatricial changes in the MDP, the probability of detecting choledocholithiasis is 88 % and 77 %, which requires careful identification of stones using other research methods.

Performing parietal impedancemetry and ultrasound with a choleretic breakfast in patients with PCS makes it possible to differentiate the diagnosis of morphofunctional changes in MDP and choose the optimal treatment method.

### **Conflict of interests**

The authors declare there is no conflict of interests.

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