#### References

- 1. Алкогольная болезнь органов пищеварения: клинические очерки. Под ред. Н. Б. Губергриц, Н. В. Харченко. Киев: Новый друк, 2009, 180 с.
- 2. Лапач С. Н. Чубенко А. В., Бабич П. Н. Основные принципы применения статистических методов в клинических испытаниях. Киев: Морион, 2002, 160 с.
- 3. Маев И. В., Зайцева Е. В., Дичева Д. Т., Андреев Д.Н. Ферментные препараты как основа лечения хронического панкреатита с внешнесекреторной недостаточностью: возможности применения и выбор в практике гастроэнтеролога. В: Consilium Medicum. Гастроэнтерология, 2013, № 1, с. 61–64.
- 4. Akshintala V. S. et al. *A population-based study of severity in patients with acute on chronic pancreatitis*. In: Pancreas, 2013, vol. 42, no. 8, p. 1245–1250.
- 5. Clinical pancreatology for practicing gastroenterologists and surgeons (ed. J. E. Dominguez-Munoz). Oxford: A Blackwell Publ. Co., 2005, 535 p.
- Domínguez-Muñoz J. E. Latest advances in chronic pancreatitis. In: Gastroenterol. Hepatol., 2013, vol. 36, suppl. 2, p. 86–89.
- 7. Feher J., Vali L., Blazovics A., Lengyel G. The beneficial effect of metadoxine (pyridoxine-pyrrolidone-carboxylate) in the treatment of fatty liver diseases. In: CEMED, 2009, vol. 3, no. 1, p. 65–79.
- 8. Gullo L., Migliory M., Brunetti M. A., Manca M. *Alcoholic pancreatitis: new insights into an old disease.* In: Curr. Gastroenterol. Rep., 2005, vol. 7, no. 2, p. 96–100.
- 9. Hernandez C. A., Nicolas J. C., Fernandez J., Pizarro P. Determination of plasma trypsin-like activity in healthy subjects, patients with mild to moderate alcoholic chronic pancreatitis, and patients with nonjaundice pancreatic cancer. In: Dig. Dis. Sci., 2005, vol. 50, no. 11, p. 2165–2169.

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# THE EFFECT OF THE PRESENCE OF DIABETES MELLITUS ON CLINICAL COURSE OF CHRONIC PANCREATITIS BY M-ANNHEIM SCORING SYSTEM

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#### Rezumat

# Efectul diabetului zaharat asupra evoluției clinice a pancreatitei cronice conform clasificării M-ANNHEIM

Pancreatita cronică este o boală dificil de diagnosticat și de tratat. Scopul studiului a fost de a evalua severitatea pancreatitei cronice prin clasificarea M-ANNHEIM conform prezenței insuficienței pancreatice endocrine. Au fost analizate prezența insuficienței pancreatice excretorii și incretorii, schimbările structurale ale pancreasului prin ultrasonografie, coprogramă, nivelul de hemoglobină glicozilată în sânge, numărul complicațiilor la 38 de pacienți cu PC(grupul 1) și pancreatită cronică asociată cu diabet (grupul 2). Corelația strânsă dintre gradul de severitate al PC în clasificarea M-ANNHEIM și criteriile obiective de leziuni cronice ale pancreasului demonstrează informativitatea sa înaltă, care este noutatea studiului nostru.

**Cuvinte-cheie**: pancreatită cronică, diabet zaharat, clasificare M-ANNHEIM, grad al pancreatitei cronice

#### Резюме

### Влияние сахарного диабета на клиническое течение и тяжесть хронического панкреатита в соответствии со шкалой M-ANNHEIM

Хронический панкреатит (ХП) относится к заболеваниям, которые сложно диагностировать и лечить. Целью исследования было оценить тяжесть хронического панкреатита в соответствии с классификацией M-ANNHEIM в зависимости от эндокринной недостаточности поджелудочной железы (ПЖ). Были проанализированы: структурные изменения ПЖ по ультразвуковым критериям, копрограмма, уровень гликозилированного гемоглобины, частота осложнений у 38 больных ХП без и с сопутствующим сахарным диабетом. Была выявлена тесная корреляция между тяжестью XII по классификации M-ANNHEIM и объективными критериями хронического поражения поджелудочной железы. Наличие эндокринной недостаточности ПЖ осложняло тяжесть заболевания в соответствии со шкалой M-ANNHEIM, коррелируя c нарушениями копрограммы (r=0.702; p<0.001), ультразвуковыми критерияи (r=0.55; p<0.05), уровнем гликозилированного гемоглобины (r=0.678; p<0.01).

**Ключевые слова**: хронический панкреатит, сахарный диабет, шкала M-ANNHEIM

#### Introduction

Chronic pancreatitis (CP) is the disease difficult to diagnose and treat. This is due to the low sensitivity of the tests and functional imaging techniques pancreas. The problem of diagnosing chronic pancreatitis cannot be considered solved. CP remains a difficult problem of clinical medicine and surgery, as in many

cases accompanied by severe complications and fatal. This is obviously due to the fact that CP is polyetiological disease because in its development play role several reasons [1]. Among them, there are the most important external toxins, metabolic changes, mediated immune factors, congenital and acquired pancreatic duct structure and others. [2]. Secretory and incretory sections of pancreas damaged at the CP, which leads to the concomitant diabetes mellitus (DM) on the later stages, course of which in this disease is studied insufficiently. It occurs in 10-90% of patients with CP [5, 8]. Such a large difference of the literature data about the frequency of diabetes in CP is associated with a different probability of endocrine disruption in various forms of pancreatitis [4, 7].

For today the evaluation criteria of the CP and CP with concomitant diabetes severity, depending on the severity of violations of excretory and incretory function of pancreas remain unclear until the end. In the world practice, to explore the severity of CP the scoring system M-ANNHEIM has been successfully used, which takes into account the multiplicity of the risk factors for CP [3]. Multifactor classification M-ANNHEIM is simple, objective, accurate, does not demand many invasive diagnostic methods that take into account the etiology, disease stage and severity of the clinical course, which opens new perspectives of a wider application in practice of the family doctor. The causal nature of the CP manifestations based on the clinical stage, disturbance of endo- and exocrine functions of the pancreas on the background DM, frequent and severity of complications are shown in the works [9, 10]. The evaluation of the CP severity by the M-ANNHEIN system [11, 12] considered only the severity and frequency of the pain syndrome manifestations and do not take into account the presence of concomitant DM with the development incretory and excretory insufficiency of the pancreas.

The purpose of the research was to assess the severity of chronic pancreatitis by the classification of M-ANNHEIM according to availability of endocrine pancreatic insufficiency.

#### **Patients and methods**

Were examined 38 patients (19 men and 19 women) with a diagnosis of CP, from them – 19 with CP and 19 – with CP and DM. The average age was  $49.8 \pm 2.2$  years. The average age of patients with CP was  $45.3 \pm 3.4$  years (8 men and 11 women), and patients with CP and DM –  $54.4 \pm 2.5$  years (11 men and 8 women). The diagnosis of CP and DM was verified in accordance with the generally accepted criteria in the clinic [6].

CP severity was assessed by the system M-ANN-HEIM based on the clinical stage and severity index. It was analyzed the presence of excretory and incretory pancreatic insufficiency, structural changes of the pancreas by ultrasound criteria, coprogram, the level of glycosylated hemoglobin in the blood, the number of complications. Evaluation of the coprogram conducted by increasing the number of muscle fibers, fiber that is digested, fatty acids, neutral fat, white blood cells, the appearance of mucus helminthic eggs. Each pathological feature was estimated as 1 point. Statistical significance of differences was assessed by averages Student t-criteria (p < 0.05).

#### **Results**

Patients were divided into 2 groups: patients with CP and CP with concomitant diabetes. By classifying M-ANNHEIM all patients belonged to the diagnostic category "defined" CP. 16 of the 38 study patients (42.11%) had II B, 3 patients (7.89%) – II C, 15 patients (39.47%) - III A and 4 patients (10.53%) – III B clinical stage.

Learning pain syndrome in the study patients showed that in 42.10% of patients with CP pain stopped when taking analgesics (2 points). In 52.64% pain had a recurring character, which corresponded to 3 points. There were times when pain was absent, regardless of the presence or absence of drug treatment in 5.26% of patients with CP – pain consistent with both 2 and 3 balls. At 21.05% of patients with CP with concomitant DM the pain abated after administration of analgesics (2 points) and in 78.95% of patients the pain had a recurring character, which corresponded to 3 points. Control of pain was assessed according to the classification of M-ANNHEIM in 1 point in all patients (100%), since all patients received narcotic analgesics.

Surgical treatment performed in 10.53% of patients with CP, and in patients with CP with concomitant DM – 26.31%, and two of them were carried out repeated operations.

In the most patients with CP with concomitant DM – 57.89% – proven exocrine insufficiency was observed, which corresponded to 2 points, in 42.11% – the presence of moderate exocrine insufficiency which did not need replacement enzyme therapy (1 point). In patients with CP 84.21% had proven – (2 points) and 15.79% – mild (1 point) – exocrine insufficiency. Patients without failure of pancreas functions in the research were not present.

Endocrine insufficiency was assessed in the absence or presence of DM and was found in 100% of CP with concomitant DM.

According to the ultrasound criteria, in 73.68% with CP revealed changes in the structure of the pancreas, which corresponded to mild severity (2 points for M-ANNHEIM), in 21.06% of patients – mo-

derate severity (3 points). Significant changes in the structure of the pancreas by ultrasound criteria were noted in 5.26% of patients, which corresponded to severe severity (4 points). 57.89% of patients with CP with concomitant DM with changes of the structure of pancreas had mild severity (2 points), 42.11% of patients had moderate severity (3 points). In patients with CP and CP with diabetes complications were found, respectively, in 3 against 4 patients.

It was found 14 (73.68%) patients with CP with moderate (B) severity and 5 (26.32%) – with medium (C) severity by the M-ANNHEIM classification. However, among the patients with CP and DM were found 2 (10.52%) patients with moderate (B) severity, 16 (68.42%) – with medium (C) severity, 2 (10.53%) – with severe (D) severity and 2 (10.53%) – with heavy (E) severity.

Analyzing the coprogram data, ultrasound data, glycosylated hemoglobin levels and points by M-ANNHEIM classification were found following changes that are shown in table. In patients with CP with concomitant DM coprogram changes were significantly more important than in patients without DM  $(5.47 \pm 0.21)$  against  $(4.73 \pm 0.14)$  points. A similar trend was observed in relation to changes of the ultrasound criteria in points (5.26  $\pm$  0.28) to (4.05  $\pm$ 0.30) points. The level of glycosylated hemoglobin in patients with DM (7.16  $\pm$  0.43) % was significantly higher than in the patients with CP without incretory failure (5.15  $\pm$  0.20) %. The points by M-ANNHEIM classification in the 2<sup>nd</sup> group were significantly more important than in the first group (13.89±0.90) against (8.63±0.52) points.

In conducting the correlation and regression analysis were found direct correlations between the severity of chronic pancreatitis by M-ANNHEIM and changes of coprogram (r=0.702; p<0.001), ultrasound criteria (r=0.55; p<0.05), the level of glycosylated hemoglobin (r=0.678; p<0.01).

Dynamics of coprogram changes, ultrasound criteria and the level of glycosylated hemoglobin in patients with chronic pancreatitis and chronic pancreatitis with concomitant diabetes mellitus

Indication	Groups of patients with CP	
	CP	CP+DM
	n=19	n=23
Coprogram, points	4.73±0.14	5.47±0.21*
Ultrasound criteria, points	4.05±0.30	5.26±0.28*
Glycosylated hemoglobin, %	5.15±0.20	7.16±0.43*
Points by M-ANNHEIM	8.63±0.52	13.89±0.90*

Note: \* - p<0.05

Quantitative criteria for the M-ANNHEIM classification are easy to use, can be easily incorporated into practice of general practitioner that will provi-

de dynamic observation of the patient, timeliness of the application is not only therapeutic but also prophylactic programs in the conditions of clinical examination and routine supervision proved in terms of their use in clinical practice.

#### **Conclusions**

- 1. The close correlation between CP severity by M-ANNHEIM classification and objective criteria of chronic lesions of the pancreas demonstrate its high information content, which is the novelty of our study. The presence of endocrine insufficiency of pancreas in the way of diabetes mellitus complicated a clinical course of chronic pancreatitis according to the M-ANNHEIM scoring system, that correlated with the changes of coprogram (r=0.702; p<0.001), ultrasound criteria (r=0.55; p<0.05), the level of glycosylated hemoglobin (r=0.678; p<0.01).
- 2. For patients with chronic pancreatitis with concomitant diabetes mellitus, a CP degree was more expressed than in the case of chronic pancreatitis without endocrine insufficiency. Average severity (S) prevailed in 68.42% of patients in the 2<sup>nd</sup> group vs. 26.32% of the first group, cases of expressed and severe severity appeared.
- 3. We consider recommending the use of M-ANNHEIM scoring system in gastroenterological practice, therapeutic institutions and practices of family physicians.

#### References

- 1. Babinets L. S. *Chronic pancreatitis*. In: Biblioteka simeinoho likaria ta medsestry, 2011, vol. 2, № 29, p. 25-35.
- 2. Vynokurova L.V. Clinical and pathogenetic mechanisms of developing external and endocrine insufficiency caused by chronic pancreatitis. Avtoref. Dis. d-ra nauk: 14.00.47. Moskva, 2009, 24 p.
- 3. Huberhrits N. B. *New international classification of chronic pancreatitis*. In: Vesnik Cluba Pankreatolohov, 2008, № 2, p. 10-25.
- N. B. Huberhrits, H. M. Lukashevich, O. A. Holubova. Pancreatogenic diabetes mellitus. In: RZHHHK, 2007, № 6, p. 11-16.
- A. P. Rebrov, M. A. Kunitsyna, E.I. Kashkina, E. E. Arkhanhelskaia. *Pancreatogenic diabetes mellitus:* actual problems of pathogenesis and treatment (overview). In: Saratovskiy nauchno-meditsynskiy zhurnal, 2012, № 3, p. 862-867.
- Modern classification and standards of treatment of common illnesses of internal organs (za red. Yi. M. Mostovoho). 11-te vyd. dop. i pererob. Vinnytsia, 2011, 511 p.
- 7. J. J Meier, B. A. Menge, T. G. Breuer. *Functional assessment of pancreatic b-cell area in humans*. In: Diabetes, 2009, vol. 58, № 7, p. 1595–1603.
- 8. H. Schrader, B. A. Menge, C. Zeidler, P. R. Ritter and oth. *Determinants of glucose control in patients with chronic pancreatitis*. In: Diabetologia, 2010, vol. 53, № 6, p. 1062-1069.

- 9. L.V.Vynokurova, I. ETrubitsyna, A.V. Hubina, B. Z. Chikunova. Functional pancreatic insufficiency and its correction depending on the stage of chronic pancreatitis. In: Russkiy meditsinsky zhurnal, 2010, № 13, p. 837-840.
- 10. Huberhrits N. B. *Chronic alcoholic pancreatitis. Uchenie zametki*. In: Materialy 2-ogo mezhdistsiplinarnogo rossiiskogo konhressa. "Chelovek, alcohol, kurenie i pishchevye addiktsii", 24-25 aprelia, 2008, p. 16-26.
- 11. Vinokurova L. V., I. E. Trubitsina. *The clinical course and therapy of chronic pancreatitis depending on the stage of the disease*. In: Ledashchii vrach, 2010, № 2, p. 45-49.
- 12. Schneider A., J. M. Lohr, M. V. Singer. *The M-ANNHEIN* classification of chronic pancreatitis: Introduction of a unifying classification system based on revien of previous classification of the disease. In: J. Gastroenterol., 2007, vol. 42, № 2, p. 101–119.

# TRATAMENTUL CHIRURGICAL AL LEZIUNILOR ŞI STRICTURILOR CĂILOR BILIARE MAGISTRALE

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#### Summary

# Surgical treatment of lesions and strictures of the main biliary ducts

During 1980-2013 year, were hospitalized 233 pacients biliary lesions and benign biliary strictures in the Clinic 1 2nd Department of Surgery of SMPU "N. Testemiţanu". Clinical evaluation included several consecutive steps: 1) setting the ethiopatogenic diagnosis; 2) pre-operatory decompression of the biliary tree; 3) reconstructive surgical act. In case of biliary strictures, following the cut of jaundice and biliary infection, bilio-digestive derivations have been performed according to the level of the obstacle, preferring the bilio-jejunal on isolated loop in Y a la Roux. The post-operatory lethality was of 6 (2.63%) cases.

The lesions of the biliary ducts have a complicated evolution, with many surgical interventions and hospital confinements. effort is needed in order to trace them timely and to prevent septic complications. In the first stage decompression of the biliary tree is performed and following the cut of the inflammatory process, bilio-digestive reconstruction is done.

Keywords: surgical treatment, strictures, biliary ducts

### Резюме

## Хирургическое лечение повреждений и рубцовых стриктур желчных протоков

За период 1980-2013 г., на кафедре Хирургии № 2, клинической базе № 1, ГМФУ им. «Н. Тестемицану» было госпитализировано 233 больных с ятрогенными повреждениями и рубцовыми стриктурами желчных протоков. Клиническое обследование пациентов включало несколько этапов: 1) этиопатогенетическая диагностика; 2) предоперационная декомпрессия желчевыводящих путей; 3) реконструктивная хирургия. В случаях желчных стриктур, после купирования желтухи и желчных инфекций, проводили реконструктивные операции, напрямую зависимые от уровня обструкции, предпочтительно гепатикоеюнальные анастомозы на изолированной петле Roux. Послеоперационная летальность была отмечена в 6 (2,63%) случаях. Ятрогенные повреждения имеют сложную эволюцию, с большим числом хирургическим операций. Необходимо приложить усилия для своевременного диагностики и профилактики септических осложнений. На первом этапе предложено применить декомпрессию желчевыводящих путей, а после – купирование воспалительного процесса, проводят реконструктивные операции.

**Ключевые слова**: хирургическое лечение, стриктуры, желчные протоки

#### **Introducere**

Odată cu prima colecistectomie efectuată în 1882 de către Langenbuch, s-au deschis noi posibilități de terapie a leziunilor iatrogene de cale biliară magistrală (LICBM). În 1899 s-a înregistrat prima reparație reușită a unei astfel de leziuni. De atunci, această tematică a fost dezbătută pe larg în articole și tratate de specialitate, atât în literatura internațională, cât și în cea autohtonă.

Principiile tehnicilor chirurgicale pe parcursul mai multor decenii au fost aduse la o cizelare impresionantă. S-a standardizat și tehnica realizării colecistectomiilor, însă ponderea leziunilor căilor biliare rămâne totuși constantă pe parcursul ultimelor decenii – 0,1-0,8% cazuri.

Colecistectomia clasică a fost un standard de aur în tratamentul colecistitei calculoase mai mult de 100 de ani. Promovarea pe scară largă a colecistectomiei laparoscopice a reaprins controversele din chirurgia clasică. Odată cu introducerea primei colecisectomii laparoscopice de către Erich Mühe von Böblingen (Germania) în 1985 și apoi de P. Mouret (Franţa) în 1987 în practica medicală, s-a constatat o ascensiune uimitoare a acestei tehnologii, care devine doar în câţiva ani un nou standard de tratament, acceptat unanim. Astfel, la moment în ţările dezvoltate se apreciază un nivel de 95% de colecistectomii laparoscopice din totalul efectuat [1, 3, 4, 5].

Literatura de specialitate prezintă o abundență de date statistice și de interpretări și concluzii privitoare la incidența, factorii de risc și mecanismele de producere a LICBM, oferind totodată și recomandări pentru a preveni o astfel de eventualitate.

Până la era laparoscopică, frecvenţa leziunilor cailor biliare, ca urmare a unei colecistectomii tradiţionale, pe parcursul ultimelor decenii a fost un