

Functional disorders in the sphincter of Oddi and possibly reflux associated diseases in the hepato-biliary-cholecysto-pancreatico-duodeno-gastro-esophageal region

Functional disorders in the sphincter of Oddi	Possibly reflux associated diseases
Type of reflux or dysfunction	Target organ – Gallbladder
Pancreaticobiliary reflux (pancreatic juice)	Chronic (enzymatic) cholecystitis. Chronic calculous cholecystitis. Intestinal metaplasia. Dysplasia. Gallbladder cancer.
Biliary type III of sphincter of Oddi dysfunction (spasm of sphincter of common bile duct)	Chronic (spastic aseptic) cholecystitis. Cholesterol gallstone disease. Chronic calculous cholecystitis. Metaplasia.
Duodenal-biliary reflux (duodenal juice) (± <i>Salmonella enterica</i> serovar Typhi)	Chronic (infectious) cholecystitis. Mixed or Pigment (brown) gallstone disease. Chronic calculous cholecystitis. Metaplasia.
Duodenal-biliary (acidic) reflux (duodenal juice and gastric juice) (± <i>Helicobacter pylori</i>)	Chronic (infectious) cholecystitis. Mixed or Pigment (brown) gallstone disease. Chronic calculous cholecystitis. Gastric metaplasia.
Type of reflux or dysfunction	Target organ – Pancreas
Biliopancreatic reflux (lithogenic bile)	Chronic biliary pancreatitis. Biliary metaplasia. Dysplasia. Pancreatic cancer.
Pancreatic type III of sphincter of Oddi dysfunction (spasm of sphincter of pancreatic duct)	Chronic (spastic aseptic) pancreatitis.
Duodenal-pancreatic alcohol reflux (duodenal juice and gastric juice and alcohol)	Chronic (alcoholic infectious) pancreatitis.
Duodenal-pancreatic reflux (duodenal juice) (± <i>Salmonella enterica</i> serovar Typhi)	Chronic (infectious) pancreatitis.
Duodenal-pancreatic (acidic) reflux (duodenal juice and gastric juice) (± <i>Helicobacter pylori</i>)	Chronic (acidic) pancreatitis. Metaplasia. Dysplasia. Pancreatic cancer.
Type of reflux or dysfunction	Target organ – Duodenum – Stomach – Esophagus
Duodenogastric reflux (duodenal juice)	Bile reflux gastritis. Atrophic antral gastritis. Intestinal metaplasia.
Duodenogastroesophageal reflux (duodenal juice and gastric juice)	Bile reflux gastritis. Gastroesophageal reflux disease. Chronic esophagitis. Gastric metaplasia. Dysplasia. Esophageal cancer.
Small intestinal bacterial overgrowth syndrome (duodenum) (duodenal hypertension)	Gallstone disease. Chronic calculous cholecystitis. Chronic pancreatitis.

Absorption function of a gallbladder, a functional status of the sphincter of Oddi, an anatomic configuration of hepatopancreatic ampulla of the sphincter of Oddi (Y-type, V-type or U-type) define development and prevalence of the certain type of pathology in each concrete patient with biliary diseases and pancreatic diseases.

Inactivation of chronic aseptic inflammation – Selective or nonselective COX-2 inhibitors.
 Inactivation of spasm – Selective or nonselective spasmolytics.
 Inactivation of *Helicobacter pylori* – Antibacterial drugs (Eradication).
 Inactivation of *Salmonella enterica* serovar Typhi – Antibacterial drugs (Eradication).
 Inactivation of lithogenic bile and toxic secondary hydrophobic bile acids – Ursodeoxycholic acid.
 Inactivation of pancreatic juice – Pancreatic enzymes (?) and/or Ursodeoxycholic acid (?).
 Inactivation of gastric juice (HCl) – Proton pump inhibitor (PPI) agents. Selective prokinetics.

1. **Selective COX-2 inhibitors** (celecoxib, nimesulide, etc.): **celecoxib** – 100 mg or 200 mg * 2 times per day during **5-7 days**;
2. **Nonselective COX-2 inhibitors** (ibuprofen, diclofenac sodium, indomethacin, naproxen sodium, ketoprofen, flurbiprofen, etc.): **ibuprofen** – 200 mg or 300 mg or 400 mg * 3 times per day during **5-7 days**;
3. **Selective spasmolytics** (pinaverium bromide, mebeverine hydrochloride, himecromone, hyoscine butylbromide, etc.): **himecromone** – 200 mg or 400 mg or 600 mg * 3 times per day during **5-7 days**;
4. **Nonselective spasmolytics** (drotaverine hydrochloride, papaverine hydrochloride, fempiverinium, etc.): **drotaverine hydrochloride** – 40 mg or 60 mg or 80 mg * 3 times per day during **5-7 days**;
5. **Antibacterial drugs** (ciprofloxacin, clarithromycin, amoxicillin, metronidazole, erythromycin, doxycycline, co-trimoxazole, etc.): **ciprofloxacin** – 500 mg * 2 times per day during **5 days**;
6. **Ursodeoxycholic acid**: **ursodeoxycholic acid** – 750 mg * 1 time before going to bed – **14-30-45 days**.
7. **Pancreatic enzymes** (mezym forte, panzynom forte, pancreoflat, festal, kreon, etc.): **kreon 10000** – 1 capsule or 2 capsules * 2-4 times per day during meal during **7-14-30 days**;
8. **Proton pump inhibitor (PPI) agents** (esomeprazole, lansoprazole, omeprazole, pantoprazole, rabeprazole, dexlansoprazole): **rabeprazole** – 20 mg * once daily – **during 4-8 weeks**.
9. **Selective prokinetics** (domperidone, cizapride, metoclopramide, etc.): **domperidone** – 10 mg * 3-4 times per day before meal and at bedtime during **14 days**.

The universal algorithm of the pathogenetic treatment of symptomatic (with biliary pain) biliary diseases with concomitant functional disorders in the sphincter of Oddi:

- 1) **selective COX-2 inhibitors** (celecoxib or nimesulide, etc.):
celecoxib – 100 mg or 200 mg * 2 times per day during 5-7 days,
+
- 2) **selective spasmolytics** (himecromone or mebeverine hydrochloride or hyoscine butylbromide or pinaverium bromide, etc.):
himecromone – 200 mg or 400 mg or 600 mg * 3 times per day during 5-7 days;
+
- 3) **antibacterial drugs** (ciprofloxacin [for eradication of *Salmonella enterica ser. Typhi*] or clarithromycin + amoxicillin or metronidazole [for eradication of *Helicobacter pylori*], etc.):
ciprofloxacin – 500 mg * 2 times per day during 5 days,
+
- 4) **after 5 days of treatment** (1+2+3):
ursodeoxycholic acid – 750 mg 1 time before going to bed – 30-45 days.

Абсорбционная функция желчного пузыря, функциональное состояние сфинктера Одди и анатомические особенности гепатопанкреатической ампулы (**тип Y** или **тип V** или **тип U**) сфинктера Одди определяют развитие и преобладание определенного типа патологии у каждого конкретного больного с заболеваниями желчевыводящих путей и/или поджелудочной железы.

Этот универсальный алгоритм позволяет впервые подобрать и предложить индивидуальный подход патогенетического лечения для каждого конкретного больного с заболеваниями желчевыводящих путей.

Патогенетическая коррекция метаболических и морфо-функциональных нарушений в желчном пузыре, печени и в желчевыводящих путях, в поджелудочной железе и в двенадцатиперстной кишке:

- у больных дисфункцией желчного пузыря позволяет снизить риск возникновения **хронического некалькулезного холецистита без билиарного сладжа**;
- у больных хроническим некалькулезным холециститом без билиарного сладжа позволит снизить риск возникновения **хронического некалькулезного холецистита с билиарным сладжем**;
- у больных хроническим некалькулезным холециститом с билиарным сладжем позволяет снизить риск образования **желчных камней в желчном пузыре** и возникновения **хронического калькулезного холецистита**;
- у больных хроническим калькулезным холециститом позволяет снизить риск возникновения **острого калькулезного холецистита**;
- у больных с постхолецистэктомическим синдромом или состоянием после перенесенной холецистэктомии позволяет снизить риск развития **холедохолитиаза**;
- ♦ у больных с синдромом избыточного бактериального роста (дуоденальная гипертензия), гипертонусом сфинктера гепатопанкреатической ампулы и панкреато-билиарным рефлюксом панкреатического сока в желчный пузырь позволяет снизить риск возникновения **хронического некалькулезного (ферментативного) холецистита, хронического калькулезного холецистита и рака желчного пузыря**;
- ♦ у больных с гипертонусом сфинктера общего желчного протока позволяет снизить риск возникновения **III билиарного типа дисфункции сфинктера Одди, возникновения хронического некалькулезного (спастического асептического) холецистита и хронического калькулезного холецистита**;
- ♦ у больных с синдромом избыточного бактериального роста (дуоденальная гипертензия), недостаточностью сфинктера гепатопанкреатической ампулы и дуодено-билиарным рефлюксом дуоденального сока позволяет снизить риск возникновения **хронического холангита, хронического некалькулезного (инфекционного) холецистита и хронического калькулезного холецистита, пиг-**

ментной желчнокаменной болезни;

- у больных с билиопанкреатическим рефлюксом литогенной желчи позволяет снизить риск возникновения хронического билиарного (желчного) панкреатита и рака поджелудочной железы;
- у больных с гипертонусом сфинктера панкреатического протока позволяет снизить риск возникновения III панкреатического типа дисфункции сфинктера Одди и возникновения хронического (спастического асептического) панкреатита;
- у больных с синдромом избыточного бактериального роста (дуоденальная гипертензия), недостаточностью сфинктера гепатопанкреатической ампулы и дуодено-панкреатическим рефлюксом дуоденального сока и/или дуоденального содержимого позволяет снизить риск возникновения хронического алкогольного («инфекционного») панкреатита, хронического («инфекционного») панкреатита и хронического («кислотного») панкреатита;
- ◆ у больных с дуоденогастральным рефлюксом дуоденального сока (смесь дуоденальной желчи и панкреатического сока) позволяет снизить риск возникновения желчного гастрита или атрофического гастрита антрального отдела желудка;
- ◆ у больных с дуоденогастроэзофагеальным рефлюксом дуоденального и желудочного сока (смесь дуоденальной желчи, панкреатического сока и желудочного сока) позволяет снизить риск возникновения желчного гастрита и желчного эзофагита.

Впервые предложенный данный патогенетически обоснованный метод лечения заболеваний желчевыводящих путей и поджелудочной железы позволяет:

- эффективно купировать болевой и диспепсический синдром в течение 1-3 дней;
- блокировать интенсивность хронического асептического воспаления в стенке желчного пузыря в течение 7-10 дней;
- способствовать полной дезагрегации и элиминации билиарного сладжа в течение 10-14 дней;
- восстанавливать накопительно-выделительную функцию печени в течение 10-14 дней;
- восстанавливать абсорбционную, концентрационную и эвакуаторную функции желчного пузыря в течение 10-14 дней;
- способствовать увеличению продолжительности ремиссии до 2-4 лет.

Таким образом, использование впервые предложенного универсального алгоритма лечения позволяет остановить рост заболеваемости желчевыводящих путей и поджелудочной железы, и уменьшить количество больных дисфункцией желчного пузыря, хроническим некалькулезным холециститом без билиарного сладжа, хроническим некалькулезным холециститом с билиарным сладжем, хроническим калькулезным холециститом и острым калькулезным холециститом, хроническим некалькулезным (ферментативным) холециститом, III билиарным типом дисфункции сфинктера Одди – хроническим некалькулезным (спастическим асептическим) холециститом, хроническим некалькулезным (инфекционным) холециститом, хроническим билиарным (желчным) панкреатитом, III панкреатическим типом дисфункции сфинктера Одди – хроническим (спастическим асептическим) панкреатитом, хроническим алкогольным («инфекционным») панкреатитом, хроническим («инфекционным») панкреатитом, хроническим («кислотным») панкреатитом, хроническим дуоденостазом (синдромом избыточного бактериального роста) и атрофическим гастритом антрального отдела желудка на 30-40%, и улучшить качество жизни больных после перенесенной холецистэктомии и уменьшить количество больных холедохолитиазом.

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Pancreaticobiliary reflux – Biliopancreatic reflux – Choledocho-pancreatic reflux – Duodenal-biliary reflux – Duodenal-pancreatic reflux

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