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ЎТКИР ГЕМОМРАГИК ГАСТРОДУОДЕНАЛ ЯРАЛАРНИНГ ЭТИОПАТОГЕНЕЗИ

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ЭТИОПАТОГЕНЕЗ ОСТРЫХ ГЕМОМРАГИЧЕСКИХ ГАСТРОДУОДЕНАЛЬНЫХ ЯЗВ

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Резюме. Ушбу мақола замонавий тиббиёт ва жарроҳликнинг асосий муаммоларидан бири бўлган гастродуоденал қон кетишларнинг ривожланиши омилларига бағишланган. Мақолада меъда ва ўн икки бармоқ ичак яра касаллигидан қон кетиш сабаблари, уларнинг ривожланиши, консерватив даволашдаги этиопатогенетик ёндашувлар ҳақида умумий маълумот берилган.

Калит сўзлар: меъда ва ўн икки бармоқ ичак яраси, *Helicobacter pylori*, қон кетиш, НЯҚП.

Abstract. The article is devoted to the factors contributing to the development of gastroduodenal bleeding, which is one of the main problems of modern medicine and surgery. The article provides general information about the causes of bleeding from gastric and duodenal ulcers, the features of their development, etiopathogenetic approaches to conservative treatment.

Key words: gastric and duodenal ulcer, *Helicobacter pylori*, bleeding, NAID.

Bleeding from gastric and duodenal ulcers is one of the most dangerous complications, occurring in 15-20% of all complications. According to the latest data [5, 18], the incidence of gastroduodenal bleeding is constantly observed throughout the world, occurring in 40 to 130 cases per 100 thousand population and has a stable trend. The main (35-58%) cause of acute gastroduodenal bleeding is gastric and duodenal ulcer. The majority of such patients are aged 22-45 years, and various conditions and pathologies observed in them (stress, excessive fatigue, consumption of various carbonated and high-calorie drinks, medication, etc.) can lead to the development of gastric and duodenal ulcers and the occurrence of bleeding complications. The second category of patients by age includes the elderly. They are mainly observed in cases of taking anticoagulants, antiplatelet agents and their analogues for preventive purposes, mainly due to the occurrence of cardiovascular pathologies. As is known, anticoagulant and antiplatelet therapy play a key role in the prevention of cardiovascular diseases (arterial hypertension, atherosclerosis, ischemic heart disease).

At the same time, according to many sources [1, 9, 23], instead of taking anticoagulant and antiplatelet therapy regularly, without following the exact dosages and indications, there are often cases of taking them haphazardly, the main complication of which is the

frequent development of hemorrhagic complications in patients prone to bleeding.

Currently, the main antiplatelet drugs include cyclooxygenase (COX) inhibitors (aspirin, i.e. acetylsalicylic acid), adenosine diphosphatase (ADP) receptor blockers (ticlopidine, clopidogrel, glycoprotein IIIa/IIb receptor blockers - tirofiban, abciximab, etc.). The "oldest" and most widely used drug among them is aspirin, which belongs to the salicylic acid derivatives. It blocks platelet cyclooxygenase and, as a result, blocks the synthesis of the monoinducer of platelet aggregation.

According to studies conducted by the Antithrombotic Research Collaboration [12, 17], patients who received acetylsalicylic acid for 12 years and were followed for 23% less than patients with other cardiovascular diseases (acute coronary syndrome, myocardial infarction, acute cerebrovascular accidents) died.

Despite the above-mentioned effectiveness of aspirin, it also has other weak points. For example, it inhibits the activation of platelets by inhibiting cyclooxygenase and the formation of thromboxane A₂. However, this mechanism is not enough to solve the problems of all patients with cardiovascular diseases. One of the most dangerous aspects of aspirin is its ability to irritate the mucous membrane of the stomach and other organs of the digestive tract. We know that the synthesis of

prostaglandin E in the gastric mucosa leads to a strengthening of its protective shell. Acetylsalicylic acid reduces the activity of cyclooxygenase enzymes, leading to erosion of the gastric mucosa and an increased risk of bleeding complications [4, 15]. Another representative of antiplatelet agents are ADP receptor antagonists (clopidogrel, ticagrelor, ticlopidine). They are antagonists of ADP receptor activators that promote platelet aggregation. The most widespread representative of them is Clopidogrel. Like other representatives of this group, it is a prodrug, and metabolites with antiplatelet activity are formed in the liver. According to the recommendations of many world scientists [8, 21], if there are contraindications to taking aspirin or its analogues due to various pathologies of the gastrointestinal tract, the most preferred drug for such patients is Clopidogrel. However, according to other data [10, 14], clopidogrel itself is not recommended as an antiplatelet agent and is considered appropriate to use as a complex therapy in combination with other anticoagulants. Ticagrelor is another modern type of ADP receptor antagonists, which increases the local endogenous adenosine concentration in the body and has functions such as vasodilation, cardioprotection, inhibition of platelet aggregation and modulation of inflammatory processes. [20].

In studies conducted by PEGASUS [6, 19], the combined combination of ticagrelor 60 mg and aspirin, when administered 2 times a day, plays a significant role in preventing antithrombotic complications in various patients, regardless of age, body weight, location and various comorbidities. However, it cannot be used together with drugs such as erythromycin, clarithromycin, fluconazole, cyclosporine. Therefore, it causes problems in the use of anti-*Helicobacter* therapy for gastric and duodenal ulcers. Therefore, it causes problems in the use of anti-*Helicobacter* therapy for gastric and duodenal ulcers. According to Jensen B.E. [2, 25], this limitation significantly narrows the therapeutic window for patients who require both eradication therapy for *Helicobacter pylori* infection and effective antiplatelet management. As a result, in patients with high cardiovascular risk and a history of gastrointestinal complications, a delicate balance must be maintained between the antithrombotic benefit and the gastrointestinal safety profile.

In this context, proton pump inhibitors (PPIs) such as omeprazole and pantoprazole are often co-prescribed with antiplatelet agents to minimize the risk of gastrointestinal bleeding. However, several studies [11, 16, 22] have pointed out that certain PPIs, particularly omeprazole, may interfere with the metabolic activation of clopidogrel through the cytochrome P450 system, especially CYP2C19, thereby reducing its antiplatelet efficacy. This has led to ongoing discussions regarding the choice of gastroprotective agents in such clinical scenarios, with pantoprazole emerging as a more suitable alternative due to its lesser impact on CYP-mediated metabolism.

Moreover, the problem of gastrointestinal bleeding is further exacerbated in elderly patients, as age-related mucosal atrophy and the frequent use of polypharmacy in this population increase their susceptibility to mucosal injury and hemorrhagic complications [13, 24]. Similarly, in pediatric cases, although ulcerative lesions are rarer, when present they tend to be more aggressive, with a

higher likelihood of rapid decompensation due to smaller blood volumes and less physiological reserve [3, 7].

Given these considerations, risk stratification remains essential. Tools such as the Rockall and Glasgow-Blatchford scoring systems are widely used to predict outcomes and guide the management of upper gastrointestinal bleeding [22, 23]. However, these tools primarily assess bleeding severity rather than etiology, and thus may need adaptation or supplementation when applied in the context of chronic antiplatelet therapy.

Conclusion. The management of gastroduodenal ulcer bleeding in patients on antiplatelet therapy, especially in vulnerable populations such as children and the elderly, remains a clinical challenge. While aspirin and newer ADP receptor antagonists like clopidogrel and ticagrelor offer significant cardioprotective benefits, they also carry considerable gastrointestinal risks. Rational drug selection, careful patient monitoring, and the use of gastroprotective co-therapy are key strategies to reduce complications. Future studies are needed to further elucidate safer and more individualized approaches for antiplatelet therapy in patients with existing or potential gastrointestinal pathologies.

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ЭТИОПАТОГЕНЕЗ ОСТРЫХ ГЕМОРРАГИЧЕСКИХ ГАСТРОДУОДЕНАЛЬНЫХ ЯЗВ

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Резюме. Статья посвящена факторам, способствующим развитию гастродуоденального кровотечения, которое является одной из основных проблем современной медицины и хирургии. В статье даны общие сведения о причинах возникновения кровотечений из язв желудка и двенадцатиперстной кишки, особенностях их развития, этиопатогенетических подходах к консервативному лечению.

Ключевые слова: язва желудка и двенадцатиперстной кишки, *Helicobacter pylori*, кровотечение, НПВП.