IMPLICATIONS OF HISTAMINE IN THE MECHANISM OF ACID GASTRIC SECRETION

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Abstract

Histamine is an important factor stimulating the control of acid gastric secretion. Works, probably, regulating the local microcirculation and is involved in the inflammation pathogenesis. It favours biochemical processes of growth and tissue repair. It Stimulates sensitive endings for pain and itching and acting as a modulator of the transmission pulse to some synapses in the brain, interfering in the arousal response in neuroendocrine regulation in thermoregulation and in regulating blood circulation. Histamine stimulates bronchial and intestinal smooth muscle, dilates small vessels and capillaries, stimulates gastric secretion.

Key words: Histamine, Gastrine, Acid secretion

I. HISTAMINE

Histamine or beta imidazoliletilamina is a widespread biogenic amine. It starts in the body by histidine decarboxilation, and it is deposited in the majority in the mastocite and basophile leukocytes. Its Depoziting is in granular formations, together with heparin and chityotactic factors for eosinophils and neutrophils. Mastocite rich tissues - skin, bronchial mucosa, intestinal mucosa - contain relatively large quantities of histamine. Small quantities of histamine are found in epidermic cells and gastric mucosa, in some neurons and in fast rowing tissues.

The main pathways of histamine metabolism in humans consist of N-methylation, followed in part by oxidative desamination (under the monoaminoxidase influence), with formation of N-metiimidazolacetic; oxidative desamination (under the influence of diaminoxidase) with formation of imidazolacetic acid, which is in part riboconjogated. Histamine is an important factor stimulating the control of acid gastric secretion. Works, probably, regulating the local microcirculation and is involved in the inflammation pathogenesis. It favours biochemical processes of growth and tissue repair. It Stimulates sensitive endings for pain and itching and acting as a modulator of the transmission pulse to some synapses in the brain, interfering in the arousal response in neuroendocrine regulation in thermoregulation and in regulating blood circulation. Histamine stimulates bronchial and intestinal smooth muscle, dilates small vessels and capillaries, stimulates gastric secretion.
2. HISTAMINE RECEPTORS.

Histamine effects are due to the operation of specific receptors, belonging to three subtypes:
- H1 - whose stimulation is responsible for bronchoconstriction and vasodilatation.
- H2 - which causes gastric hypersecretion of hydrochloric acid, heart depression and vasodilatation.
- H3 - which acts as presynaptic selfreceptors with inhibitory function at the level of histaminergic endings in the central nervous system.

Location and principal actions of agonists and antagonists at histamine receptors level are presented in Table 1.

Table 1

<table>
<thead>
<tr>
<th>Receiver</th>
<th>Location</th>
<th>Action mech.</th>
<th>Actions</th>
<th>Agonists</th>
<th>Antagonists</th>
</tr>
</thead>
<tbody>
<tr>
<td>H1</td>
<td>smooth muscle, brain, vascular</td>
<td>GQ PLC</td>
<td>Bronchoconstriction and capillary permeability.</td>
<td>2-methyl-histamine</td>
<td>astemizole, clemastina, clorfenamines</td>
</tr>
<tr>
<td></td>
<td>endothelium</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>H2</td>
<td>gastric mucosa, myocardium</td>
<td>Gs Ac</td>
<td>Gastric secretion of hydrochloric acid</td>
<td>4(5)-methyl-histamine</td>
<td>cimetidine, ranitidine</td>
</tr>
<tr>
<td></td>
<td>brain, mastocite</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>H3</td>
<td>brain, plexus mienteric</td>
<td>GQ Ca2</td>
<td>Inhibition of some neurotransmitter release</td>
<td>A-methyl-histamine</td>
<td>Impomidin, tioperamide</td>
</tr>
</tbody>
</table>

3. IMPLICATIONS IN ACID GASTRIC SECRETION.

Stimulating effect of histamine on gastric acid secretion is known since 1920. Physiological role of histamine in gastric secretion was elucidated only after the discovery of histamine H2-blocker. Regarding the gastric acid secretion mechanism confrontations were summarized 2 approaches:
1. Final comon mediator theory (theory of Code.), in which histamine is the
final mediator to other secretagogue (gastrine, acetylcholine), resulting that the parietal cell has receptors only for histamine.

2. Grossman’s theory - Konturek, or receptor theory, which postulates that all three secretagogue (histamine, acetylcholine, gastrine) stimulate parietal cells directly and that each of them potentiates the other. According to this theory, these 3 receptors are in a dynamic interaction so that activation of one receptor increase sensibility to other specific receptors for agonists and inhibition of one receptor decreases sensitivity of other antagonists receptors.

It can be said after all the current data that histamine plays a central role in regulating gastric acid secretion. Recent research shows that histamine is produced and released from neuroendocrine cells (NEC) of the gastric mucosa of the fundus and gastrine and vagus nerve stimulate the formation and release of histamine from NEC.

Regarding the relationship with calcium, it is noted that high concentrations of calcium inhibit histamine release stimulated by gastrine inhibiting also the gastric acid secretion. At the same time, serum calcium levels did not influence gastric acid secretion stimulated by histamine.

H2 receptor transduction is mediated by GTP linked to G protein with 3 subunits: alpha, beta, gamma. These subunits are associated or dissociated related to receptor occupation or vacancy. H2 receptor G protein has not yet been isolated, but its existence is strongly suggested by the observation that GTP could mimic histamine stimulation of gastric acid secretion. This would be of subtype Gs (adenilat cyclase stimulator), in contrast to Gi protein associated with somatostatine PG inhibites adenilat cyclase. Histamine increases intracellular cyclic AMP considered the second messenger of H2 receptor. Histamine increases the circulatory flow of gastric mucosa which is inhibited by cimetidine but not by omeprazole, and also serves as a vasodilator on gastric arterioles via H1-receptor, or via the H1-H2-receptor.

4. CONCLUSION:

1. Histamine or beta imidazoliletilamina is a widespread biogenic amine.
2. Histamine is an important factor stimulating the control of acid gastric secretion.
3. It favours biochemical processes of growth and tissue repair.
4. It Stimulates sensitive endings for pain and itching and acting as a modulator of the transmission pulse to some synapses in the brain, interfering in the arousal response in neuroendocrine regulation in thermoregulation and in regulating blood circulation.
5. Histamine stimulates bronchial and intestinal smooth muscle, dilates small vessels and capillaries, stimulates gastric secretion.

6. Histamine effects are due to the operation of specific receptors, belonging to three subtypes:
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REFERENCES