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## GASTRO PROTECTIVE EFFECTS OF *USNEA LONGISSIMA* METABOLITES ON PROBIOTIC *LACTOBACILLUS CASEI*

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**ABSTRACT:** Protective effect of the probiotic combination of usnic acid and *Lactobacillus casei* in experimentally induced ulcer in rats was investigated. Rats inward usnic acid (100 mg/kg), *L. casei* ( $10^{-8}$  con.), usnic acid (100 mg/kg) + *L. casei* ( $10^{-8}$  con.), and omeprazole (30 mg/kg) twice daily for 5 days for prevention against aspirin (ASP), ethanol (EtOH), cold restraint stress (CRS) and pylorus ligation-induced ulcer (PL). The results of the present study showed the first time that the usnic acid (100 mg/kg) + *L. casei* ( $10^{-8}$  con.) as probiotic combination significantly inhibited the ulcer index in ASP, EtOH, CRS and PL to  $3.4 \pm 0.12$ ,  $6.3 \pm 1.8$ ,  $3.4 \pm 0.8$  and  $4.3 \pm 0.9$  ( $p < 0.001$ ) respectively, as compared to control group ( $19.2 \pm 1.6$ ,  $22.5 \pm 6.3$ ,  $24.2 \pm 3.2$ ,  $14.2 \pm 2.7$ ). Besides usnic acid (100 mg/kg) + *L. casei* ( $10^{-8}$  con.) offered protection (72%,  $p < 0.001$ ) against ethanol-induced depletion of gastric wall mucus. The usnic acid (100 mg/kg) + *L. casei* ( $10^{-8}$  con.) showed significant inhibition of lipid peroxidation and superoxide dismutase ( $5.46 \pm 1.30$ ,  $120.6 \pm 3.2$ ) ( $p < 0.01$ ,  $p < 0.001$ ) respectively and enhance activity of catalase ( $32.2 \pm 1.3$ ,  $p < 0.001$ ) to healthy group range ( $34.2 \pm 2.7$ ,  $p < 0.001$ ). However, it elevated the decreased level of  $\text{PGE}_2$  from  $0.62 \pm 0.12$  ( $p < 0.05$ ) to  $2.03 \pm 0.51$  ( $p < 0.001$ ) as compared to omeprazole. These results suggest that probiotic combination could attenuate the severity of gastric ulcer and prevent the toxicity level of usnic acid in the liver.

**INTRODUCTION:** Abnormal secretion of gastric acid and pepsin are the main reason for gastric ulcer disease. Nowadays, critical issue for gastric ulcer disease is gastric hyper secretion-linked with gastrinoma in Zollinger-Ellison syndrome, raise in parietal-cell mass, antral G-cell hyperplasia, and a physiological disproportion between the antagonistic gastric hormones gastrin and somatostatin.

It is known that cholinergic hypersensitivity and parasympathetic dominance both are related to pepsin<sup>1</sup>. Besides having anti-inflammatory and analgesic activity, NSAID especially aspirin; significantly increase the risk of gastrointestinal infection in Asian countries. Individually those are related to gastric injury: Ulcer complications especially bleeding. NSAIDs cause an injurious effect by the inhibition of COX1 and its function in standard mucosal protection mechanisms, and also by the inhibition of thromboxane  $\text{A}_2$ , which accord platelet function and cause in gastrointestinal bleeding<sup>2-5</sup> but simultaneous low-doses of aspirin reduces the effect<sup>2</sup>. However, the interaction between microbial flora and pathogenic organisms has been reported<sup>6</sup>.

