

Circulating level of interleukin (IL)-18 in *Helicobacter pylori*-infected patients, and its associations with bacterial CagA and VacA virulence factors

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Background: We analyzed the impact of Interleukin (IL)-18 promoter polymorphisms on IL-18 serum levels in *H. pylori*-infected duodenal ulcer (DU) patients and healthy asymptomatic (AS) carriers. We also aimed to determine the association of the *H. pylori* virulence factors, CagA and VacA antibodies with serum concentrations of IL-18 as to elucidate any correlation between them.

Methods: Three groups including DU patients (67 individuals), AS carriers (48 individuals), and *H. pylori*-negative subjects (26 individuals) were enrolled. Serum concentrations of IL-18 were determined by ELISA. Patient sera were tested by Western blot method to determine the presence of serum antibodies to bacterial CagA, VacA. Genotyping of IL-18 promoter polymorphisms at positions -137 G/C and -607 C/A were performed by Allele-Specific Primer PCR protocol.

Results: Our study revealed that serum IL-18 levels are positively influenced by CagA-positive *H. pylori* strains, so that maximum levels of IL-18 were detected in DU patients with the CagA+ phenotype, regardless of the presence of the anti-VacA antibody. Regarding IL-18 promoter polymorphisms, the AA genotype and A allele at position -607 C/A found to be significantly lower in DU patients than in AS carriers and *H. pylori*-negative subjects ($P=0.032$ and 0.043 , respectively).

Conclusions: IL-18 -607 C variant was associated with higher levels of serum IL-18 and increased risk of DU. Moreover, our findings indicated that serum concentrations of IL-18 were influenced by CagA factor, irrespective of the VacA status, suggesting that high levels of IL-18 in CagA positive subjects are predisposed to susceptibility to DU.

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