Lack of Association between Interleukin-12 Gene Polymorphisms and Recurrent Aphthous Stomatitis

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Recurrent Aphthous Stomatitis (RAS) is the most common oral inflammatory disease, which is a painful, ulcerative condition of the oral cavity 1 and is characterized by episodic, small, round ulcers with erythematous halos 2. Although several factors such as systemic diseases, nutritional factors, psychological stress, local trauma, allergies, smoking and hormonal alterations could be associated with RAS, genetic factors seem to have an important role in predisposition to this condition whereas the exact pathogenesis of the disease has not clearly been understood 3.

Interleukin (IL)-12 which is secreted by macrophages and dendritic cells has a key role in differentiation of Th0 cells into Th1 cells 4, and therefore it could theoretically have a role in RAS pathogenesis. Considering the fact that SNP could affect the cytokine secretion, an attempt was made to evaluate the alleles and genotypes frequencies of IL12 gene in a group of patients with RAS.

In this investigation, 5 ml blood from sixty four Iranian patients with confirmed diagnosis of RAS 5 was collected in the EDTA tubes. DNA was extracted using a phenol-chloroform method. This project was approved by Ethics Committee of Tehran University of Medical Sciences. Written informed consent was obtained from all subjects before sampling. IL12 gene typing was performed by Polymerase Chain Reaction with Sequence-Specific Primers (PCR-SSP) assay (PCR-SSP kit, Heidelberg University, Heidelberg, Germany), similar to what explained before 6. The allele and genotype frequencies of IL12 gene at position -1188 correlates with low cytokine secretion 7. In the present study, IL12 SNP was evaluated at that position which is located in the promoter region of the gene. However, no significant difference on IL12 alleles and genotype frequencies between the patients and the controls (Table 1).

In several studies, association of number of cytokine gene polymorphisms in pathogenesis of RAS has been investigated 8,9. In this study, the possible role of IL12 SNP with RAS was investigated and no association was found which is similar to previous studies 9. It has been documented that an A to C exchange in the 3' UTR of IL12 gene at position -1188 correlates with low cytokine secretion 10. In the present study, IL12 SNP was evaluated at that position which is located in the promoter region of the gene. However, no significant difference on IL12 (A -1188 C) alleles and genotypes between the patients and the controls was found.

Lack of association between IL12 (A -1188 C) polymorphisms and RAS could indicate that IL-12 has no significant role in pathophysiology of RAS.

Keywords: Interleukin 12, Recurrent aphthous stomatitis, Single nucleotide polymorphisms
Table 1. Comparison of alleles, genotype frequencies of \textit{IL12} between patients with RAS and the control group

<table>
<thead>
<tr>
<th>Position</th>
<th>Genotypes/Haplotypes</th>
<th>RAS (n=60), n(%)</th>
<th>Controls (n=140), n(%)</th>
<th>p-value</th>
<th>Odds Ratio (95% Confidence Interval)</th>
</tr>
</thead>
<tbody>
<tr>
<td>-1188</td>
<td>A</td>
<td>91(77.1)</td>
<td>204(72.9)</td>
<td>0.446</td>
<td>1.26(0.74-2.15)</td>
</tr>
<tr>
<td></td>
<td>C</td>
<td>27(22.9)</td>
<td>76(27.1)</td>
<td>0.446</td>
<td>0.80(0.47-1.36)</td>
</tr>
<tr>
<td></td>
<td>AA</td>
<td>34(57.6)</td>
<td>72(51.4)</td>
<td>0.519</td>
<td>1.28(0.67-2.48)</td>
</tr>
<tr>
<td></td>
<td>CA</td>
<td>23(39)</td>
<td>60(42.9)</td>
<td>0.727</td>
<td>0.85(0.44-1.66)</td>
</tr>
<tr>
<td></td>
<td>CC</td>
<td>2(3.4)</td>
<td>8(5.7)</td>
<td>0.387</td>
<td>0.58(0.08-3.08)</td>
</tr>
</tbody>
</table>

References


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