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Title: The prognostic value of IFN- γ and IL-4 cytokines ratio and CD25+ lymphocytes blood content for asthma exacerbation

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Body: Background: Asthma is a disease characterized by a chronic airway inflammation, so it is important to find biomarkers to assess the activity of inflammation and to predict asthma exacerbation. Objective: To evaluate IFN- γ and IL-4 cytokines ratio and CD25+ lymphocytes blood content in patients with asthma and to predict asthma exacerbation. Methods: We included 48 patients with asthma exacerbation and 48 with asthma remission and 30 matched control subjects. All underwent detailed clinical examination and spirometry. IFN- γ and IL-4 cytokines concentration in blood was measured by enzyme immunoassay. Investigation of CD25+ lymphocytes in blood was carried out by flow cytometry. Results: Patients with asthma exacerbation had significantly lower values of IFN- γ and IL-4 ratio than patients with asthma remission (0,026 (0,016-0,046) vs 0,092 (0,066-0,157); p<0,001) and controls (1,44 (1,17-1,69); p<0,001), they also had significantly higher values of CD25+ cells than patients with asthma remission (10,22 (4,92-13,09)% vs 5,85 (4,14-7,83)%; p<0,001) and controls (5,49 (3,69-6,01)%; p<0,001). The regression equation for asthma exacerbation prediction is:

$Z = \exp(1,40005 + 0,107702X - 54,223Y) / (1 + \exp(1,40005 + 0,107702X - 54,223Y))$, where X – IFN- γ and IL-4 ratio; Y – % of CD25+ lymphocytes. If $Z \leq 0,5$, the patient has remission; if $Z > 0,5$, the patient has exacerbation; the above Z, the above asthma exacerbation probability. The value of correct predictions is 89%. Conclusion: Decrease of IFN- γ and IL-4 cytokines ratio and increase of CD25+ lymphocytes blood content can be predictor for asthma exacerbation and allows predicting asthma exacerbation with the probability of 89%.