

**PS5:97 ROLE OF CD107A+ (LAMP-1) CYTOTOXIC CD8+ T-CELLS IN PATIENTS WITH SLE**

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**Objectives** Cytotoxic T-cells are a thought to play a pivotal role in the pathogenesis of systemic lupus erythematosus (SLE) in particular in lupus nephritis. The aim of this study was to investigate the activation marker CD314 (NK2GD) and CD107a (LAMP-1) on cytotoxic CD8+ T-cells in patients with systemic lupus erythematosus.

**Methods** Peripheral blood of patients with systemic lupus erythematosus (n=30) and healthy controls (n=21) was stained with anti-CD8 (PB), -CD3 (Chrom Orange), -CD45RO (FITC), -CD197 (PE), -CD314 (APC), -CD107a (APC) antibodies and analysed by flow cytometry. Kidney biopsies of lupus nephritis patients were investigated by immunohistoc-chemistry and immunofluorescence for the presence of CD8+CD107a+ cells. Patients disease activity was assessed by systemic lupus erythematosus disease activity index (SLEDAI).

**Results** The percentages of CD314+CD8+ T-cells were not different between SLE-patients and healthy controls. The percentages of CD107a+CD8+ T-cells were significantly decreased in SLE-patients as compared to healthy controls (40.2%±18.5% vs 47.9±15.0%, p=0.02). This was even more significant in SLE-patients with inactive disease. The evaluation of lupus nephritis biopsies showed a significant number of CD107a+CD8+ T-cells mainly located in the peritubular infiltrates.

**Conclusions** These results demonstrate that CD8+ T-cells of patients with systemic lupus erythematosus have an altered activation status which seems to be associated with disease activity. The proof of intrarenal CD107a+CD8+ suggests a role in the pathogenesis of lupus nephritis.

**PS5:98 TCR ZETA DIM LYMPHOCYTES IN PATIENTS WITH SYSTEMIC LUPUS ERYTHEMATOSUS**


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T lymphocytes with low or absent expression of TCR zeta chain (TCR zeta -/dim cells) are supposed to have pathogenic effects in the development of autoagression and chronic inflammation. Systemic lupus erythematosus (SLE) is a model of chronic inflammatory disease with autoimmune background.

**In vitro** studies have shown, that microenvironement rich in proinflammatory cytokines, such as TNF-alfa and IL-6, especially promotes the impairment of TCR zeta chain expression, what leads to the increase of the percentage of T cells with defects of the synthesis of that molecule.

**Objective** The aim of this study was to define the size of the population of autoreactive TCR zeta -/dim lymphocytes in the blood of patients with SLE and the functional characteristics of TCR zeta -/dim and TCR zeta bright cells based on the assessment of their capacity to synthesize certain cytokines (IFN-gamma and IL-2) and to compare it with healthy controls.

**Results** Our study showed decreased percentage of CD4+ and CD8+ lymphocytes with TCR zeta chain expression in patients with SLE. The significantly diminished TCR zeta chain expression in lymphocytes of patients with SLE which in addition was correlated with longer disease duration, renal involvement and higher level of serum anti-dsDNA antibodies confirms former studies indicating not only the association of this defect with disease development, but also with more active clinical course.

The in vitro stimulation of T lymphocytes in SLE patients lead to overproduction of IFN-gamma especially in the population of autoreactive TCR zeta -/dim lymphocytes.

On the other hand, T lymphocytes of SLE patients have lower capacity of the synthesis of IL-2, which is strongly correlated with TCR zeta chain expression and may be observed even in patients with very low clinical disease activity.

**Conclusion** SLE is characterised by the expansion of TCR zeta -/dim lymphocytes in peripheral blood, which are regarded as autoreactive cells, and is correlated with clinical course of SLE. The decreased expression of TCR zeta may be due to the presence of exogenous inhibiting factor in blood of SLE patients. Higher synthesis of INF-gamma and lower of IL-2 may contribute to chronic inflammation and autoimmune processes in SLE.

**PS5:99 EXAMINING THE MODULATORY EFFECTS OF ANTI-SERINE PROTEASE ANTIBODIES UPON FACTOR XA, THROMBIN AND COMPLEMENT INTERACTIONS**


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**Purpose** Serine protease (SP) enzymes play a critical role in both the coagulation and complement cascades. Anti-SP antibodies are found in approximately 40% of patients systemic lupus erythematosus (SLE) and antiphospholipid Syndrome (APS). Complement activation is important in SLE and APS. However, little is known about the effects of anti-SP antibodies on complement activation in these diseases. We sought to investigate whether affinity-purified antibodies to the SP Factor (F)xa and Thrombin (Thr) alter the effects of these SP on complement cleavage in the presence or absence of the physiological inhibitor anti-thrombin (AT).

**Methods** Serum was obtained by informed consent from patients with SLE (n=10) and/or APS (n=2) under long-term follow-up at University College London Hospital. A novel method was developed to affinity purify anti-Fxa and anti-Thr IgG separately. We incubated Fxa (2 μM) and Thr (2.7 μM) separately with complement component C3 to determine baseline C3 cleavage. We then repeated the experiment in the presence of AT3, the appropriate anti-SP antibody. We operated C3 cleavage by 1.8-fold. AT mediated inhibition was prevented by addition of anti-Fxa...