CIRCULATING CD4+ T CELLS LEVELS IN ACTIVE AND NON ACTIVE SYSTEMIC LUPUS ERYTHEMATOSUS PATIENTS

Ernesto Cairoli,****** Alfonso Cayota,****** María José Iriarte,***

Sebastián Irureta,*** Alex Rocha***

Dear Editor,

We read with great interest the article written by Ferreira et al¹ recently published in your journal. Considering the comments made in discussion by the authors, we highlight some differences obtained in our work. The objectives of the present study were to quantify the levels of circulating CD4+ T cells in systemic lupus erythematosus (SLE) patients and further correlate their levels with the degree of disease activity. A prospective study was performed in the Unit of Systemic Autoimmune Disease, Hospital de Clínicas, School of Medicine, Uruguay. Thirty consecutive (hospitalized and ambulatory) patients with SLE were included. All patients fulfilled four or more of the revised classification criteria for SLE of ACR.2 Disease activity was scored based on the SLE disease activity index (SLE-DAI),³ with one group comprising patients with non active disease (SLEDAI < 5; n = 16) and another group with active disease (SLEDAI ≥ 5 ; n = 14) with or without immunosuppressive treatment. Peripheral blood samples were drawn for simultaneous measurements of total white blood cells and CD4+ T cells (theses by flow cytometry). In all cases informed consent was obtained according to local approved ethical rules. Comparison between the different groups was performed using Mann-Whitney U test and correlations between absolute number of CD4+T cells and SLEDAI scores were assessed by nonparametric Spearman correlation. A *p*< 0.05 was considered statistically significant.

29 out of the 30 patients were female. The mean

Lymphopenia correlates with disease flares that may contribute to the development of susceptibility to infections, ^{4,5} however, CD4+T cells abnormalities were not generalized to all SLE patients. ^{6,7}

In the context of a retrospective study, the results of Ferreira et al,¹ could be influenced by the inclusion of patients with severe immunosupression. In our case, the prospective inclusion of consecutively patients could better reflect the status of CD4+ T cell deficits in SLE. Although the sample size is small, our series included only one male patient, better reflecting the gender distribution observed in the SLE in this age range. We have not found significant differences between CD4+ T cell number and either non active or active SLE patients. This result, could be explained at least in part, by changes induced by high doses of prednisone in patients with active disease. Our results do

age was 38.5 ± 15 years and duration of disease was 8.5 ± 10 years in the total of SLE patients included. 16 patients (53%) had non active disease (SLEDAI 1.0 ± 1) and 14 patients (47%) had active disease (SLEDAI 13 \pm 6). No significant differences of age, disease duration, percentage and total number of lymphocyte among the groups were detected. A decrease in the concentrations of complement C3 and treatment with high doses of prednisone were found in SLE active group (Table I). The absolute number of CD4+ T lymphocyte (cell/µl) and the percentage in the non active and active SLE patients was 508 ± 153 and 471 ± 288 and $39.7 \pm 8.5\%$ and $36.5 \pm 11.0\%$ respectively. The active patients seemed to have lower mean levels of CD4+ T cells than inactive patients, however the difference was not statistically significant. No significant correlation between absolute cell numbers of CD4+T cells and SLEDAI score in the active SLE patients was detected (Spearman r = -0.347). There were no opportunistic infections and only in 3 patients (with active disease) bacterial infection was confirmed.

^{*}Unidad de Enfermedades Autoinmunes Sistémicas, Hospital de Clínicas, Facultad de Medicina, Universidad de la República, Uruguay.

^{**}Clínica Médica «C», Hospital de Clínicas, Facultad de Medicina, Universidad de la República, Uruguay.

^{***}Departamento Básico de Medicina, Hospital de Clínicas. Facultad de Medicina, Universidad de la República, Uruguay. ***Institut Pasteur, Montevideo, Uruguay.

| | Total SLE | Non active SLE | Active SLE | p value |
|-------------------------------|-------------|----------------|-------------|---------|
| Total (n) | 30 | 16 | 14 | _ |
| Age (years) | 38.5 ± 15 | 38.3 ± 12 | 38.7 ± 18 | 0.471 |
| SLE duration (years) | 8.5 ± 10 | 7.8 ± 8 | 9.3 ± 12 | 0.359 |
| SLEDAI | _ | l ± 1.4 | 13 ± 6.0 | 0.001 |
| Complement C3 | _ | 123 ± 26.8 | 65.1 ± 17.7 | 0.001 |
| Prednisone dose (mg/day) | _ | 6.7 ± 11.4 | 44.3 ± 17.4 | 0.001 |
| Total Lymphocytes | 1572 ± 1033 | 1502 ± 791 | 1646 ± 1270 | 0.357 |
| Absolute CD4+ T cell (n°/ µl) | 492 ± 218 | 508 ± 153 | 471 ± 288 | 0.340 |
| % CD4+ T cell | 38.3 ± 9.6 | 39.7 ± 8.5 | 36.5 ± 11.0 | 0.195 |

not support the view that CD4+T cell counts could constitute a marker of disease activity.

Correspondence to

Ernesto Cairoli.

Unidad de Enfermedades Autoinmunes Sistémicas. Clínica Médica «C». Hospital de Clínicas. Avenida Italia s/n. piso 8. telefax: +5982 487 87 02.

E-mail: ecairoli@hc.edu.uy

Acknowledgments

This work was supported in part by "Programa para la Investigación Biomédica" (PROINBIO) and Fundación Manuel Pérez. Facultad de Medicina. Universidad de la República. Uruguay.

References

- Ferrerira S, Vasconcelos J, Marinho A, et al. CD4 lymphocytopenia in systemic lupus erythematosus. Acta Reumatol Port 2009; 34:200 - 206.
- 2. Tan E, Cohen A, Fries J, et al. The 1982 revised criteria for the classification of systemic lupus erythematosus. Arthritis Rheum 1982; 25:1271 1277.
- 3. Bombardier C, Gladman D, Urowitz M, et al. Derivation of the SLEDAI: a disease activity index for lupus patients. the Committee on Prognosis Studies in SLE. Arthritis Rheum 1992; 35:630 640.
- Duffy KN, Duffy CM, Gladman D. Infection and disease activity in systemic lupus erythematosus: a review of hospitalized patients. J Rheumatol 1991; 18:1180 1184.
- Iliopoulos A, Tsokos G, Immunopathogenesis and spectrum of infections in systemic lupus erythematosus. Semin Arthritis Rheum 1996; 25:318 - 336.
- 6. Via C, Tsokos G, Bermas B, et al. T cell-antigenpresenting cell interactions in human systemic lupus erythematosus. J Immunol 1993; 151:3914 3922.
- Wouters C, Diegenant C, Ceuppens J, et al. The circulating lymphocyte profiles in patients with discoid lupus erythematosus and systemic lupus erythematosus suggest a pathogenetic relationship. Br J Dermatol 2004; 150:693 700.