Remission or low disease activity as a target in systemic lupus erythematosus

We read the report by Zen *et al*¹ on the impact of lupus low disease activity state (LLDAS) on damage accrual with great interest and would like to congratulate the authors on their work. In this single-centre Caucasian cohort, followed for 7 years, only 38 of their 293 patients (11.3%) failed to achieve LLDAS for at least 1 year; moreover, of the 255 patients who achieved LLDAS for at least 1 year, 246 (96.5%) also satisfied the definition of remission for the same length of time. And, in 214 (83.9%) patients, the duration of remission was the same as that of LLDAS. Being in LLDAS for at least 2 years was associated with a reduced risk of damage; but, when remission was included in the multivariable model, remission for at least 2 years was associated with a reduced risk of damage whereas LLDAS was not. Based on these findings, the authors suggest that the effects of LLDAS could be overlapping with those of remission. However, the prevalence of systemic lupus erythematosus (SLE) in non-Caucasian populations is higher and the disease is more severe with patients having, in general, less favourable outcomes; this limits the applicability of Zen et al's results to patients with different characteristics. In fact, our previously published data from the multiethnic, multinational inception Latin American cohort² differs quite a bit from Zen *et al*'s dataⁱ. Remission was a relatively rare event with only 273 of 1350 patients (20.2%) achieving it at least once during their follow-up, and another 192 patients (14.2%) achieving low disease activity state (LDAS) while 885 (65.6%) did not achieve either status, with a follow-up of 2.4 years. Since we have a larger number of patients with pure LDAS, it was possible to evaluate the real impact of this status on SLE prognosis. Additionally, since we evaluated these statuses as intervals, we could independently ascertain the impact of remission and LDAS; we found that both statuses were protective of new damage occurrence whereas remission was protective of severe new damage, defined as an increase of at least 3 points in the Systemyc Lupus International Collaborating Clinics/American College of Rheumatology (SLICC/ACR) Damage Index. Moreover, remission and LDAS were protective of both, new and severe new damage excluding those items clearly related to glucocorticoid use suggesting that the effect of LDAS and remission on damage accrual is primarily related to the level of disease activity. Although the definitions of LLDAS and LDAS used by Zen et al and by us are not exactly the same, they nevertheless do not explain by themselves the differences between our publications (both definitions required a Systemic Lupus Erythematosus Disease Activity Index (SLEDAI) \leq 4, and similar treatment; the definition used by Zen *et al* included no new lupus disease activity compared with the previous assessment, Physician Global Assessment ≤ 1 which ours did not require).

Studies from other countries have reported an incidence of remission and LLDAS more comparable to our's than to Zen *et al*'s suggesting than in disadvantaged SLE populations, LLDAS may be an alternative, yet not the ideal, treatment target.³

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ⁱOur data were not cited by Zen *et al* as the dates of both the publications, theirs and ours, were quite close in time.