Glucocorticoid withdrawal in lupus – to do or not to do?

I read with great interest, the recently published article in your journal titled “Withdrawal of low-dose prednisone in SLE patients with a clinically quiescent disease for more than 1 year: a randomised clinical trial” by Mathian *et al.*¹ The discussion on the feasibility of completely stopping glucocorticoids in systemic lupus erythematosus (SLE) has been ongoing and it poses a practical challenge to every physician treating SLE. Besides the adverse effects, studies also suggest that long term glucocorticoid therapy for remission maintenance in SLE leads to increased accrual of organ damage.² The effect of withdrawal of glucocorticoids on SLE remission has been studied in a number of observational studies. Data from a randomised trial was lacking except for a small pilot study (SIMPL) conducted by Galbraith *et al* where the effect of glucocorticoid withdrawal on lupus nephritis remission maintenance was studied in 15 patients.³

The above-mentioned study by Mathian *et al* (CORTICOLUP study) provides scientific evidence addressing this issue. The study has successfully shown the superiority of glucocorticoid maintenance therapy over its withdrawal both in terms of all flares and severe flares.¹ The follow-up period was only 52 weeks, yet the number of flares in the withdrawal group were quite high (27% vs 7%) with a HR of 0.2. The results of the CORTICOLUP study are in contrast to the results of previous studies.^{3,4}

Majority of the patients in the CORTICOLUP study in either groups (28 and 25 in maintenance and withdrawal groups, respectively) were not on any immunosuppressive therapy besides hydroxychloroquine, and 11 patients were not on hydroxychloroquine.¹ It would be interesting to know if there was any difference in the flare rates among those who were on immunosuppression compared with others. Another avenue to explore is the time of steroid withdrawal. A retrospective study by Tani *et al* suggests that the time interval from the last flare to the steroid withdrawal may also play a role in future flares.⁵ Interestingly, the quiescent time was more in the withdrawal group in CORTICOLUP study, although not significant.¹

Research is ongoing to study the steroid sparing effects of biologics in remission induction. Belimumab, epratuzumab, and tabalumab have been successfully shown to reduce the steroid doses by 25% or more in remission induction in a meta-analysis by Oon *et al.*⁶ The use of biologics and other immunosuppressants as steroid sparing therapy for remission maintenance in SLE is

yet to be explored. In the absence of randomised trials favouring steroid withdrawal, the question remains – to withdraw or not to withdraw?

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