Dr. Genovese comments

To the Editor:

The Abatacept Comparison of sub(QU)cutaneous versus intravenous in Inadequate Responders to methotrexatE (ACQUIRE) trial was a multinational, Phase IIIb, randomized, double-blind study that evaluated the comparable efficacy and safety of subcutaneous (SC) and intravenous (IV) abatacept (ABA) over 6 months¹. At Month 6, similar proportions of SC and IV ABA-treated patients achieved an American College of Rheumatology 20 response (estimated difference: 0.3%, 95% CI -4.2, 4.8), confirming noninferiority of SC to IV ABA. The onset and magnitude of efficacy responses were equal for both formulations, and similar patient retention was also reported (94.2% for SC ABA vs 93.8% for IV ABA at Month 6). Overall safety was also similar between groups, including discontinuations due to adverse events and serious adverse events, serious infections, malignancies, and autoimmune events. However, the trial did not look directly at the question of switching from IV to SC in the parent trial. That aspect was considered in the longterm extension (LTE) study², in which all patients who completed the 6-month double-blind period received SC ABA 125 mg weekly for up to ~3.5 years of exposure, to assess its longterm safety, efficacy, and tolerability. Clinical and functional benefits were maintained longterm during the ACQUIRE LTE study, regardless of whether patients received SC ABA throughout or switched from IV ABA to SC ABA at the start of the LTE.

These observations support findings from the ATTUNE study, which evaluated safety and efficacy in patients who switched from IV to SC ABA. The results of 2 large studies, ATTUNE³ and the ACQUIRE LTE², reached different conclusions from those of Reggia, *et al*⁴. However, one needs to remain cautious regarding overinterpretation of clinical trials data, because

ACQUIRE and ATTUNE had fairly homogeneous patient populations followed in the context of fairly rigorous protocols, and trial results may not always represent individual smaller non-trial patient populations.

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