Reviewer's report

Title: Old drugs, old problems: Where do we stand in prediction of rheumatoid arthritis responsiveness to methotrexate and other synthetic DMARDs?

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Reviewer: Raimon Sanmarti

Reviewer's report:

This is a comprehensive and exhaustive revision on the topic of prognostic factors of therapeutic response to MTX and other DMARD in patients with RA. I have only minor concerns

- In the some of the articles analysed it is difficult to ascertain if the prognostic factors are related to MTX itself or may be related to other DMARDS or the combination of both. I think that this statement should be addressed in the introduction section.

- Taking into account that this review is addressed not only to rheumatologists, the impact and the recommendations of use of MTX in RA in the therapeutic strategy of RA may be of interest for the readers; therefore a small summary on this topic is appropriate.

- The section of Genetic biomarkers of respone seems to me to be too extensive and I suggests to shorten it of substantial form.

Quality of written English: Acceptable

Statistical review: No, the manuscript does not need to be seen by a statistician.

Declaration of competing interests:

I declare that I have no competing interests