Reviewer’s report

Title: Effects of low-dose computed tomography on lung cancer screening: a systematic review, meta-analysis, and trial sequential analysis

Version: 1 Date: 13 Jun 2019

Reviewer: José Belda

Reviewer's report:

Congratulations on the manuscript, it is extraordinary. I would like to make some simple questions.

Did you find any difference in terms of death and major complications between diagnostic and therapeutic procedures?

What does mean: "2 studies were excluded because there were no relevant results"?

After excluding the MILD trial, the LCCT screening showed a statistically significant reduction in lung cancer mortality; on the contrary, the same result was not observed when excluding the DANTE trial. For what reason?

What's the reason for such significant difference in the lung cancer mortality between subgroups of higher versus lower quality trials?

Could you speculate about how a high quality RCT for lung cancer screening should be designed to minimize outcomes bias?

Are the methods appropriate and well described?
If not, please specify what is required in your comments to the authors.

Yes

Does the work include the necessary controls?
If not, please specify which controls are required in your comments to the authors.

Yes

Are the conclusions drawn adequately supported by the data shown?
If not, please explain in your comments to the authors.

Yes
Are you able to assess any statistics in the manuscript or would you recommend an additional statistical review?
If an additional statistical review is recommended, please specify what aspects require further assessment in your comments to the editors.

I recommend additional statistical review

**Quality of written English**
Please indicate the quality of language in the manuscript:

Acceptable

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