Reviewer’s report

Title: The NAPRESSIM Trial: The use of low dose prophylactic naloxone infusion to prevent respiratory depression with intrathecal morphine in elective hepatobiliary surgery: study protocol and statistical analysis plan for a randomised controlled trial

Version: 1 Date: 06 Nov 2017

Reviewer: Nuria Porta

Reviewer’s report:

They have addressed all my previous comments and have made changes to improve readability of the manuscript. The only bit outstanding is, in my view, to add further clarification in the manuscript regarding the follow-up period. In their answers to both Reviewer 1 and myself (questions 4 and 15) they have explained all patients will have received treatment by 2pm on the surgery date, and followed up until 8am on the following day.

I think the bit of follow-up starting before 2pm on the day of surgery is missing from the manuscript, and it would explain why follow-up is between 18-24h (this could be explicitly stated. They make a point that this reflect clinical practice, so I would add it explicitly in the protocol.

And then clarify in lines 229: Pts will be monitored up to 24h - and then say at least 18h, so I think it just needs to be simplified and say follow-up for min 18h to up to 24h max of follow-up.

I am happy to recommend its publication.

Level of interest
Please indicate how interesting you found the manuscript:

An article whose findings are important to those with closely related research interests

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

Acceptable

Quality of figures
All images and figures within the manuscript should be genuine i.e. without evidence of manipulation. No specific feature within an image may be enhanced, obscured, moved, removed, or introduced. If you have concerns about the veracity of the figures you should choose the first option below.
Statistical review
Is it essential that this manuscript is seen by an expert statistician? If so, please give your reasons in your report.

Declaration of competing interests
Please complete a declaration of competing interests, considering the following questions:

1. Have you in the past five years received reimbursements, fees, funding, or salary from an organisation that may in any way gain or lose financially from the publication of this manuscript, either now or in the future?

2. Do you hold any stocks or shares in an organisation that may in any way gain or lose financially from the publication of this manuscript, either now or in the future?

3. Do you hold or are you currently applying for any patents relating to the content of the manuscript?

4. Have you received reimbursements, fees, funding, or salary from an organization that holds or has applied for patents relating to the content of the manuscript?

5. Do you have any other financial competing interests?

6. Do you have any non-financial competing interests in relation to this paper?

If you can answer no to all of the above, write 'I declare that I have no competing interests' below. If your reply is yes to any, please give details below.

I declare that I have no competing interests.

I agree to the open peer review policy of the journal. I understand that my name will be included on my report to the authors and, if the manuscript is accepted for publication, my named report including any attachments I upload will be posted on the website along with the authors' responses. I agree for my report to be made available under an Open Access Creative Commons CC-BY license (http://creativecommons.org/licenses/by/4.0/). I understand that any comments which I do not wish to be included in my named report can be included as confidential comments to the editors, which will not be published.

I agree to the open peer review policy of the journal