Reviewer’s report

Title: PrEvention of posttraumatic contractuRes with Ketotifen 2 (PERK 2) – Protocol for a multicenter randomized clinical trial.

Version: 0 Date: 26 Nov 2019

Reviewer: Fabricio Fogagnolo

Reviewer's report:

Dear authors,

Congratulations for such interesting protocol. The subject has paramount importance in the field of articular trauma and the study has an enormous potential of interesting findings.

I have taken some notes about minor corrections (and suggestions):

Abstract / Discussion:

The results of this study will provide evidence for the use of KF in treating post-traumatic joint contractures and improving quality of life after joint injuries. (Maybe preventing or reducing post-traumatic joint contractures would be a more appropriate definition of the objective of intervention - use of medication)

Line 6, page 2:

Background: Injuries and and resulting stiffness… (remove and)

Line 55, page 2:

Discussion: The results of this study will provide evidence for the use of KF in treating post-traumatic joint contractures and improving quality of life after joint injuries. Same as above: maybe preventing or reducing would be a more appropriate definition of the objective of intervention - use of medication

Line 46, page 6

..fracture healing, our interest is the number of participants with disappearance of radiographic fracture lines overtime - Maybe more specific criteria should be used to assess and classify bone healing, such as RUST or mRUST scores (J Bone Joint Surg Am. 2018;100:1871-8)
Line 21, page 7 - same as above for fracture healing assessment

Line 28, page 9 - The TSC will meet quarterly to review trial performance and adjust the RCT (remove will, duplicated)

Line 12, page 11: Contractures complicate other orthopaedic conditions or procedures, such as total knee arthroplasty (please review sentence or it needs a better explanation - knee?) and positive results in post-traumatic (post-traumatic) contractures could apply to these other orthopaedic conditions or procedures.

Line 49, page 11: an externally funded...

**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 am able to assess the statistics

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

Acceptable

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