Author's response to reviews

Title: TP53 mutations in ovarian carcinomas from sporadic cases and carriers of two distinct BRCA1 founder mutations; relation to age at diagnosis and survival

Authors:

Pedro Kringen (pedrok@broadpark.no)
Yun Wang (yun.wang@medisin.uio.no)
Vanessa Dumeaux (vanessa.dumeaux@medisin.uio.no)
Jahn M Nesland (j.m.nesland@medisin.uio.no)
Gunnar Kristensen (gunnar.kristensen@klinmed.uio.no)
Anne-Lise Borresen-Dale (a.l.borresen-dale@medisin.uio.no)
Anne Dorum (anne.dorum@dnr.uio.no)

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Author's response to reviews:

Dear Editor

Please find the revised version of our manuscript entitled TP53 mutations in ovarian carcinomas from sporadic cases and carriers of two distinct BRCA1 founder mutations; relation to age at diagnosis and survival by Pedro Kringen, Yun Wang, Vanessa Dumeaux, Jahn M. Nesland, Gunnar Kristensen, Anne-Lise Borresen-Dale and Anne Dorum.

The referees comments were very helpful and constructive and we have carefully taken their suggestions into account.

The following is a point-by-point listing of our responses to the comments.

Reviewer: David Goldgar

Minor Compulsory Revisions

Regarding figure 1 new and fewer regression lines for all sporadic and all familial cases are drawn
Goodness of fit are added on figure 1.

Text out of line on page 9 is corrected.

Reviewer: Richard Buller

The 8 cases positive on TTGE but not confirmed by sequencing are removed from the study. All statistics now reflect this throughout the manuscript.

Reviewer: Roddam

Major
1) Information regarding type of test used to calculate difference in age at onset are now in the paragraph with the results, p10

2) Log-rank test performed, p11

3) We think that the relationship between age at onset and survival time is not presented clearly in a Kaplan-Meier Plot and that the current figure 1 gives a good visual presentation of the relationship. However, we acknowledge that the cases still alive is not always an exact survival time. We have therefore added a mark for the cases still alive in figure 1.

4) Word changed from "eotime" to "erate".

5) We are not quite sure what part needs to be summarized in a table. The first part is regarding the number of transitions vs. transversions and could possibly be presented in a 2x2 table but still would need some text explaining the results. The part explaining the univariate analyses are compact and adding a table will require more space and even more text without adding substantial information.

6) There are now references to table 3 in the TP53/BRCA1 results, p11

7) Parts of discussion and summary are re-written according to new results.

Minor

1) "eRelative risk" are now termed "ehazard risk".

2) The statistical significance level is corrected to "eset at 0.05".

3) Text error corrected, p9

Statistical review

All statistics rerun. The basis for the statistics is clearly stated in the manuscript.

Yours sincerely

Dr. med PhD Anne Dorum