Author's response to reviews

Title: Burkitt lymphoma beyond MYC translocation: NMYC and DNA methyl transferases deregulation

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Author's response to reviews: see over
Dear Editor,

on behalf of all the authors I would like to thank you and the reviewers for your consideration and criticisms that surely improved the paper.

Please find following a point-by-point letter of response.

Reviewer 1

The authors show results and discuss the difference in miRNA profiling in BL w. and w.o. c-myc translocation, but do not clarify if some of these miRNA could be directly under the transcriptional control of c-myc. Similarly, the authors suggest that N-myc could be directly regulated by the hsa-miR-29a, but do not discuss the possibility that N-myc is affecting the miRNA profile as a transcriptional factor.

The possibility that MYC is regulating itself the expression of some microRNAs and that MYCN is impacting the molecular profile has now been discussed in the Discussion section and some supporting references have been added

English needs some language corrections before being published

English language has been revised.

Reviewer 2

Discretionary and minor revisions were all addressed as you suggested.

In the paragraph on “The microRNA pattern impacts on the gene expression profiling (GEP) of BL cases”…….

A figure displaying the expression of DNMT3b at the mRNA and protein level has now been added to the results

There are no data regarding the EBV status

EBER analysis was performed on all the cases and the results are has been included in the revised form of the manuscript.

In the paragraph on “MYC translocation positive and MYC translocation negative BL cases”…….

We described the characteristic of the previously reported miRNA signature and quoted the corresponding reference as correctly requested. If needed, Nature Publishing Group (Leukemia) might be asked for permission for reproduction and a table listening all the 38 miRNA might be added.
In the same paragraph I suggest to clarify what “DLBCL morphologically mimicking classical BL” exactly means.

In the revised form we have added what does DLBCL morphologically mimicking classical BL namely the BL/DLBCL category according to the WHO. A reference have also been added.

In the Discussion, indirect regulation of DNMT1……..

The relevant references has been cited in the revised form of the manuscript.

At the end of the Discussion, decreased expression of has-miR29 family member…..

The text has been modified accordingly.

Reviewer 3

Please revise in the abstract where you write that MYC (italics) is a transcription factor either to MYC (the protein) or the gene encoding the transcription factor MYC. Please use MYCN (italics) for the gene, now sometimes MYCN and sometimes NMYC (italics).

As you suggested, names of genes and proteins have been changed accordingly.

Best personal regards

Lorenzo Leoncini