Author's response to reviews

Title: Hidden chromosomal abnormalities in Pleuropulmonary Blastomas identified by Multiplex FISH

Authors:

Benoit Quilichini (Benoit.Quilichini@ap-hm.fr)
Nicolas Andre (nicolas.andre@ap-hm.fr)
Corinne Bouvier (corinne.bouvier@mail.ap-hm.fr)
Marie-Anne Chrestian (mchrestian@ap-hm.fr)
Angelique Rome (angelique.rome@ap-hm.fr)
Dominique Intagliata (helene.cannoni@ap-hm.fr)
Carole Coze (carole.coze@ap-hm.fr)
Gabriel Lena (gabriel.lena@ap-hm.fr)
Helene Zattara (helene.cannoni@ap-hm.fr)

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Author's response to reviews: see over
CORRECTIONS IN THE MANUSCRIPT

1-Corrections suggested by Reviewer 2, concerning nomenclature karyotypes, have been applied.

Case 1:

38~44,X,-X,der(8)t(8;18)(p1?;q1?),dic(11;21)(p11;p11),-15,der(17)t(15;17)(q1?;p1?3),
-18,del(20)(p12pter),21[cp11]/81~82<4n>,idemx2[cp2]/46,XX[7].ish der(8)t(8;18)
(D8Z1+,wcp8+,wcp18+),dic(11;21)(wcp11+,wcp21+),der(17)t(15;17)(D17Z1+,
wcp15+,wcp17+,TP53-)

Case 2:

80~109<4n>,XXYY,+X,der(1)t(1;8)(p1?;q1?)x2,dup(3)(?q)x2,+der(?4)t(1;4),-5,dup(7)
(q?),+8,-9,der(9)t(9;13),-10,-10,-11,-16,-17,der(19)t(10;19)x2[cp8]/46,XY[7].ish
der(1)t(1;8)(wcp1+,wcp8+),dup(3)(?q)(wcp3+),der(?4)t(1;4)(wcp1+,wcp4+),dup(7)(q?)
(wcp7+),t(9;13)(wcp9+,wcp13+),t(10;19)(wcp10+,wcp19+)
2-In order to shorten the manuscript, modifications have been made:

Page 4 – line 9: suppression of She was then referred to our hospital.

Page 9 – lines 2-4: suppression of Treatment is based surgical resection of the tumor combined with chemotherapy but clear evidences and recommendations concerning optimum therapy are still lacking

Page 9 – line 7: suppression of To the best of our knowledge

Page 9 – lines 14-17: suppression of In order to identify recurrent genetic abnormalities of importance in the pathogenesis of PPB, and to obtain a refined karyotypic description, we used multicolor FISH, Locus-Specific and Enumeration FISH probes.

Page 9 – lines 22-23: suppression of If this numerical abnormality seems to represent the most common feature in PPB,

Page 10 -lines 18-19: suppression of In our first case, TP53 deletion was demonstrated on metaphase FISH

Page 11 – lines 2-3: suppression of The fact that only few cases have been analyzed, the understanding and the prognosis value of this genetic event remains unclear in PPB.

Page 11 – lines 16-20: suppression of Molecular cytogenetic analysis by Multiplex FISH technique of both our cases reveals complex structural chromosomal abnormalities involving chromosomes 1, 3, 4, 7, 8, 9, 10, 11, 13, 15, 17, 18, 19 and 21. The karyotype complexity seems to be a feature of PPB and a genetic profile of the pleuropneumoblastoma entity is difficult to establish due to the lack of accurate genetic data published.

Page 12 – line 3: suppression of From this point of view, interestingly,
CORRECTIONS IN FIGURE LEGENDS:

*In order to shorten the figure legends, modifications have been made:*

Figure 3: suppression of : The refined tumor karyotype is:

42, X, -X, der(8), der(11), -15, der(17), -18, del(20)(p12pter), -21 .ish
der(8)t(8;18)(D8Z1+,wcp8+,wcp18+) .ish der(11)dic(11;21)(wcp11+,wcp21+) .ish
der(17)t(15;17)(D17Z1+,wcp15+,wcp17+,TP53-)

Figure 4: suppression of: The refined tumor karyotype is: 81, XXXYY, +der(1)x2, +2, +2,
+der(3)x2, +4, +4, +der(?4), +5, +6, +7, +der(7), +8, +8, +der(9), +11, +12, +12,
+14, +16, +18, +18, +der(19)x2, +20, +20, +21, +22, +22 .ish der(1)t(1;8)(wcp1+,wcp8+)
.ish der(3) dup(3)(?q)(wcp3+) .ish der(?t(1;4)(wcp1+,wcp4+) .ish
der(7)dup(7)(q?)(wcp7+) .ish der(9)t(9;13)(wcp9+,wcp13+) .ish
der(19)t(10;19)(wcp10+,wcp19+)

CORRECTIONS IN TABLE I:

The karyotypes have been corrected:
1- suppression of spaces (see Table I- version 3) have been made
2- “idem” for case n°4 was added – for the other cases, respecting the data published we can’t modify the karyotypes.