Author's response to reviews

Title: Novel PTRF-CAVIN mutation in a child with mild myopathy and very mild congenital lipodystrophy

Authors:

Anna Ardissone (anna.ardissone@istituto-besta.it)
Cinzia Bragato (cinzia.bragato@istituto-besta.it)
Lorella Caffi (npibergamo@ospedaliruniti.bergamo.it)
Flavia Blasevich (flavia.blasevich@istituto-besta.it)
Lucia Morandi (lmorandi@istituto-besta.it)
Isabella Moroni (isabella.moroni@istituto-besta.it)
Marina Mora (mmora@istituto-besta.it)

Version: 5 Date: 23 July 2013

Author's response to reviews: see over
To: BMC Medical Genetics

RE: Manuscript MS: 105092838846767 - Novel PTRF-CAVIN mutation in a child with mild myopathy and very mild congenital lipodystrophy.

Dear Editor,

We are now submitting a new version, revised in response to the reviewers’ comments.

Our replies to the reviewers are detailed below.

Please note that the changes have required further studies with inclusion of 2 new co-authors.

We hope the paper may now be acceptable for publication in BMC Medical Genetics.

Sincerely yours,

Marina Mora, PhD
Dept. of Neuromuscular Diseases
Istituto Nazionale Neurologico “C. Besta”
Via Temolo 4
20126 Milano
Italy
Tel: +39 02 23942632
fax: +39 02 23942619
Email: mmora@istituto-besta.it

First reviewer

1. I recommend to cite a recent article by Murakami et al. (PMID: 23489663) and discuss overlapping clinical findings and differences. We have now cited this article and discuss differences in Discussion page 6, 2nd paragraph.

2. The official name of the gene is PTRF We have now changed PTRF-CAVIN into PTRF throughout the manuscript.
3. Second paragraph p. 4: the first sentence should be reworded. One suggestion: The patient was born, after an uncomplicated pregnancy, with APGAR scores of 8 at 1 minute and 9 at 5 minutes. He was noticed to have supinated left foot varus, mild dysmorphic features and (reduced motility?) (muscle weakness?hypotonia?)?
We have now changed the sentence as follows: “The patient was born after an uncomplicated pregnancy with APGAR 8 at 1 minute and 9 at 5 minutes. He was noticed to have supinated left foot varus, mild dysmorphic features and reduced motility without hypotonia.”

4. Last line p.6: “correct stop codon” should be changed to “original stop codon”.
We have changed “correct” into “original”

Second reviewer
1. Case Report Para 6: Authors claim that the novel mutation introduces a STOP codon 27 bases downstream of the correct STOP codon. This needs to be substantiated by the DNA sequence. Evidence from Fig 1 indicates a protein larger than the expected size.
Needs clarification.
We have changed supplementary figure 1 with one that shows the DNA sequence.

Minor Essential Revisions:
1. Discussion Para 3: Delete “therefore unsurprising”. Insert “not surprising”.
We have changed “it is therefore unsurprising” into “it is not surprising”.

Third reviewer
1. One of the major findings of CGL is the decrease of serum leptin and adiponectin levels. These levels should be mentioned in this article
We now provide leptin and adiponectin levels in Case Report, page 5 and mention it in the Discussion page 6, 2nd paragraph.

2. In the case of heterozygous mutation of one nucleotide deletion, double waves should be appeared downstream from the deleted nucleotide. However, sequence results of parents in figure 3D, double waves are appeared upstream from the deleted nucleotide. The reason should be mentioned.
We thank this reviewer for noticing the confusing sequence that had been obtained using the reverse primer. We have now changed the sequence with the one obtained using the forward primer, which indeed shows double waves downstream of the deleted nucleotide as expected.