**Author’s response to reviews**

**Title:** GERMLINE c.1A>C HETEROZYGOUS PATHOGENIC VARIANT IN SDHA REPORTED FOR THE FIRST TIME IN A YOUNG ADULT WITH A GASTRIC GASTROINTESTINAL STROMAL TUMOUR (GIST): A CASE REPORT

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Reviewer reports:

I am satisfied with the response to the reviewers comments. I have been through the manuscript and recommend additional changes as set out below.

When using italicized letters for the name of genes, remove the word "genes" after the italicized word - Do this throughout the manuscript.

IT HAS BEEN MADE.

Background

Paragraph 1. Change "diagnosed of GIST..." to diagnosed with a GIST...."

Paragraph 2. Change "..related condition, being the neurofibromatosis type...." to "related condition, with neurofibromatosis type..."
Case Presentation

Paragraph 1
A 20 year old female was referred to our Medical Oncology department after a gastric antrum GIST resection. A subtotal gastrectomy revealed a pathologic specimen compatible with an epithelioid GIST variant localised to the muscularis propria

Change the sentence where it is written "....CD117 (KIT) and DOG1, which supported GIST diagnosis." to "....CD117 (KIT) and DOG1, which supported a GIST diagnosis."

Change the sentence starting with "This corresponded to pT1pN0 (0/1) high mitotic rate - stage II - of the eighth edition of TNM classification." to "The tumor was classified as a pT1pN0 (0/1) with a high mitotic rate - stage II - according to the eighth edition of TNM classification."

Change ".... calculate the risk of this tumor relapse and/or progression as zero..." to ".... calculate the risk of this tumor relapsing and/or progressing as zero..."

Change the sentence "These procedures revealed us the presence...." to "These procedures revealed the presence..."

Paragraph 2
...molecular features (WT) of GIST tumor..." to "...molecular features of her tumor...."

Change "The family history of the patient seemed to be, at first sight, irrelevant: she had a 19 years old healthy brother. Her parents, 51 years old, had no history of any diseases." to "The family history of the patient seemed to be unremarkable: she has a 19 year old healthy brother. Her parents, 51 years of age, had no history of any diseases."

Change "Maternal family history was anodyne but among paternal family history the fact that the grandfather died at the age of 52 as a consequence of a pituitary adenoma was worth mentioning." to "The maternal family history was anodyne but the paternal family history included a grandfather who had died at the age of 52 as a consequence of a pituitary adenoma."
Change "....when he was 30 years old and he received...") to "....when he was 30 years of age and he received...")

Change "... maternal and paternal lines." to "...maternal or paternal lines."

Paragraph 3. Change "...presentation and in a spite of an..." to "presentation and in spite of an..." and further in this sentence, remove the word "sample"

Results

Paragraph 1. Change "... the presence , in heterozygous state, of the c.1A>C ;...." to ...the presence of a heterozygous c.1A>C;..."

Change "....responsible for starting translation from the messenger RNA..." to "...responsible for translation initiation from messenger RNA..."

Paragraph 2. Change "Her brother was found to be no carrier of the variant;" to "Her brother was found not to be a carrier of this variant;"

Change "grandmother was no carrier of the variant." to "grandmother was also negative for this variant."

Discussion and Conclusions

Paragraph 1. Change the first sentence to "An exhaustive review of databases and the medical literature was undertaken in order to assess the pathogenicity of this variant. And the second sentence to "The c.1A>C; p.(Met1?) variant has previously been reported in the a patient diagnosed with Leigh syndrome, in which genetic analysis revealed the presence of a compound heterozygote..." In the second last sentence (line 22) remove the words "performed" and "high".

Paragraph 2. Last sentence, line 31, change the word "Other" to "Another" and on line 32 change "published" with "reported".

Paragraph 3. At the end of line 39 add the word "the" after "consequence", line 40 change the word "to" to "with". Line 43 insert "the" between "on" and "basis" and the word "this" after "basis of" and change "patient" to "patient's". Line 51, remove "in the performed" and put in "by". At the end of this line remove "must". Line 54, change "mutation" to "pathogenic variants". Line55, delete "On the other hand" and change deficient to deficiency". Line 56 remove "are" and change "overlapped" to "overlap". Line 57 remove "in a", change "extensive" to "extensively" and delete "way"/
Paragraph 4. Line 9, remove "patient" and change "diagnosed" to "diagnosis" and remove after the word "higher" the word "is". Line 14, change "been" to "survived" and change "the" to "her". Line 58, change "she has" to "with". Remove the rest of the sentence starting "in the performed image..." through to line 59. Line 20, insert the word "the" between "in" and "absence"

Paragraph 5. Line 24, change "in searching for" with "SDH screening" and add at the end of this sentence "should be undertaken".

ALL THE CORRECTIONS HAVE BEEN MADE.