Reviewer’s report

Title: Premature ovarian insufficiency as a variable feature of blepharophimosis, ptosis, and epicanthus inversus syndrome associated with p.Leu75Phe FOXL2 mutation: a case report.

Version: 0 Date: 06 Jan 2019

Reviewer: Yueqiu Tan

Reviewer's report:

This paper described a BPES patient with POI, and identified a reported missense mutation (p.Leu75Phe) in FOXL2. This mutation was found in three women of two Chinese families, with typical eyelid malformations, but free of POI. The result indicate possibility of different phenotypes should be considered in patients with a similar genetic mutation. This is a well-written study. However, I do have two major concerns over the manuscript.

1. The authors should add the strategy of variant filtering in Whole exome sequencing.

2. Other causes of POI, such as endocrinopathies, autoimmune disorders, karyotype, and FMR1 CGG repeats, should be excluded.

Other minor points:

1. Lines 68: the authors should declare the structure of FOXL2, which contains two domains: the DNA-binding forkhead domain (FHD) and the 14-residue polyAla tract.

2. Lines 179-180: There is a grammatical error in this sentence.

3. Lines 92; Lines185; Lines214: BPES type I manifests with palpebral malformations associated with POI, whereas patients with BPES type II exhibit isolated eyelid malformations. However, the patients of this study were classified as BPES type I?

Are the methods appropriate and well described?
If not, please specify what is required in your comments to the authors.

Yes

Does the work include the necessary controls?
If not, please specify which controls are required in your comments to the authors.

Unable to assess
Are the conclusions drawn adequately supported by the data shown?
If not, please explain in your comments to the authors.

Yes

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If an additional statistical review is recommended, please specify what aspects require further assessment in your comments to the editors.

Not relevant to this manuscript

Quality of written English
Please indicate the quality of language in the manuscript:

Needs some language corrections before being published

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