Reviewer's report

Title: A Novel Mutation in the Glycogen Synthase 2 Gene in a Child with Hypoglycemia Due to Glycogen Storage Disease Type 0

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Reviewer: Tjin-shing Jap

Reviewer's report:

In this paper, Soggia et al. have reported an interesting case of Glycogen storage disease (GSD0) in a patient with 6 months of age. They found a compound heterozygous for one previously nonsense mutation (c.736 C>T; R243X) which was inherited down from the mother and a novel frame-shift mutation (966_967 delGA/insC) from the father, respectively.

Major comments:
1. In background, please describe a little more about the properties of hepatic glycogen synthase including the KDalton, when GYS2 gene was cloned, and how many exons, etc. In addition, how many cases with GSD0 have been reported in your home country?
2. The glucose profile after moderate alcohol challenge study may be repeated again in the father who experienced hypoglycemia after alcohol intake; otherwise, the result was unable to suggest haploinsufficiency.
3. Please specify the gender of patient and his or her glycated hemoglobin level.
4. The context of discussion is disproportional to the results found and please shorten the discussion not more than 2 pages in the present form.

Minor comments:
1. The title of manuscript should be concise and informative. Please delete “hypoglycemia due to”
2. All data should be expressed in SI units.
3. Please delete irrelevant data including TSH, free T4, LH and FSH.
4. In background, the statement of “an invasive approach that can be inconclusive since assays to measure the activity of the enzyme GS are not performed routinely in clinical laboratory” is not appropriate because the authors may send it to research laboratory elsewhere.
5. The first paragraph of discussion may be moved to Introduction.
6. Redundancy is not uncommon in the manuscript, e.g.,
   a. At 2 years of age, diabetes mellitus was suspected because of healing difficulties at foot wound.
   b. Tanner stage
c. Further studies are needed to investigate this issue.

In summary,

In conclusion, this paper requires substantial re-writing to improve the fluency of the text.

I was unable to read the tables from the manuscript.