

July 15, 2013

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 3796-review.doc).

**Title:** *ABCB4* mutations underlie hormonal cholestasis but not pediatric idiopathic gallstones

**Author:** Milan Jirsa, Jiří Bronský, Lenka Dvořáková, Jan Šperl, Vít Šmajstrla, Jiří Horák, Jiří Nevoral, Martin Hřebíček

**Name of Journal:** *World Journal of Gastroenterology*

**ESPS Manuscript NO:** 3796

The manuscript has been improved according to the suggestions of reviewers:

1 Format has been updated

2 Revision has been made according to the suggestions of the reviewer

(1) Some minor points

1. According to patients and methods and Fig. 1, there were 41 patients with likely idiopathic gallstones including in this study. Only were 35 subjects selected with gallstone family history. Did the other 6 subjects have no the family history?

Yes, this is correct. The other 6 subjects had negative family history of gallstones since their parents and grandparents had no gallstones. To make this clear we added the words "with positive family history" to the bracket on page 5 row 2: "These subjects (15 males and 20 females with positive family history), all unrelated Caucasians..."

2. The 'patients' in the article should be 'patients'.

The typing error was corrected.

3. Five young females and their families (15 additional family members) were indicated in the methods of abstract, however, in the results of abstract were 16 members of the studied 5 LAPC kindreds.??

Five young females and 15 additional family members (i.e. 20 individuals) provided their DNA samples as stated in the **methods** section of the abstract. DNA samples from the other 3 living family members were not accessible (not mentioned in the abstract, these individuals are indicated as "n.a." in family trees on Fig. 2). One proband was a compound heterozygote for two different mutations in *ABCB4*, the remaining 4 probands and 10 family members were heterozygotes. The father of the proband **III** (F1-1) was not examined; however, he is most likely a heterozygous carrier of c.523A>G (p.Thr175Ala) as depicted in Fig. 2. So in summary, 15 members of the studied 5 LAPC kindreds were confirmed and another one was highly suspected to carry predictably pathogenic mutations in *ABCB4*. The sentence was changed in the revised abstract to make this more clear.

The original document was converted from pdf to MS Word, formatted according to the "Format for brief letter", references and typesetting were checked and corrected and COMMENTS were added in

line with the "Format for highlighted contents". Major changes were highlighted in yellow. Finally, the revised version was edited by a language editor employed in our institute. The language certificate is enclosed.

Thank you again for publishing our manuscript in the *World Journal of Gastroenterology*.

Sincerely yours,

A handwritten signature in blue ink, appearing to read 'Milan Jirsa', is shown within a light blue rectangular border.

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