

ANSWERING TO REVIEWERS



February 12, 2014

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 8899-P Zezos E Renner REVISED.doc).

Title: Liver Transplantation and Non-Alcoholic Fatty Liver Disease

Author: Petros Zezos and Eberhard L. Renner

Name of Journal: *World Journal of Gastroenterology*

ESPS Manuscript NO: 8899

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. Genetic predisposition and/or setting of donor, can present a role in the recurrence of NAFLD after OLT? And the type of diet (i.e. Mediterranean diet)?

- a. Recent data suggest that the genetic predisposition plays role in the post-LT recurrence of NAFLD. It is well established that the patatin-like phospholipase domain-containing protein 3 (PNPLA3) rs738409-G allele is associated with an increased fat accumulation in the liver, and is a risk factor for developing more advanced liver disease in NAFLD patients^[42; Rotman et al, 2010]. A recent study showed that the presence of the rs738409-G allele of the PNPLA3 in the recipients, but not in the donors, is an independent risk factor for post-LT steatosis^[43; Finkenstedt et al, 2013]. These findings are in keeping with a previous study demonstrating that the presence of the PNPLA-3 rs738409 G- allele was associated with the development of post-LT obesity and diabetes mellitus^[44; Watt et al, 2013].

The above paragraph has been incorporated in the manuscript.

- b. Dietary modifications could affect the progression of NAFLD. In a recent study, Kontogianni *et al* ^[49; Kontogianni et al, 2013] found that patients with NAFLD adhering to a Mediterranean diet had less severe liver disease and lower degree of insulin resistance (IR)^[49]. The above have been incorporated to the manuscript.
2. Is present in literature an intention to treat analysis on the insulin-resistance treatment before OLT in these patients?

We are not aware of solid data on the effects on post liver transplant outcomes of treatment of insulin-resistance prior to liver transplantation in patients with NAFLD.

The sentence has been added to the manuscript.

3. Information about liver transplantation cost in the text, may be useful.

It is difficult to estimate the cost of transplantation because there are many differences between countries and liver transplantation programs.

4. Please use the acronym IR in the text

Corrected

5. Several double-spaces are present in the text

Corrected

- 3 References and typesetting were corrected

- 4 Figures have been sent in a separate file (ppt)

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

Sincerely yours,

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