ANSWERING REVIEWERS

June 09, 2014

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 10990-review.doc).

Title: Employing Extracellular Vesicles for Non-invasive Renal Monitoring: A captivating prospect

Author: Gabriella Pocsfalvi, Christopher Stanly, Annalisa Vilasi, Immacolata Fiume, Rosarita Tatè, Giovambattista Capasso

Name of Journal: *World Journal of Clinical Urology*

ESPS Manuscript NO: 10990

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

1 Format has been updated according to BPG' Revision Policies for Review.
2 Authorship: Authors contribution has been updated.
3 Fax number was added to corresponding author's address.
4 Abstract was revised for more-fluent English and for clarity.
5 An additional key word was added: urine analysis.
6 Figure 1 legend has been corrected for clarity: "Figure 1 Schematics of Nephron showing the urinary extracellular vesicles-derived proteins identified in the different segments: A) Glomerulus/podocyte, B) Proximal Convoluted Tubule, C) Ascending limb, E) Distal Convoluted Tubule and D) Collecting Duct."
7 Figure 2 legend has been corrected for clarity: "Figure 2 A) Transmission electron microscopy and B) Nanoparticle Tracking Analysis images of urinary extracellular vesicles isolated in the 1 M sucrose/deuterium-oxide fraction of urinary extracellular vesicles purified from healthy male urine samples by the sucrose cushion ultracentrifugation method."
8 Typesetting were corrected.
9 Revision has been made according to the suggestions of the reviewer.
   (1) In particular, we implemented the Discussion part of the review with a paragraph describing the so far achieved results and future perspectives of urinary EVs in the study of renal transplantation. Exosomes are involved in important signal transduction events during organ transplantation in general. More specifically, the expression level of NDAG has been studied and it was found to sensitively change upon renal transplantation. (ref 90) To highlight these important finding the following was added into the revised version of the paper: "NGAL is suggested to be used as a urinary biomarker of delayed graft function (DGF), a frequent complication after kidney transplantation too. Recently Alvarez *et al* have demonstrated that NGAL is mainly secreted into urinary vesicles and that the expression level of NGAL in uEVs is a sensitive measure of DGF,[90] findings which might support the clinical management of patients undergoing kidney transplantation."
   (2) PMID and/or doi were added to References.

Thank you again for revising and publishing our manuscript in the *World Journal of Clinical Urology*.

Sincerely yours,

Gabriella Pocsfalvi PhD

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