

March 12, 2014

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 9046-review.doc).

**Title:** In 2014, can we do better than CA125 in the early detection of ovarian cancer?

**Author:** Joshua G. Cohen MD, Matthew White BS, Ana Cruz MS, and Robin Farias-Eisner MD, PhD

**Name of Journal:** *World Journal of Biological Chemistry*

**ESPS Manuscript NO:** 9046

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

1) The format has been updated in response to the revision policies for topic highlights. We have added the "Core tip" and author contributions in detail. We have structured the manuscript per the example provided in the format for topic highlights. With adjustment of font type to Book Antiqua with 10 point size, numbering of the lines has changed.

2) Revisions have been made according to the suggestions of the reviewers

Reviewer 1

- (1) Lines 283-285: we have added these results for ROMA and RMI from reference 89 to Table 2. The sensitivity and specificity percentages for ROMA and RMI in reference 89 reflect a broader spectrum of values provided for the reader in Table 2. Various results are cited in Table 2 to demonstrate this spectrum. We have not added sensitivity and specificity for skilled ultrasonographers in Table 2, as the table is meant to focus on algorithm based screening strategies.
- (2) Lines 289-291: we have corrected the order of sensitivity and specificity percentages listed for ROMA and LR2 and they now reflect the same percentages as given in Table 2.
- (3) Line 103: caner has been corrected to cancer
- (4) Table 2 references (ie 44-52) have been removed from lines 178 and 296 based on revision policies. This was also done with Table 3 references (ie 54-64) on line 200. Table 2 and 3 both have cited references within the tables to identify pertinent sources.
- (5) Line 373: sited has been changed to cited

Reviewer 2

- (1) Lines 87-88: we have adjusted the sentence structure to clarify that a woman in the US has a 1 in 70 risk of developing ovarian cancer over her lifetime.
- (2) Line 254: pathology has been changed to morphology.
- (3) Lines 262-270: we have incorporated additional text to clarify that ultrasound screening does not increase disease-free five year survival by itself.
- (4) Lines 360-363: we have provided our opinion that the UKCTOCS represents the best opportunity to date in multimodality screening because of its use of an individualized algorithm in the form of ROCA.

Additional edits:

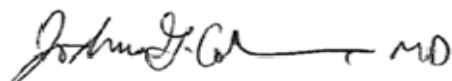
Lines 338-344 were shortened with the statistics updated for sensitivity and specificity specific to invasive epithelial ovarian and tubal cancers.

All three tables have been placed at the end of the paper in addition to the legend for Figure 1. Figure 1 has been uploaded separately as instructed.

References and typesetting were corrected per the revision guidelines. All references have a PMID except reference 69 and reference 113. Reference 69 is not indexed in PubMed and a copy of the article has been uploaded to the journal submission site. Reference 113 refers to the National Comprehensive Cancer Center (NCCN) Guidelines available online through [nccn.org](http://nccn.org). If the journal desires to place an electronic link to these guidelines within the bibliography itself, written permission from the NCCN will be required. We can assist with this if so desired by the editors of *WJBC*. DOI information has been provided for all references when this information was available.

We truly appreciate the time that the reviewers and editors have taken to provide thoughtful and constructive comments to improve the manuscript. Thank you again for the opportunity to contribute to the sixth anniversary issue of the *World Journal of Biological Chemistry*.

Sincerely yours,

A handwritten signature in black ink, appearing to read 'Joshua G. Cohen MD', with a stylized flourish at the end.

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