

Submitted Manuscript:

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Answering to the Reviewers' comments

We thank all the reviewers for their valuable suggestions. We tried our best to answer them. Please note, all the corrections/ modifications suggested by the reviewers are mentioned in color in the text manuscript.

Reviewer 1:

The authors reviewed roles of Wnt/beta-catenin signaling in urothelial carcinoma of bladder and urothelial cancer stem cells (UCSCs). The review described molecular components including signaling proteins, miRNAs, small agents and drug that affect Wnt/beta-catenin signaling. Through the review, I came to know many components that affect Wnt/beta-catenin signaling without search other papers. It seems to me that the review is well organized and written. In particular, Table and figures really helped their description to be understood because many molecular components were introduced in this review.

Reviewer 2:

The mini-review is interested. However, there are minor comments as follows: - There are some typo errors. - Figure 2 needs correction, yellow color is confusing with white background and some works did not appear.

Manuscript was read carefully. Typological errors at places were corrected. Corrections are marked in color. Color scheme in Fig.2 has been modified to make it readable.

Reviewer 3:

This is a well-written review paper about WNT/b-catenin signaling in urothelial carcinoma of bladder (UCB). The authors discuss molecular mechanisms involved in urothelial cancer stem cells and therapeutic potentials of anti-WNT/b-catenin signaling strategy in details. I have only minor comments.

1. Please make clearer canonical WNT signaling and non-canonical WNT signaling in Fig. 1.

Fig. 1 has been modified. Canonical Wnt pathway is shown in black color arrows whereas non-canonical Wnt pathway is shown in red color arrows.

2. Please compare the involvement of WNT/b-catenin signaling in UCB with that in other types of cancer. What mechanisms are specific to UCB?

Few recent papers on aberrant Wnt signaling in other human cancers are cited in the manuscript (please refer page no. 10; lines no.: 14-28 and page no. 11: lines no. 1-2). New references (ref no. 29 to 33) are added. Nevertheless detailed description on aberrant Wnt signaling in human cancers other than bladder cancer is out of the scope of this manuscript.