Author’s response to reviews

Title: Cold exposure induced oxidative stress and apoptosis in the myocardium by inhibiting the Nrf2-Keap1 signaling pathway

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Author’s response to reviews:

Dear Editors,

Thank you for your last letter. We have read your comments seriously and uploaded revised manuscript online which includes a report of additional experiments done at the referee’s suggestion. The followings are the responses to the comments of reviewers.

To Reviewer 1:

Dear Dr. Cai,

Thank you for your referee’s comments concerning our manuscript. We have studied your comments carefully and have made correction which we hope meet with your approval.
1. Firstly, we corrected the grammar mistakes in our manuscript.

2. Second, we rechecked our primers again and corrected sequences of Bad, Nrf2 and Keap1, and add Bcl-2 in them. “Bad 5′-GAGCAGGAAGACGCTAGTGC-3′ (forward), 5′-GGTACGAAGACGCTAGTGC-3′ (reverse); Bcl-2 5′-TGAGTACCTGAACCGGCATC-3′ (forward), 5′-AAGCAGAAGACGCTAGTGC-3′ (reverse); Nrf2 5′-CCCAGCAGCATGCGATTTGA-3′ (forward), 5′-AGCTCATATCCTTCTGTCGC-3′ (reverse); Keap1 5′-CCAGATATCCTGTCGC-3′ (forward), 5′-GGTGAAGAAGACGCTAGTGC-3′ (reverse).”

3. Third, in “Cold exposure increased apoptosis in the myocardium” section, we followed your suggestion and detected Bcl-2 for western blot and real time PCR.

4. Fourth, we cited five new literatures, including the two you mentioned.


Thanks again for your concern and pertinent comments. Your patience and attentive attitude inspire us to be better.

To Reviewer 2:

Dear Dr. Ren,

Thank you for your referee’s comments concerning our manuscript. We have studied your comments carefully and have made correction which we hope meet with your approval.
1. We have used English language editing service to ask professional help in revising this manuscript. And the edit tracked copy has been uploaded.

2. We changed pictures of figure 2. They are more sharpen. And we will prolong exposure time to explore the new changes in the future. Your comments provided new direction for us.

3. We have attempted to quantitate figure 3, but couldn’t find proper primers. So we made relative density (3’-NT or 4HNE/GAPDH) instead.

As for the further suggestions about cell signaling pathway, we have realized that make an intensive study of Nrf2 could better explain our findings. And we will improve the depth of cell signaling pathway in further study.

4. We have read two references you mentioned, and did preliminary discussion in the revised manuscript.

Thanks again for your concern and pertinent comments. Your professional suggestions enlightened our mind and attentive attitude inspired us to be better.

That’s our point-by-point response to the comments of reviewers. We have changed more sharpen figure 2, detected Bcl-2 for western blot and real-time PCR and add new citations to discuss better. And we have used English language editing service to ask professional help in revising this manuscript. We are so grateful to be given such constructive suggestions to our manuscript.