

Review of: "[Research Note] Endoplasmic Reticulum Stress: Unfolding the Impact on Cellular Environment, Anaerobic Respiration, Tumor Activity, And the pre-glucolipotoxicity stage"

Thomas Simmen¹

1 University of Alberta

Potential competing interests: No potential competing interests to declare.

Akl et al. have written a short letter exploring the connections between anaerobic respiration and tumor growth. The text lists multiple proven facts and hypotheses without links between each other. This makes the current letter not accessible.

Specific Points:

- 1. What is meant by "tumor activity" in the title?
- 2. Which epigenetic effects are exerted by ER stress?
- 3. The link between the sentence describing immune responses and the preceding text is missing. Also, it is not clear why autoimmune diseases are mentioned.
- 4. ER stress activates mitochondria, unlike what the authors mention. Maybe a shift to fermentation occurs during long-term ER stress, but any such effects should be delineated.
- 5. Instead of providing a vague link to p53, the authors should elaborate on the significance of hypoxia, ER stress, and tumor growth/metastasis.
- 6. The second paragraph currently completely veers off-topic, and there is no link with the preceding text. This paragraph also contains a multitude of fact listings with unknown connections and significance. The authors should decide what they want to write on and stick to that topic, describing the findings in a logical, accessible manner. This is most important in the second paragraph and to a lesser extent in the first paragraph.

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