

## Endoplasmic reticulum stress plays significant role in type 2 diabetes

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A new research report published in the October 2015 issue of the *Journal of Leukocyte Biology*, suggests that the endoplasmic reticulum plays a more important role in type 2 diabetes and its complications than previously believed. Specifically, when this cell compartment undergoes disruptions (stress) patients experienced poor glycemic control, dyslipidemia, insulin resistance and inflammatory and oxidative stress. Targeting ER stress might be an effective treatment strategy for type 2 diabetes.

"In a way, we observed a 'disharmonious stress signaling alliance' in diabetes," said Muthuswamy Balasubramanyam, a senior researcher and disease-biologist involved in the work from the Madras Diabetes Research Foundation in Chennai, India. "The future of potential practical applications of our research is that this would lead to the development of 'chemical chaperones' as a new generation class of antidiabetic agents that should improve ER folding capacity, help to stabilize protein conformation and ameliorate ER stress," he added.

To make their discovery, scientists identified several <u>endoplasmic</u> <u>reticulum</u> stress markers increased in cells from patients with type 2 diabetes as compared to healthy control subjects. These endoplasmic reticulum stress markers also correlated significantly and positively with <u>poor glycemic control</u>, dyslipidemia, <u>insulin resistance</u> and inflammatory and <u>oxidative stress</u> markers. In addition, the scientists found that one of the microRNAs called "miR-146a" (which is supposed to regulate inflammatory responses) was also impaired and negatively correlated to



endoplasmic reticulum stress and inflammation in patients with diabetes.

"In an increasing number of diseases we are finding links between stress and inflammation," said John Wherry, Ph.D., Deputy Editor of the *Journal of Leukocyte Biology*. "There has also been great work in being able to translate the vague notions of stress, the way we think about it non-scientifically, into true molecular events associated with physiological changes in tissues and cell. This work represents an interesting advance in thinking about how to use that molecular knowledge to develop possible molecular targets for reducing <u>stress</u>, inflammation and ameliorating disease."

**More information:** Raji Lenin, Aravind Sankaramoorthy, Viswanathan Mohan, and Muthuswamy Balasubramanyam. Altered immunometabolism at the interface of increased endoplasmic reticulum (ER) stress in patients with type 2 diabetes. *J. Leukoc. Biol.* October 2015 98:615-622; DOI: 10.1189/jlb.3A1214-609R

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