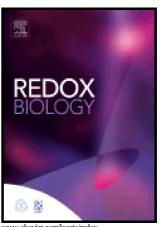
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Erdi Sozen, Nesrin Kartal Ozer



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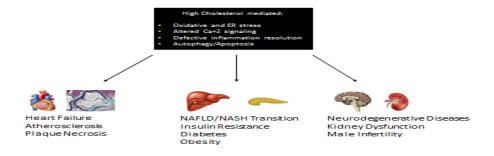
Impact of High Cholesterol and Endoplasmic Reticulum Stress on Metabolic Diseases: An Updated Mini-Review

Erdi Sozen, Nesrin Kartal Ozer*
Department of Biochemistry, Faculty of Medicine, Genetic and Metabolic Diseases Research and Investigation Center (GEMHAM), Marmara University, 34854, Maltepe, Istanbul, Turkey

Abstract

Endoplasmic reticulum (ER) is the major site of protein folding and calcium storage. Beside the role of ER in protein homeostasis, it controls the cholesterol production and lipid-membrane biosynthesis as well as surviving and cell death signaling mechanisms in the cell. It is well-documented that elevated plasma cholesterol induces adverse effects in cardiovascular diseases (CVDs), liver disorders, such as non-alcoholic fatty liver disease (NAFLD), non-alcoholic steatosis hepatitis (NASH), and metabolic diseases which are associated with oxidative and ER stress. Recent animal model and human studies have showed high cholesterol and ER stress as an emerging factors involved in the development of many metabolic diseases. In this review, we will summarize the crucial effects of hypercholesterolemia and ER stress response in the pathogenesis of CVDs, NAFLD/NASH, diabetes and obesity which are major health problems in western countries.

Graphical abstract



Abbreviations

ATF6, activating transcription factor 6; CHOP, C/EBP-homologous protein; CVD, cardiovascular disease; eIF2α, eukaryotic translation initiator factor 2α; ER, endoplasmic reticulum; GRP78, glucose regulated protein 78; IDL, intermediate-density lipoprotein; IRE1, inositol requiring kinase 1; JNK, c-Jun N-terminal kinase; LDL, low-density lipoprotein; NAFLD, non-alcoholic fatty liver disease; NASH, non-alcoholic steatosis hepatitis; Ox-LDL, oxidized-LDL; PERK, RNA-activated protein kinase-like endoplasmic reticulum kinase; ROS,

^{*}Corresponding author. nkozer@marmara.edu.tr

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