Mycotoxins, in a Prospection in the Causes and Treatment of Metabolic Syndrome

Abstract
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Mycotoxins, in chicks showed a strong malabsorption syndrome. Aflatoxin, in chicks showed a strong malabsorption syndrome occurring in MS. Till date, a number of risk factors as well as treatment strategies have been introduced in MS. Metcotoxins are one of them. This paper features mycotoxins as a risk factor as well as therapeutic tools of MS.

Keywords: Mycotoxins; Metabolic syndrome; Prospaction

Abbreviations: CVD: Cardiovascular Disease; MS: Metabolic Syndrome; HDL: High-Density Lipoprotein; NDDs: Chronic Neurodegenerative Diseases; LPS: Lipopolysaccharides; NTIS: Non-Thyroidal Illness Syndrome

Introduction
Abdominal (central) obesity, elevated blood pressure, elevated fasting plasma glucose, high serum triglycerides and low high-density lipoprotein (HDL) levels are the key features of the metabolic syndrome (MS) that is associated with the risk of developing cardiovascular disease (CVD) and type 2 diabetes [1].

Various strategies have been proposed to prevent/management the development of MS. Among them, changing of lifestyle and food habit is most common. A number of drug therapies are also recommended to treat or control of MS. To date, a number of risk factors also identified for the development of MS. Mycotoxins, toxic secondary metabolites produced by fungus kingdom [2] are not only evident to cause MS but also helpful to treat MS. This article depicts a perspection on myotoxins in MS.

Methodology
A search was made in the PubMed database for published articles on the following keywords: mycotoxins in metabolic syndrome; mycotoxin-induced metabolic syndrome; mycotoxins in the treatment of metabolic syndromes. Articles from the past to till date, concerning the search topic was collected and used for this review. Articles falling outside the scope of this study were discarded by reading the titles, abstracts and contents.

Mycotoxins Causing MS
Aflatoxin, in chicks showed a strong malabsorption syndrome characterized by steatorrhea, hypocarotenoidemia, and decreased concentrations of bile salts and pancreatic lipase, trypsin, amylase, and RNase while T-2 characterized by steatorrhea and decreased levels of pancreatic lipase, trypsin, amylase, and RNase [3]. In the same study, ochratoxiosis was evident to show severe hypocarotenoidemia. Though, T-2 toxicosis exhibited lipid malabsorption in the absence of hypocarotenoidemia, ochratoxiosis exhibited hypocarotenoidemia in the absence of lipid malabsorption, but aflatoxiosis exhibited both symptoms.

In low content of vitamin E (6mg/kg as compared with 100 mg/kg in control), T-2 mycotoxin in rats decreased in activity of lysosomal enzymes, aniline hydroxylase, carboxyl esterase and in content of cytochrome P-450 in liver tissue simultaneously with twofold activation of epoxide hydrolase and UDP-glucuronosyl transferase; decreased in non-sedimented activity of lysosomal enzymes; decreased in activity of alkaline phosphatase and of lysosomal enzymes in blood serum [4].

The role of large-conductance Ca$^{2+}$-activated K$^+$ (BK (Ca)) channels in the regulation of coronary microvascular function is widely appreciated. In a study, male Ossabaw miniature swine consumed in 3-6 mo a normal diet (11% kcal from fat) or an excess-calorie atherogenic diet that induces MetS (45% kcal from fat, 2% cholesterol, 20% kcal from fructose). MetS significantly impaired coronary vasodilation to the BK(Ca) opener NS-1619 in vivo (30 - 100µg) and reduced the contribution of these channels to adenosine-induced microvascular vasodilation in vitro (1 - 100µM). MetS reduced whole cell penitrem A (1 µM)-sensitive K$^+$ current and KS-1619-activated (10µM) current in isolated coronary vascular smooth muscle cells. MetS increased the concentration of free intracellular Ca$^{2+}$ and augmented coronary vasoconstriction to the L-type Ca$^{2+}$ channel agonist BAY K 8644 (10PM - 10nM). BK (Ca) channel alpha and beta (1) protein expression was increased in coronary arteries from MetS swine. Coronary vascular dysfunction in MetS is related to impaired BK (Ca) channel function and is accompanied by significant increases in L-type Ca$^{2+}$ channel-mediated coronary vasoconstriction [5]. In another study of this research group, penitrem A (10µg/kg, i.v) revealed that the exercise-induced increases in blood pressure were significantly elevated in MetS swine [6].

Chronic neurodegenerative diseases (NDDs) are associated with obesity and diabetes worldwide, mainly the Parkinson’s and Alzheimer’s disease. MetS involves lipoprotein abnormalities and insulin resistance is the major cause of induction of NDDs. The effects of bacterial lipopolysaccharides (LPS) on dyslipidemia...
and NAFLD indicate that the clearance and metabolism of fungal mycotoxins are linked to hypercholesterolemia and amyloid beta oligomers. LPS and mycotoxins are associated with membrane lipid disturbances with effects on cholesterol interacting proteins, lipoprotein metabolism, and membrane apo E/amyloid beta interactions relevant to hypercholesterolemia with close connections to NDDs [7].

Long-term exposure to dampness microbiota induces multi-organ morbidity. One of the symptoms related to this disorder is non-thyroidal illness syndrome (NTIS). A retrospective study was carried out in nine patients with a history of mold exposure, experiencing chronic fatigue, cognitive disorder, and different kinds of hypothyroid symptoms. Mycotoxin exposure via the diet is underlined as DIO2 genetic polymorphism and dysfunction of DIO2 play an important role in the development of symptoms [8].

**Mycotoxins Against MS**

Drugs or products acting as chemo-modulators were assayed for their effect on the chemotactic activity. Cytochalasin B is evident by interfering with the cytoskeletal elements (microtubules and microfilaments) cyclic nucleotides involved in the metabolic activity during the cell motility [9]. In another study, AFB1 (0.1, 0.3 or 1 µg/mL) in Li-Fraumeni patient, but not cells from normal individuals, can induce immortalization [10].

**Conclusion**

Mycotoxins are one of the important causes of MS. However, some MS along with the related phenomena could be taken into account that there is an option to select mycotoxins and their laboratory derivatives to be treated. Mycotoxins are important lead compounds in MS and MS-related complications.

**Conflict of Interest**

None declared.

**References**