

A commentary on Effect of Chebulagic Acid from the Fruits of *Terminalia chebula* Retz

Abstract

In the present study, we firstly compared rat intestinal α glucosidase inhibitory activity by different ethanol-aqueous extractions from the dried fruits of *Terminalia chebula* Retz. The enzymatic assay showed that the 80% ethanol extract was more potent against maltase activity than both 50% and 100% ethanol extracts. By HPLC analysis, it was determined that the 80% ethanol extract had a higher content of chebulagic acid than each of 50% or 100% ethanol extract. Next, we investigated how efficiently chebulagic acid could inhibit sugar digestion by determining the glucose level on the apical side of the Caco-2 cell monolayer. The results presented here suggest that chebulagic acid from *T. chebula* can be used to control blood glucose and manage type 2 diabetes, although clinical trials are needed.

Keywords: Terminalia • chebulachebulagic acid • α -glucosidase inhibitor • anti-hyperglycemia

Introduction

Diabetes mellitus (DM) is a common metabolic disorder characterized by hyperglycemia, which is the main cause of complications related with micro- and macro-vascular diseases. DM is one of the three leading causes of death worldwide and constitutes a major health problem. Postprandial hyperglycemia results from abnormal insulin secretion by β -cells in response to a meal, impaired hepatic glucose production, and defective glucose uptake by peripheral insulin-sensitive tissues, particularly the skeletal muscles. Therefore, control of postprandial plasma levels is critical in treatment of not only diabetic patients but also individuals with impaired glucose tolerance. Mammalian intestinal α glucosidase (EC 3.2.1.20) is the key enzyme, which catalyzes the final step in the digestive process of carbohydrates. Hence, α -glucosidase inhibitors can reduce postprandial blood glucose levels and absorption of starch and disaccharides.

we compared rat intestinal α -glucosidase inhibitory activity of different ethanol-aqueous extractions and detected chebulagic acid in these extracts by HPLC analysis. Meanwhile, we determined α -glucosidase inhibitory activity of chebulagic acid with the Caco-2 cell monolayer, together with evaluating the postprandial blood glucose lowering effect of chebulagic acid after sugar (maltose, sucrose or glucose) loading in Sprague-Dawley (SD) rats.

Description

Extraction of phenolic compounds from *T. chebula* is generally carried out using various types of organic solvents such as 95% ethyl acetate, hot water, 70% methanol, and 95% ethanol. Ethanolic extraction of plant bioactives has displayed a higher yield compared with the aqueous extract. In the present study, we compared rat intestinal α -glucosidase inhibitory activity of different ethanol-aqueous extractions. the maltose-hydrolysis inhibitory activity of 50–100% ethanol extracts of *T. chebula* fruits. Each of these ethanol

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extracts significantly inhibited the maltase activity, and the enzymatic inhibition was dose independent. In the same assay condition, the IC₅₀ value of chebulagic acid was determined to be 37 µg/mL. On the other hand, the IC₅₀ values of 50%, 80% and 100% ethanol extracts against maltase were determined to be 173.6 µg/mL, 51.7 µg/mL and 85.7 µg/mL, respectively. Hence, this result revealed that the 80% ethanol extract was more potent regarding its effect on maltase activity than both 50% and 100% ethanol extraction.

In the present study, we compared rat intestinal α-glucosidase inhibitory activities of different ethanol-aqueous extractions from the dried fruits of *T. chebula*. The enzymatic assay showed that the 80% ethanol extract had a more potent effect on maltase activity than both 50% and 100% ethanol extracts. HPLC analysis revealed that the 80% ethanol extract contained a higher content of chebulagic acid than either the 50% or 100% ethanol extract. In a Caco-2 cell model, α-glucosidase activity for maltose hydrolysis was down-regulated by chebulagic acid, which is a reversible inhibitor of maltase. On the other hand, chebulagic acid showed a weak inhibition of sucrose-hydrolysis activity and did not affect intestinal glucose uptake by Caco-2 cells. Furthermore, chebulagic acid

significantly reduced postprandial blood glucose level in maltose-loaded SD-rats. So, it was suggested that chebulagic acid from *T. chebula* may be useful for suppressing postprandial hyperglycemia as a potent anti-diabetic agent, although clinical trials are needed.

Acknowledgement

None

Conflict of Interest

No conflict of interest

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