Therapeutic exploration of betulinic acid in chemically induced hypothyroidism

Muhammad Afzal · Imran Kazmi · Susmita Semwal · Fahad A. Al-Abbasi · Firoz Anwar

Received: 17 May 2013 / Accepted: 26 September 2013 © Springer Science+Business Media New York 2013

Abstract Hypothyroidism is a chronic condition characterized by abnormally low thyroid hormone production. The decreased serum level (>5.1 mIU/l) of thyroid-stimulating hormone (TSH) in blood indicates hypothyroidism. The study was an attempt to access the effect of betulinic acid on chemically induced hypothyroidism in female albino rats. Betulinic acid is a naturally occurring pentacyclic triterpenoid, which has antiretroviral, antimalarial, and anti-inflammatory properties, as well as anticancer potential, by inhibiting topoisomerase. Hypothyroidism was induced in female albino rats using propylthiouracil (PTU) at a dose of 60 µg/kg body weight orally for 1 month. Induction of hypothyroidism was confirmed by increased TSH level. At the end of second month, blood was collected, centrifuged and serum was analyzed for TSH, T3, and T4 level and protocol was terminated by killing of animals. The animals exposed to PTU were treated with pure standard drug thyroxine at a dose of 10 µg/kg of body weight by oral route and the test drug betulinic acid 20 mg/kg by oral route through force feeding in their respective groups. Treatment was carried out for a period of 2 months. Group with PTU-induced hypothyroidism showed an elevation in serum TSH and reduction level, which was restored by the betulinic acid in treated female albino rats. Betulinic acid also reduced the damage caused in the thyroid tissues by PTU, thus minimizing the symptoms of hypothyroidism. Histopathological examinations of the thyroid tissue showed changes in the thyrocytes of PTU-treated group while thyroxine group showed normal thyroid follicles cell architecture and the group treated with betulinic acid also showed marked improvement in the follicles integrity which shows that betulinic acid has some protective activity. This study shows that the betulinic acid has thyroid-enhancing potential by lowering down the TSH levels and reducing the damage caused in the thyroid tissues, thus minimizing the symptoms of hypothyroidism when used anaphylactically in rats.

Keywords Betulinic acid · Hypothyroidism · Propylthiouracil · Thyroxine · Thyroid-stimulating hormone

Introduction

Hypothyroidism is an autoimmune thyroid disease known to common endocrine disorder in the adult population, and its prevalence increases with age and highly reported in women. Hypothyroidism occurs when thyroid gland cannot make adequate thyroid hormone to keep the body running normally [1, 2]. Physiologically thyroid hormones (THs), thyroxine (T4), and 3,5,3'-triiodothyronine (T3) are recognized as key metabolic hormones that play a critical role to control the rate of metabolic reactions in the body, which are essential to normal body development, especially to structural and functional formation of the central nervous system (CNS) [3, 4], throughout life and also regulate the metabolism and an assortment of functions of neurotransmitters [5, 6]. The iodine deficiency is the most common cause of hypothyroidism globally, but either an absence of the thyroid gland or a