

## Thyroid-Insulin Dysfunction during Development

Ahmed R.G.

*Division of Anatomy and Embryology, Zoology Department, Faculty of Science, Beni-Suef University,  
Beni-Suef, Egypt*

**\*Corresponding Author:** *Ahmed R.G., Division of Anatomy and Embryology, Zoology Department,  
Faculty of Science, Beni-Suef University, Beni-Suef, Egypt*

### Letter to Editor

Thyroid hormones (THs) are key hormones regulating the development (Ahmed, 2011, 2012a,b, 2013, 2014, 2015a-c, 2016a-d, 2017a-f; Ahmed et al., 2013a,b, 2014; 2015a,b; Ahmed and Incerpi, 2013; Van Herckel et al., 2013; Ahmed and El-Gareib, 2014; Incerpi et al., 2014; Candelotti et al., 2015; De Vito et al., 2015; El-Ghareeb et al., 2016; Ahmed and El-Gareib, 2017), particularly glucose homeostasis (Fain and Bahouth, 1998), metabolism and energy balance (Wang et al., 2008; Ahmed, 2013). Also, insulin is a vital hormone modifying these biological processes during the development (Hytinanti et al., 2008; Ahmed, 2013; Candelotti et al., 2015; Cremaschi et al., 2016). There is a synergistic mechanism between THs and insulin levels during the perinatal development and growth (Tsai et al., 2004; Hytinanti et al., 2008; Ahmed, 2013). In hypothyroid state, the level of insulin was delayed (Robson et al., 2002). Reversibly, the disturbance in insulin action, in hypothyroid state, might be responsible for the hypothalamic-pituitary-thyroid-axis (HPTA) disorders (Ahmed, 2013, 2016d). Additionally, a symptomatic hypoglycaemia, dyslipidemia, elevated low density lipoprotein (LDL) cholesterol and triglyceride were recorded in hypothyroidism (Mohn et al., 2002; Hueston & Pearson, 2004; American Diabetes Association (ADA), 2011; Marwaha et al., 2011). Also, the reduction in the high density lipoprotein (HDL) cholesterol, overall glucose turnover and peripheral glucose utilization was reported by Miura et al. (1994), Cettour-Rose et al. (2005) and Johnson (2006). The reduction in the glucose level might be caused growth retardation and delayed the sexual maturity (Gutch et al., 2013). Also, the diminution in the level of insulin growth factor-I (IGF-I) is correlated to the disturbance between the activities of THs and insulin that may retard the development and growth (Ramos et al., 2001 & 2002; Ahmed, 2013). Thus, these disorders can reduce the uteroplacental passage to fetuses (Ahmed and El-Gareib, 2017). In conclusion, any disruption in the THs-insulin axis may possibly influence the developmental mechanisms. Further studies are necessary to distinguish the molecular mechanisms of this axis during the development.

**Conflict of interest:** The author declares that no competing financial interests exist.

### References

- Ahmed, R.G., 2011. Perinatal 2, 3, 7, 8-tetrachlorodibenzo-p-dioxin exposure alters developmental neuroendocrine system. *Food Chem. Toxicology*, 49, 1276–1284.
- Ahmed, R.G., 2012a. Maternal-newborn thyroid dysfunction. In *the Developmental Neuroendocrinology*, pp. 1-369. Ed R.G. Ahmed. Germany: LAP LAMBERT Academic Publishing GmbH & Co KG.
- Ahmed, R.G., 2012b. Maternal-fetal thyroid interactions, *Thyroid Hormone*, Dr. N.K. Agrawal (Ed.), ISBN: 978-953-51-0678-4, In Tech Open Access Publisher, Chapter 5, pp. 125-156.
- Ahmed, R.G., 2013. Early weaning PCB 95 exposure alters the neonatal endocrine system: thyroid adipokine dysfunction. *J. Endocrinol.* 219 (3), 205-215.
- Ahmed, R.G., 2014. Editorial: Do PCBs modify the thyroid-adipokine axis during development? *Annals Thyroid Res.* 1(1), 11-12.
- Ahmed, R.G., 2015a. Chapter 1: Hypothyroidism and brain development. In *advances in hypothyroidism treatment*. Avid Science Borsigstr.9, 10115 Berlin, Berlin, Germany. Avid Science Publications level 6, Melange Towers, Wing a, Hitec City, Hyderabad, Telangana, India. pp. 1-40.
- Ahmed, R.G., 2015b. Hypothyroidism and brain developmental players. *Thyroid Research J.* 8(2), 1-12.

- Ahmed, R.G., 2015c. Editorials and Commentary: Maternofetal thyroid action and brain development. *J. of Advances in Biology*; 7(1), 1207-1213.
- Ahmed, R.G., 2016a. Gestational dexamethasone alters fetal neuroendocrine axis. *Toxicology Letters*, 258, 46–54.
- Ahmed, R.G., 2016b. Neonatal polychlorinated biphenyls-induced endocrine dysfunction. *Ann. Thyroid. Res.* 2 (1), 34-35.
- Ahmed, R.G., 2016c. Maternal iodine deficiency and brain disorders. *Endocrinol. Metab. Syndr.* 5, 223. <http://dx.doi.org/10.4172/2161-1017.1000223>.
- Ahmed, R.G., 2016d. Maternal bisphenol A alters fetal endocrine system: Thyroid adipokine dysfunction. *Food Chem. Toxicology*, 95, 168-174.
- Ahmed, R.G., 2017a. Developmental thyroid diseases and GABAergic dysfunction. *EC Neurology* 8.1, 02-04.
- Ahmed, R.G., 2017b. Hyperthyroidism and developmental dysfunction. *Arch Med.* 9, 4.
- Ahmed, R.G., 2017c. Anti-thyroid drugs may be at higher risk for perinatal thyroid disease. *EC Pharmacology and Toxicology* 4.4, 140-142.
- Ahmed, R.G., 2017d. Perinatal hypothyroidism and cytoskeleton dysfunction. *Endocrinol. Metab. Syndr.* 6, 271. doi:10.4172/2161-1017.1000271
- Ahmed, R.G., 2017e. Developmental thyroid diseases and monoaminergic dysfunction. *Advances in Applied Science Research* 8(3), 01-10.
- Ahmed, R.G., 2017f. Hypothyroidism and brain development. *J. Anim. Res. Nutr.* 2(2), 13.
- Ahmed, R.G., Abdel-Latif, M., Ahmed F., 2015a. Protective effects of GM-CSF in experimental neonatal hypothyroidism. *International Immunopharmacology* 29, 538–543.
- Ahmed, R.G., Abdel-Latif, M., Mahdi, E., El-Nesr, K., 2015b. Immune stimulation improves endocrine and neural fetal outcomes in a model of maternofetal thyrotoxicosis. *Int. Immunopharmacol.* 29, 714-721.
- Ahmed, R.G., Davis, P.J., Davis, F.B., De Vito, P., Farias, R.N., Luly, P., Pedersen, J.Z., Incerpi, S., 2013a. Nongenomic actions of thyroid hormones: from basic research to clinical applications. An update. *Immunology, Endocrine & Metabolic Agents in Medicinal Chemistry*, 13(1), 46-59.
- Ahmed, R.G., El-Gareib, A.W. 2014. Lactating PTU exposure: I- Alters thyroid-neural axis in neonatal cerebellum. *Eur. J. of Biol. and Medical Sci. Res.* 2(1), 1-16.
- Ahmed, R.G., El-Gareib, A.W., 2017. Maternal carbamazepine alters fetal neuroendocrine-cytokines axis. *Toxicology* 382, 59–66.
- Ahmed, R.G., El-Gareib, A.W., Incerpi, S., 2014. Lactating PTU exposure: II- Alters thyroid-axis and prooxidant-antioxidant balance in neonatal cerebellum. *Int. Res. J. of Natural Sciences* 2(1), 1-20.
- Ahmed, R.G., Incerpi, S., 2013. Gestational doxorubicin alters fetal thyroid–brain axis. *Int. J. Devl. Neuroscience* 31, 96–104.
- Ahmed, R.G., Incerpi, S., Ahmed, F., Gaber, A., 2013b. The developmental and physiological interactions between free radicals and antioxidant: Effect of environmental pollutants. *J. of Natural Sci. Res.* 3(13), 74-110.
- American Diabetes Association (ADA), 2011. Standards of medical care in diabetes- 2011. *Diabetes Care*, 34(Suppl. 1), S11–S61 [www.care.diabetesjournals.org](http://www.care.diabetesjournals.org).
- Candelotti, E., De Vito, P., Ahmed, R.G., Luly, P., Davis, P.J., Pedersen, J.Z., Lin, H-Y., Incerpi, I., 2015. Thyroid hormones crosstalk with growth factors: Old facts and new hypotheses. *Immun., Endoc. & Metab. Agents in Med. Chem.*, 15, 71-85.
- Cettour-Rose, P., Theander-Carrillo, C., Asensio, C., Klein, M., Visser, T.J., Burger, A. G., 2005. Hypothyroidism in rats decreases peripheral glucose utilisation, a defect partially corrected by central leptin infusion. *Diabetologia*, 48(4), 624–633.
- Cremaschi, G.A., Cayrol, F., Sterle, H.A., DíazFlaqué, M.C., Barreiro Arcos, M.L., 2016. Thyroid hormones and their membrane receptors as therapeutic targets for T cell lymphomas. *Pharmacol. Res.* pii: S1043-6618(16)00040-2.
- De Vito, P., Candelotti, E., Ahmed, R.G., Luly, P., Davis, P.J., Incerpi, S., Pedersen, J.Z., 2015. Role of thyroid hormones in insulin resistance and diabetes. *Immun., Endoc. & Metab. Agents in Med. Chem.*, 15, 86-93.
- El-Ghareeb, A.A., El-Bakry, A.M., Ahmed, R.G., Gaber, A., 2016. Effects of zinc supplementation in neonatal hypothyroidism and cerebellar distortion induced by maternal carbimazole. *Asian Journal of Applied Sciences* 4(04), 1030-1040.
- Fain, J.N., Bahouth, S.W., 1998. Effect of tri-iodothyronine on leptin release and leptin mRNA accumulation in rat adipose tissue. *Biochem J* 332(Pt 2), 361-366.

- Gutch, M., Philip, R., Saran, S., Tyagi, R., Gupta, K.K., 2013. Re-emergence of a rare syndrome: A case of mauriac syndrome. *Indian J Endocrinol Metab.* 17(Suppl1), S283–S285.
- Hueston, W.J., Pearson, W.S., 2004. Subclinical hypothyroidism and the risk of hypercholesterolemia. *Annals of Family Medicine*, 2, 351–355.
- Hytinanti, T., Kajantie, E., Karonen, S.L., Andersson, S., 2008. Postnatal changes in ghrelin, adiponectin, insulin and leptin concentrations in term newborns. *Int J Endocrinol Metab* 3, 127–134.
- Incerpi, S., Hsieh, M-T., Lin, H-Y., Cheng, G-Y., De Vito, P., Fiore, A.M., Ahmed, R.G., Salvia, R., Candelotti, E., Leone, S., Luly, P., Pedersen, J.Z., Davis, F.B., Davis, P.J., 2014. Thyroid hormone inhibition in L6 myoblasts of IGF-I-mediated glucose uptake and proliferation: new roles for integrin  $\alpha\beta3$ . *Am. J. Physiol. Cell Physiol.* 307, C150–C161.
- Johnson, J.L., 2006. Diabetes control in thyroid disease. *Diabetes Spectrum*, 19(3), 148–153.
- Marwaha, R.K., Tandon, N., Garg, M.K., Kanwar, R., Sastry, A., Narang, A., 2011). Dyslipidemia in subclinical hypothyroidism in an Indian population. *Clinical Biochemistry*, 44(14–15), 1214–1217.
- Miura, S., Iitaka, M., Yoshimura, H., 1994. Disturbed lipid metabolism in patients with sub clinical hypothyroidism: effect of L-thyroxine therapy. *Internal Medicine*, 33(7), 413–417.
- Mohn, A., Di Michele, S., Di Luzio, R., Tumini, S., Chiarelli, F., 2002. The effect of subclinical hypothyroidism on metabolic control in children and adolescents with type 1 diabetes mellitus. *Diabetic Medicine*, 19(1), 70–73.
- Ramos, S., Goya, L., Alvarez, C., Martín, M.A., Pascual-Leone, A.M., 2001. Effect of thyroxine administration on the IGF/IGF binding protein system in neonatal and adult thyroidectomized rats. *Endocrinology* 169, 111–122.
- Ramos, S., Goya, L., Martín, M.A., Escriva, F., Pascual-Leone, A.M., 2002. Influence of hypothyroidism on circulating concentrations and liver expression of IGF-binding proteins mRNA from neonatal and adult rats. *Endocrinology* 172, 363–373.
- Tsai, P.J., Yu, C.H., Hsu, S.P., Lee, Y.H., Chiou, C.H., Hsu, Y.W., Ho, S.C., Chu, C.H., 2004. Cord plasma concentrations of adiponectin and leptin in healthy term neonates: positive correlation with birth weight and neonatal adiposity. *Clin Endocrinol (Oxf)* 61(1), 88–93.
- Van Herck, S.L.J., Geysens, S., Bald, E., Chwatko, G., Delezie, E., Dianati, E., Ahmed, R.G., Darras, V.M., 2013. Maternal transfer of methimazole and effects on thyroid hormone availability in embryonic tissues. *Endocrinol.* 218, 105–115.
- Wang, Y., Lam, K.S., Yau, M.H., Xu, A., 2008. Post-translational modifications of adiponectin: mechanisms and functional implications. *Biochem J* 409, 623–633.

**Citation:** R.G. Ahmed, " Thyroid-Insulin Dysfunction during Development ", *International Journal of Research Studies in Zoology*, vol. 3, no. 4, p. 73-75, 2017.

**Copyright:** © 2017 Authors. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.