

# Effects of L-tryptophan on diencephalic tryptophan hydroxylase gene expression in heat-stressed broilers

Ladan Emadi<sup>1\*</sup>, Saeed Esmaeili-Mahani<sup>2</sup> and Yadollah Badakhshan<sup>3</sup>

<sup>1</sup> Associated Professor, Department of Basic Sciences, Faculty of Veterinary Medicine, Shahid Bahonar University of Kerman, Kerman, Iran

<sup>2</sup> Professor, Department of Biology, Faculty of Sciences, Shahid Bahonar University of Kerman, Kerman, Iran

<sup>3</sup> Assistant Professor, Department of Animal Sciences, Faculty of Agriculture, University of Jiroft, Jiroft, Iran

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## Abstract

The brain monoaminergic system is changed during heat stress condition. Research has shown that commercial poultry are very sensitive animals to heat stress. Hence, this study investigated the effects of L-tryptophan on diencephalic tryptophan hydroxylase type 1 and 2 (TH1 and TH2) gene expression in heat-stressed broiler chicks as animal model. Forty eight, seven-day old broiler chicks were divided into three groups. Chicks were intraperitoneally injected L- tryptophan (25 and 50 mg/Kg) and normal saline. Then they were exposed to the heat stress (39°C) for 5 hours. After 5 hours of treatment, the birds were anesthetized with isoflurane before being euthanized. The brain samples were taken for gene expression evaluation. The data showed declined diencephalic gene expression of TH1 and TH2 in heat stress condition. Tryptophan administration at dose of 50 mg/kg significantly increased the expression levels of TH1 and TH2 in heat stress exposed chicks. It can be concluded that diencephalic serotonergic pathway may have an important role in tryptophan ameliorating effect during heat stress condition.

**Key words:** Broiler, Diencephalon, High ambient temperature, Tryptophan hydroxylase

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\* **Corresponding Author:** Ladan Emadi, Associated Professor, Department of Basic Sciences, Faculty of Veterinary Medicine, Shahid Bahonar University of Kerman, Kerman, Iran  
E-mail: ladan.emadi@ut.ac.ir



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## References

- Amori, L., Wu, H.-Q., Marinozzi, M., Pellicciari, R., Guidetti, P., & Schwarcz, R. (2009). Specific inhibition of kynurenate synthesis enhances extracellular dopamine levels in the rodent striatum. *Neuroscience*, *159*(1), 196–203.
- Chauhan, N. R., Kapoor, M., Singh, L. P., Gupta, R. K., Meena, R. C., Tulsawani, R., Nanda, S., & Singh, S. B. (2017). Heat stress-induced neuroinflammation and aberration in monoamine levels in hypothalamus are associated with temperature dysregulation. *Neuroscience*, *358*, 79–92.
- Chen, Y., Xu, H., Zhu, M., Liu, K., Lin, B., Luo, R., Chen, C., & Li, M. (2017). Stress inhibits tryptophan hydroxylase expression in a rat model of depression. *Oncotarget*, *8*(38), 63247.
- Crane, J. D., Palanivel, R., Mottillo, E. P., Bujak, A. L., Wang, H., Ford, R. J., Collins, A., Blümer, R. M., Fullerton, M. D., & Yabut, J. M. (2015). Inhibiting peripheral serotonin synthesis reduces obesity and metabolic dysfunction by promoting brown adipose tissue thermogenesis. *Nature Medicine*, *21*(2), 166–172.
- Davis, B. P., Engle, T. E., Ransom, J. I., & Grandin, T. (2017). Preliminary evaluation on the effectiveness of varying doses of supplemental tryptophan as a calmative in horses. *Applied Animal Behaviour Science*, *188*, 34–41.
- de Lima, A. P. N., Sandini, T. M., Reis-Silva, T. M., & Massoco, C. de O. (2017). Long-lasting monoaminergic and behavioral dysfunctions in a mice model of socio-environmental stress during adolescence. *Behavioural Brain Research*, *317*, 132–140.
- Fujigaki, H., Yamamoto, Y., & Saito, K. (2017). L-Tryptophan-kynurenine pathway enzymes are therapeutic target for neuropsychiatric diseases: Focus on cell type differences. *Neuropharmacology*, *112*, 264–274.
- Goekint, M., Roelands, B., Heyman, E., Njemini, R., & Meeusen, R. (2011). Influence of citalopram and environmental temperature on exercise-induced changes in BDNF. *Neuroscience Letters*, *494*(2), 150–154.
- Höglund, E., Øverli, Ø., Andersson, M. Å., Silva, P., Laursen, D. C., Moltesen, M. M., Krogdahl, Å., Schjolden, J., Winberg, S., & Vindas, M. A. (2017). Dietary l-tryptophan leaves a lasting impression on the brain and the stress response. *British Journal of Nutrition*, *117*(10), 1351–1357.
- Ishiwata, T., & Greenwood, B. N. (2018). Changes in thermoregulation and monoamine release in freely moving rats during cold exposure and inhibition of the ventromedial, dorsomedial, or posterior hypothalamus. *Journal of Comparative Physiology B*, *188*, 541–551.
- Ito, K., Erwan, E., Nagasawa, M., Furuse, M., & Chowdhury, V. S. (2014). Changes in free amino acid concentrations in the blood, brain and muscle of heat-exposed chicks. *British Poultry Science*, *55*(5), 644–652.
- Ito, Kentaro, Bahry, M. A., Hui, Y., Furuse, M., & Chowdhury, V. S. (2015). Acute heat stress up-regulates neuropeptide Y precursor mRNA expression and alters brain and plasma concentrations of free amino acids in chicks. *Comparative Biochemistry and Physiology Part A: Molecular & Integrative Physiology*, *187*, 13–19.
- Kojima, D., Nakamura, T., Banno, M., Umemoto, Y., Kinoshita, T., Ishida, Y., & Tajima, F. (2018). Head-out immersion in hot water increases serum BDNF in healthy males. *International Journal of Hyperthermia*, *34*(6), 834–839.
- Kumar, P., Pal, A. K., Sahu, N. P., Jha, A. K., Kumar, N., Christina, L., & Priya, P. (2018). Dietary L-Tryptophan potentiates non-specific immunity in *Labeo rohita* fingerlings reared under elevated temperature. *Journal of Thermal Biology*, *74*, 55–62.
- Li, G., Wang, Y., Yan, M., Ma, H., Gao, Y., Li, Z., Li, C., Tian, H., & Zhuo, C. (2016). Time-dependent correlation of BDNF and CREB mRNAs in adult rat brains following acute psychological stress in the communication box paradigm. *Neuroscience Letters*, *624*, 34–41.
- Mellor, A. L., & Munn, D. H. (1999). Tryptophan catabolism and T-cell tolerance: immunosuppression by starvation? *Immunology Today*, *20*(10), 469–473.
- Mota, C. M. D., Branco, L. G. S., Morrison, S. F., & Madden, C. J. (2020). Systemic serotonin inhibits brown adipose tissue sympathetic nerve activity via a GABA input to the dorsomedial hypothalamus, not via 5HT1A receptor activation in raphe pallidus. *Acta Physiologica*, *228*(3), e13401.
- Nakagawa, H., Matsumura, T., Suzuki, K., Ninomiya, C., & Ishiwata, T. (2016). Changes of brain monoamine levels and physiological indexes during heat acclimation in rats. *Journal of Thermal Biology*, *58*, 15–22.

- Newman-Tancredi, A., Depoortère, R., Carilla-Durand, E., Tarayre, J. P., Kleven, M., Koek, W., Bardin, L., & Varney, M. A. (2018). NLX-112, a highly selective 5-HT<sub>1A</sub> receptor agonist: Effects on body temperature and plasma corticosterone levels in rats. *Pharmacology Biochemistry and Behavior*, *165*, 56–62.
- Schaaf, M. J. M., de Jong, J., de Kloet, E. R., & Vreugdenhil, E. (1998). Downregulation of BDNF mRNA and protein in the rat hippocampus by corticosterone. *Brain Research*, *813*(1), 112–120.
- Sinha, R. K. (2008). Serotonin synthesis inhibition by pre-treatment of p-CPA alters sleep-electrophysiology in an animal model of acute and chronic heat stress. *Journal of Thermal Biology*, *33*(5), 261–273.
- Siuciak, J. A., Clark, M. S., Rind, H. B., Whittemore, S. R., & Russo, A. F. (1998). BDNF induction of tryptophan hydroxylase mRNA levels in the rat brain. *Journal of Neuroscience Research*, *52*(2), 149–158.
- Tang, J., Zheng, X., Xiao, K., Wang, K., Wang, J., Wang, Y., Wang, K., Wang, W., Lu, S., & Yang, K. (2016). Effect of boric acid supplementation on the expression of BDNF in African ostrich chick brain. *Biological Trace Element Research*, *170*, 208–215.
- Tanizawa, H., Shiraishi, J., Kawakami, S.-I., Tsudzuki, M., & Bungo, T. (2014). Effect of short-term thermal conditioning on physiological and behavioral responses to subsequent acute heat exposure in chicks. *The Journal of Poultry Science*, *51*(1), 80–86.
- Tanke, M. A. C., Alserda, E., Doornbos, B., van der Most, P. J., Goeman, K., Postema, F., & Korf, J. (2008). Low tryptophan diet increases stress-sensitivity, but does not affect habituation in rats. *Neurochemistry International*, *52*(1–2), 272–281.
- Tomonaga, S., Okuyama, H., Tachibana, T., & Makino, R. (2018). Effects of high ambient temperature on plasma metabolomic profiles in chicks. *Animal Science Journal*, *89*(2), 448–455.
- U. Bello, A., Idrus, Z., Yong Meng, G., Awad, E. A., & Soleimani Farjam, A. (2018). Gut microbiota and transportation stress response affected by tryptophan supplementation in broiler chickens. *Italian Journal of Animal Science*, *17*(1), 107–113.
- Walther, D. J., & Bader, M. (2003). A unique central tryptophan hydroxylase isoform. *Biochemical Pharmacology*, *66*(9), 1673–1680.
- Yoshida, J., Erwan, E., Chowdhury, V. S., Ogino, Y., Shigemura, A., Denbow, D. M., & Furuse, M. (2015). Comparison of centrally injected tryptophan-related substances inducing sedation in acute isolation stress-induced neonatal chicks. *Pharmacology Biochemistry and Behavior*, *129*, 1–6.
- Zhang, R., Tachibana, T., Takagi, T., Koutoku, T., Denbow, D. M., & Furuse, M. (2004). Serotonin modifies corticotropin-releasing factor-induced behaviors of chicks. *Behavioural Brain Research*, *151*(1–2), 47–52.
- Zhou, J., Li, L., Tang, S., Cao, X., Li, Z., Li, W., Li, C., & Zhang, X. (2008). Effects of serotonin depletion on the hippocampal GR/MR and BDNF expression during the stress adaptation. *Behavioural Brain Research*, *195*(1), 129–138.