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## CHRONIC UNPREDICTABLE ENVIRONMENTAL STRESS IMPAIRS BIOCHEMICAL AND PHYSIOLOGICAL HOMEOSTASIS: ROLE IN DIABETES MELLITUS

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**C**hronic unpredictable environmental stress (CUES) may induce predisposition to diabetes mellitus. This study investigates the role of CUES on impaired homeostasis. Stressed group mice (n=20) were exposed to CUES for 16 weeks. Weekly body weight, feed consumption, feed efficiency ratio, fasting blood glucose were monitored. Plasma HbA1c, plasma cortisol, plasma epinephrine and plasma insulin, serum lipids, antioxidants and carbohydrate metabolizing enzymes activity were assessed along with DNA damage and histopathological examination of liver, kidney, pancreas, spleen and skeletal muscles. Semi-quantitative expression of IL-4, IL-6 and  $\beta$ -actin was also assessed. Fasting blood glucose levels & HbA1c in the stressed were significantly higher compared to control ( $p<0.001$ ). Serum lipids were found

insignificantly higher in stressed mice compared to control. Body weights of the stressed mice and feed efficiency ratio were found significant ( $p<0.001$ ). Plasma corticosterone, plasma epinephrine, HOMA-IR (Homeostatic model assessment- insulin resistance) was found to be significantly higher in the stressed group ( $p<0.001$ ). Plasma insulin level was found to be significantly lower in the stressed group ( $p<0.001$ ). Significant changes were observed in antioxidants level, carbohydrate metabolizing enzymes activity, peripheral tissues and DNA integrity. Expression of IL-4, IL-6 was found significantly higher in the stressed group. CUES initiates pathogenesis of diabetes.

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