ORAL GALACTOSE TREATMENT IMPROVES METABOLIC CHANGES IN A RAT MODEL OF SPORADIC ALZHEIMER'S DISEASE

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OBJECTIVES

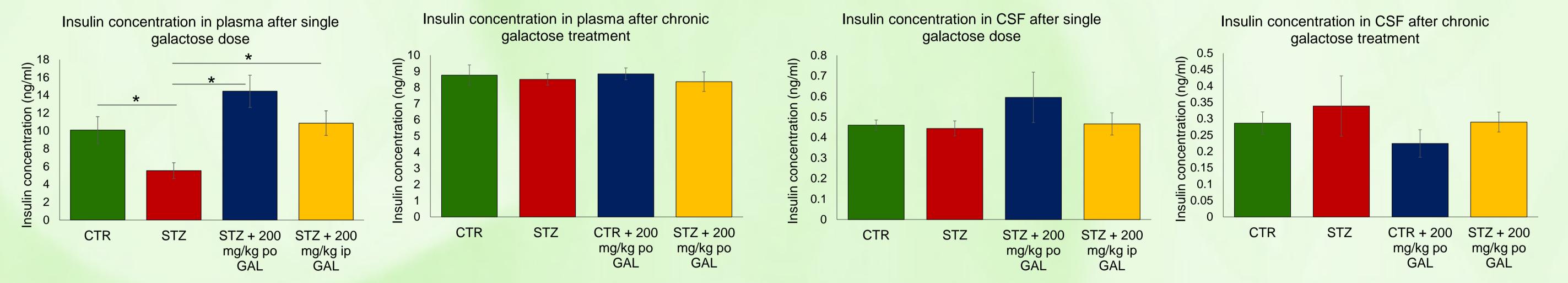
Chronic oral galactose treatment improves learning and memory functions in streptozotocin-induced (STZ-icv) rat model of sporadic Alzheimer's disease (sAD), contrary to its parenteral administration, as shown in our previous research. We aimed to explore the effects of long- and short-term oral galactose treatment on metabolic changes in STZ-icv rat model.

MATERIALS & METHODS

Adult male Wistar rats were given STZ-icv (3 mg/kg) while controls received vehicle only. Oral galactose treatment (200 mg/kg) was initiated 1 month after icv injections and continued for 2 months on daily basis until sacrifice (long-term). In a single-dose experiment animals were sacrificed 15 min after the oral or intraperitoneal galactose treatment (200 mg/kg), performed 1 month after icv injection. Insulin and glucagon-like peptide 1 (GLP-1, active/total) were measured by ELISA in plasma and cerebrospinal fluid (CSF). Data were analyzed by Kruskal-Wallis and Mann-Whitney U-test (p<0.05).

RESULTS

Decreased plasma insulin levels found after 1 month in STZ-icv rats were normalized to the control level after a single oral galactose dose. Unchanged CSF insulin levels in STZ-icv rats were found regardless the galactose treatment.



Plasma GLP-1 activity was unchanged in STZ group after one month and decreased 3 months after icv treatment. Single oral galactose dose tended to increase plasma GLP-1 activity while chronic treatment normalized GLP-1 activity in STZ group.

GLP-1 concentration in plasma after single galactose Active GLP-1 concentration in plasma after single GLP-1 ratio in plasma after single galactose dose dose galactose dose p=0.0833 concentration (pM) 30 25 1.8 p=0.0703 concentration (pM) 1.6 25 p=0.0703 p=0.0851 20 1.4 20 1.2 15 Ratio 15 0.8 10 0.6 10 0.4 GLP-1 5 GLP-1 5 0.2 CTR STZ STZ + 200 STZ + 200 CTR STZ STZ + 200 STZ + 200 CTR STZ STZ + 200 STZ + 200 mg/kg po GAL mg/kg ip GAL mg/kg po GAL mg/kg ip GAL mg/kg po GAL mg/kg ip GAL Active GLP-1 concentration in plasma after chronic GLP-1 concentration in plasma after chronic galactose GLP-1 ratio in plasma after chronic galactose treatment galactose treatment treatment concentration (pM) 0.5 concentration (pM) 2 0.45 1.8 12 0.4 * 0.35 p=0.0703 0.3 1.2 Ratio 8 0.25 6 0.2 0.8 0.15 0.6 GLP-1 0.1 GLP-1 0.4 2 0.05 0.2 STZ CTR CTR + 200 STZ + 200 CTR STZ CTR + 200 STZ + 200 STZ CTR CTR + 200 STZ + 200 mg/kg po GAL mg/kg po GAL

CONCLUSION

ORAL GALACTOSE IMPROVES METABOLIC CHANGES INDUCED IN A NON-TRANSGENIC RAT MODEL OF sAD FOLLOWED UP TO 3 MONTHS POST STZ-ICV TREATMENT



