

Designing a late-stage type 2 diabetes mellitus model with brain insulin resistance and oxidative stress

Rocío Redondo-Castillejo^{1,2}; Marina Hernández-Martín^{1,2}; Luis García-García^{1,3}; Juana Benedí¹; Adrián Macho-González⁴; Francisco J. Sánchez-Muniz⁴; Sara Bastida⁴; Alba Garcimartín¹; M. Elvira López-Oliva²; Aránzazu Bocanegra¹.

¹ Department of Pharmacology, Pharmacognosy and Botany. Faculty of Pharmacy. Complutense University of Madrid. Madrid 28040. Spain.

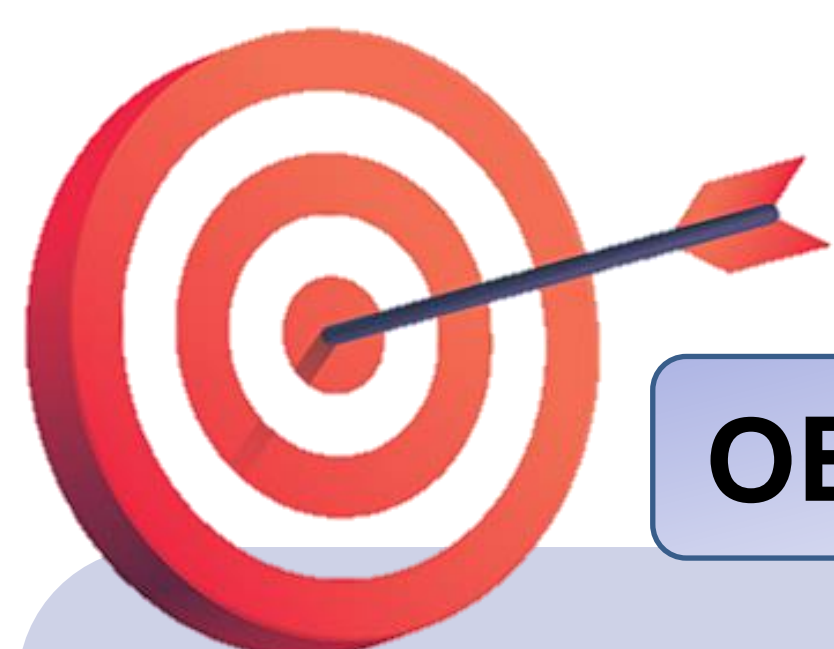
² Departmental Section of Physiology. Faculty of Pharmacy, Complutense University of Madrid. Madrid 28040. Spain.

³ Brain Mapping Unit, Pluridisciplinary Institute. Complutense University of Madrid. Madrid 28040. Spain.

⁴ Department of Nutrition and Food Science (Nutrition). Faculty of Pharmacy, Complutense University of Madrid. Madrid 28040. Spain.

INTRODUCTION

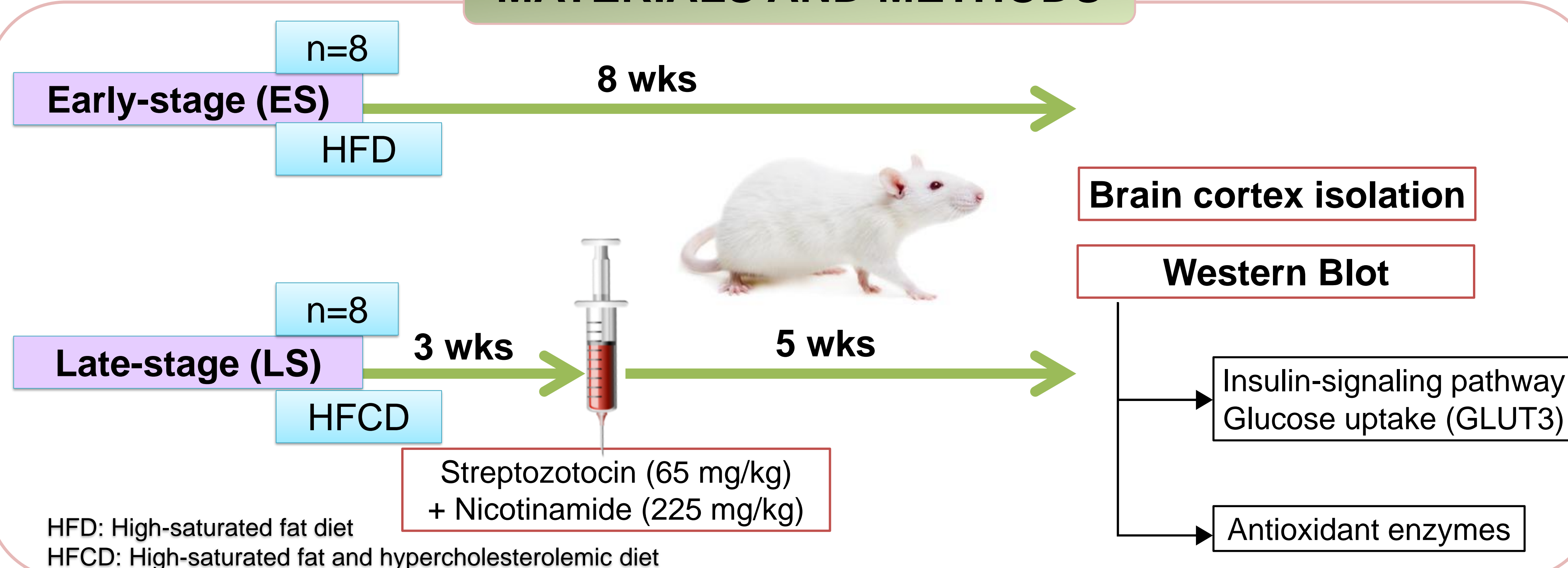
- ❖ The impairment in insulin-signaling pathway induces neurological damage [e.g., Alzheimer's disease (AD)]. When AD and type 2 diabetes mellitus (T2DM) coexists, it is known as type 3 diabetes mellitus (T3DM).
- ❖ The mitochondrial dysfunction promotes oxidative stress and plays an important role in brain insulin resistance instauration, which finally leads to T3DM.



OBJECTIVE

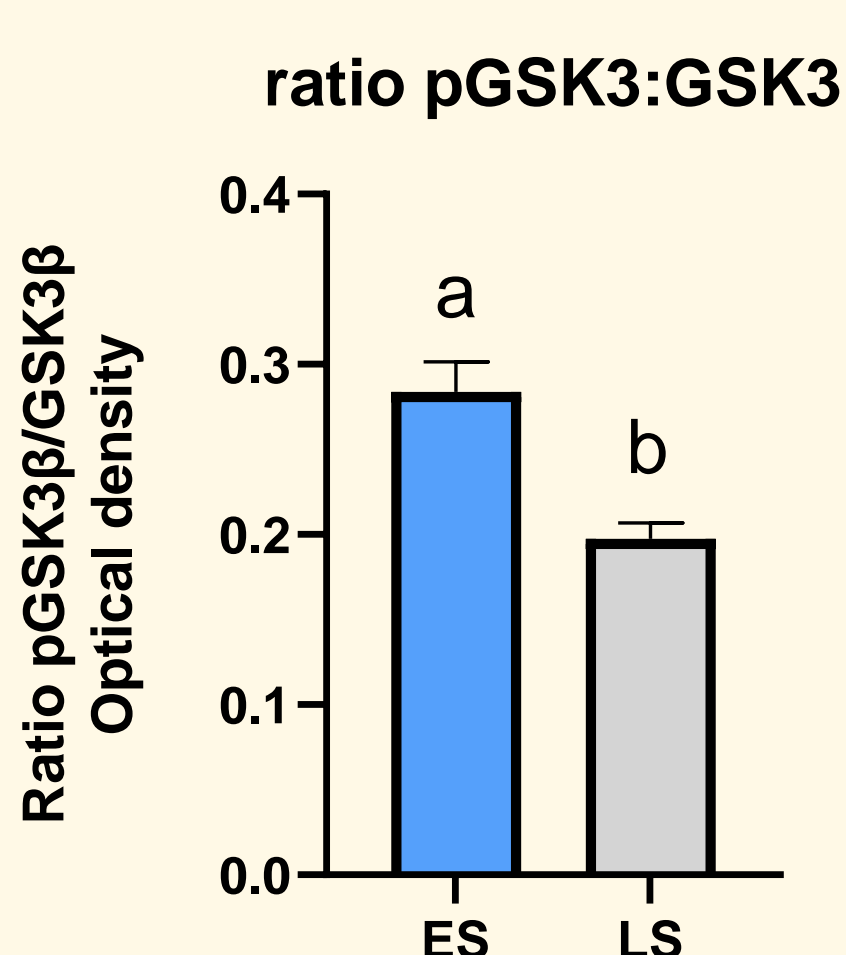
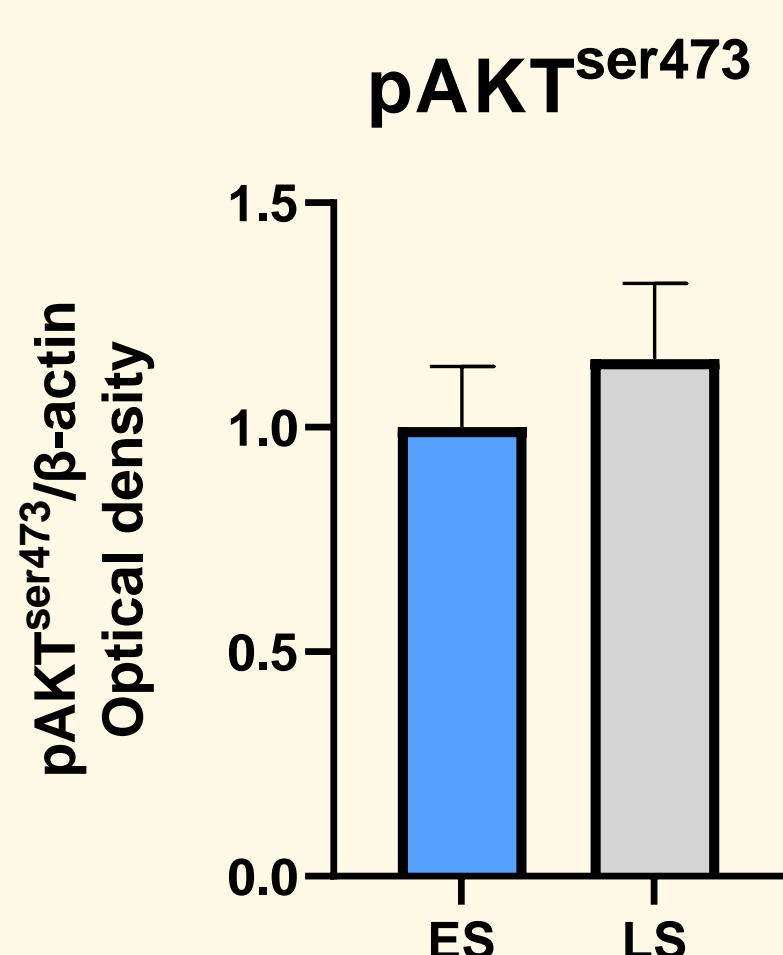
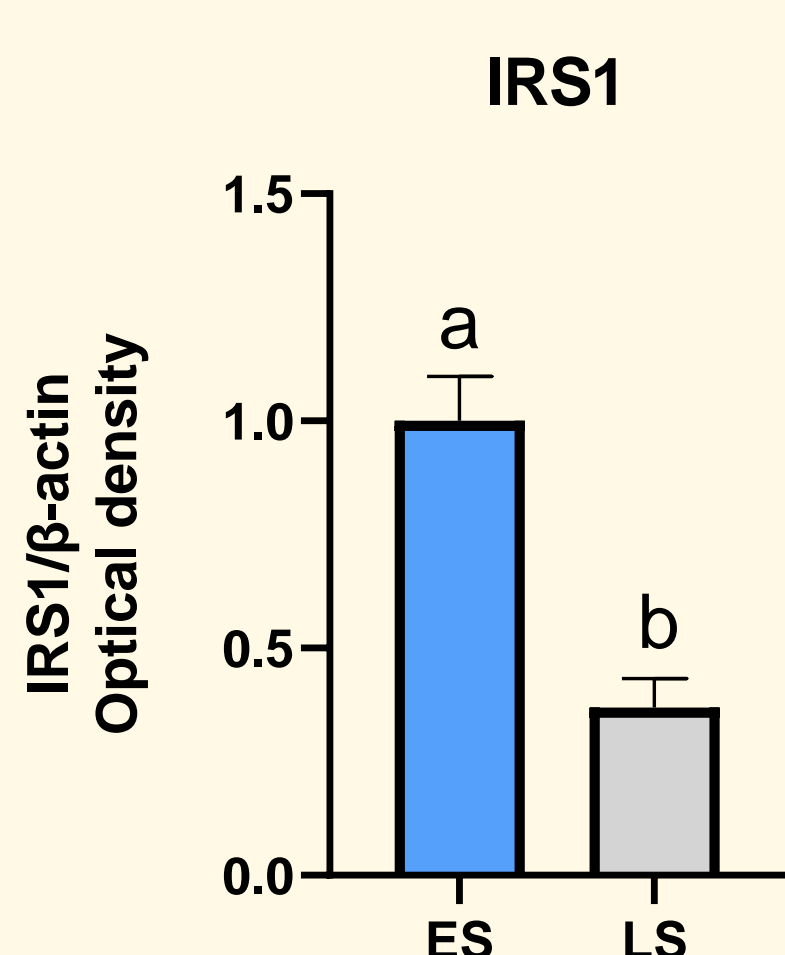
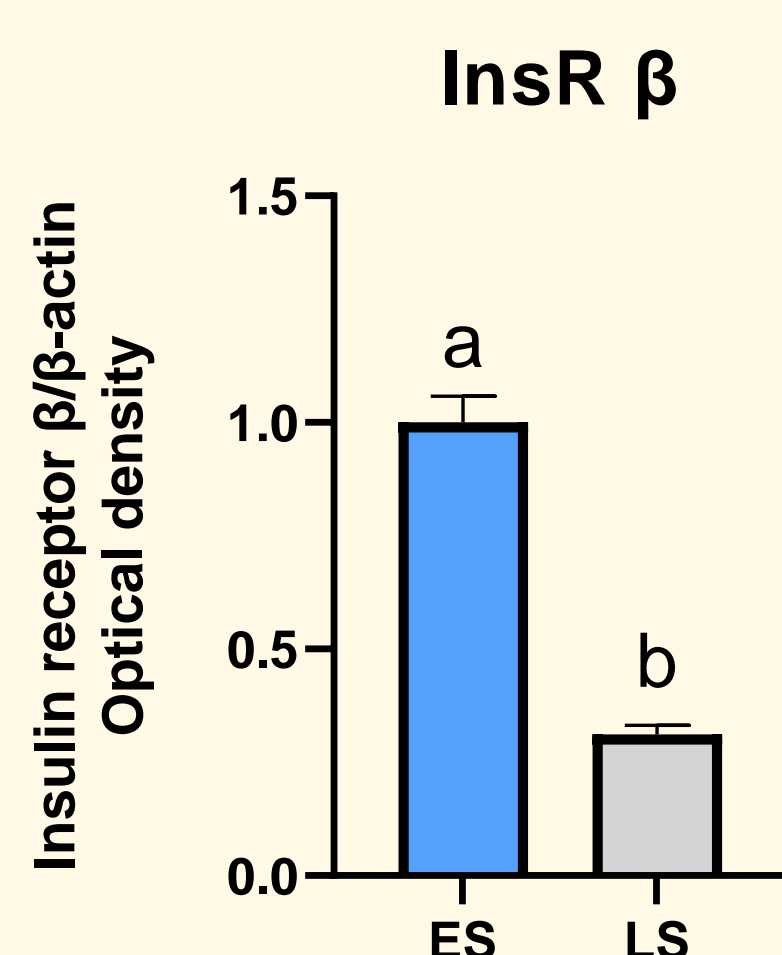
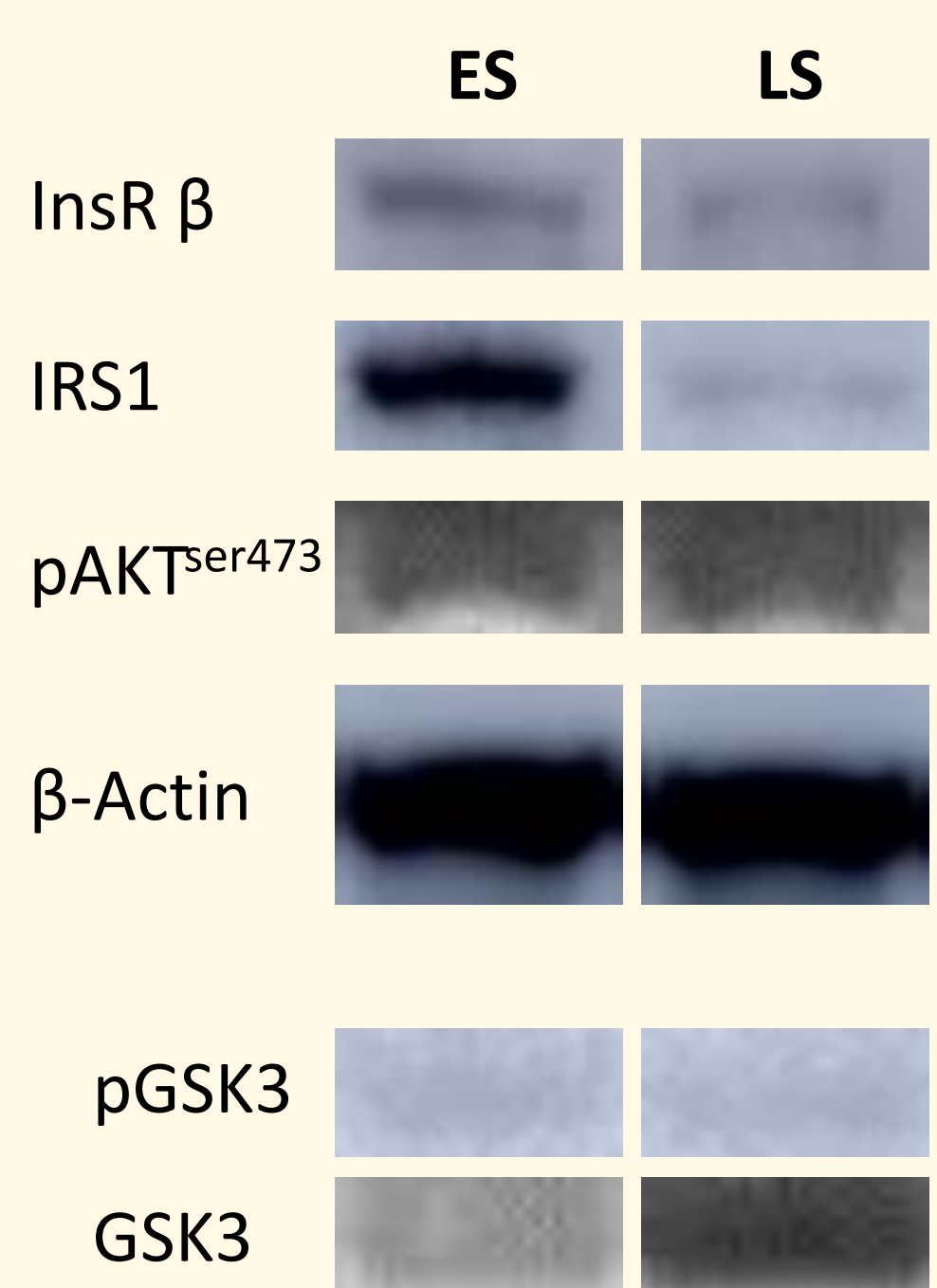
To reveal the suitability of a late-stage T2DM rat model induced by high-saturated fat and hypercholesterolemic diet and streptozotocin-nicotinamide (STZ-NAD) injection as a T3DM model.

MATERIALS AND METHODS

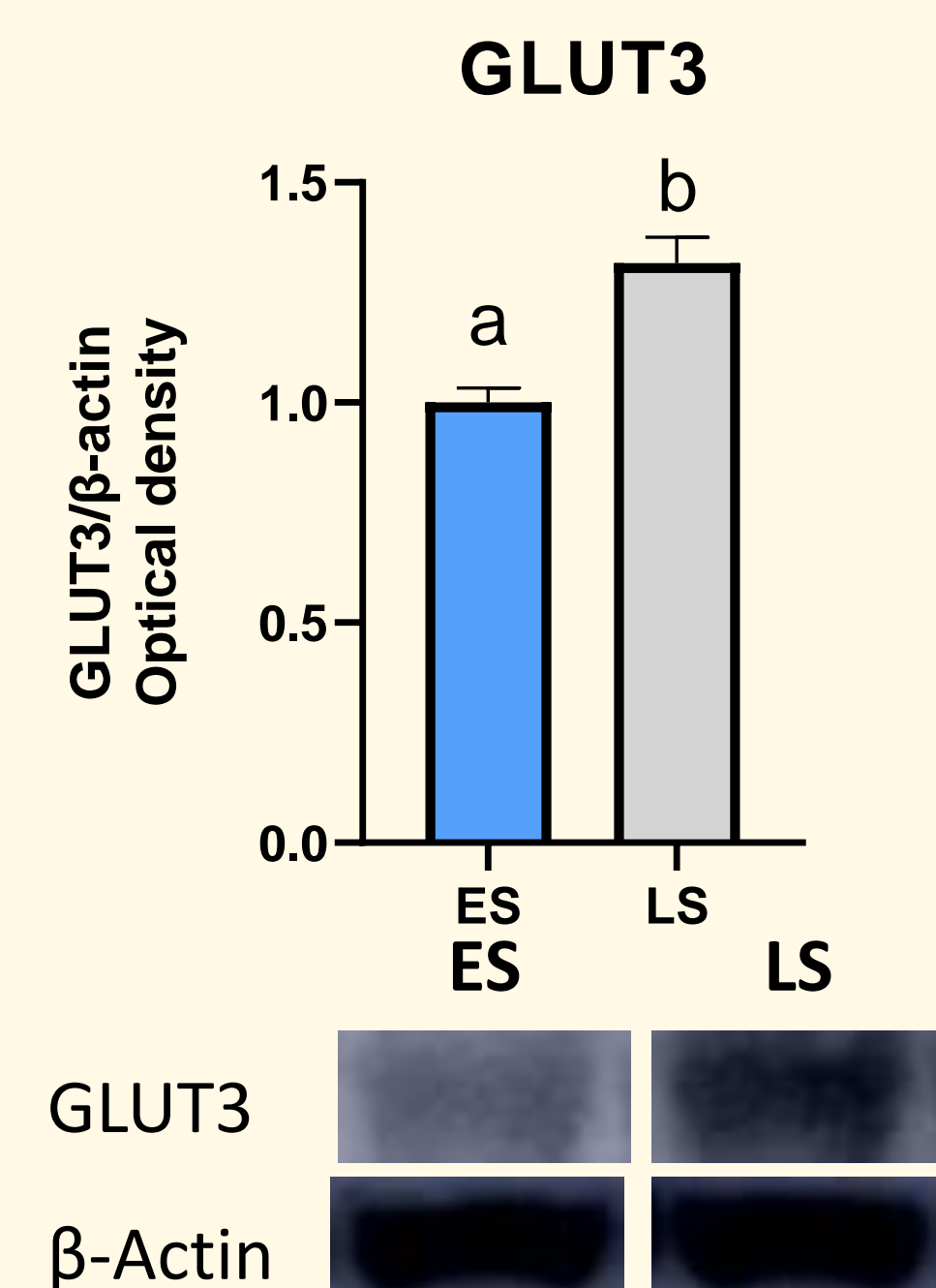


RESULTS

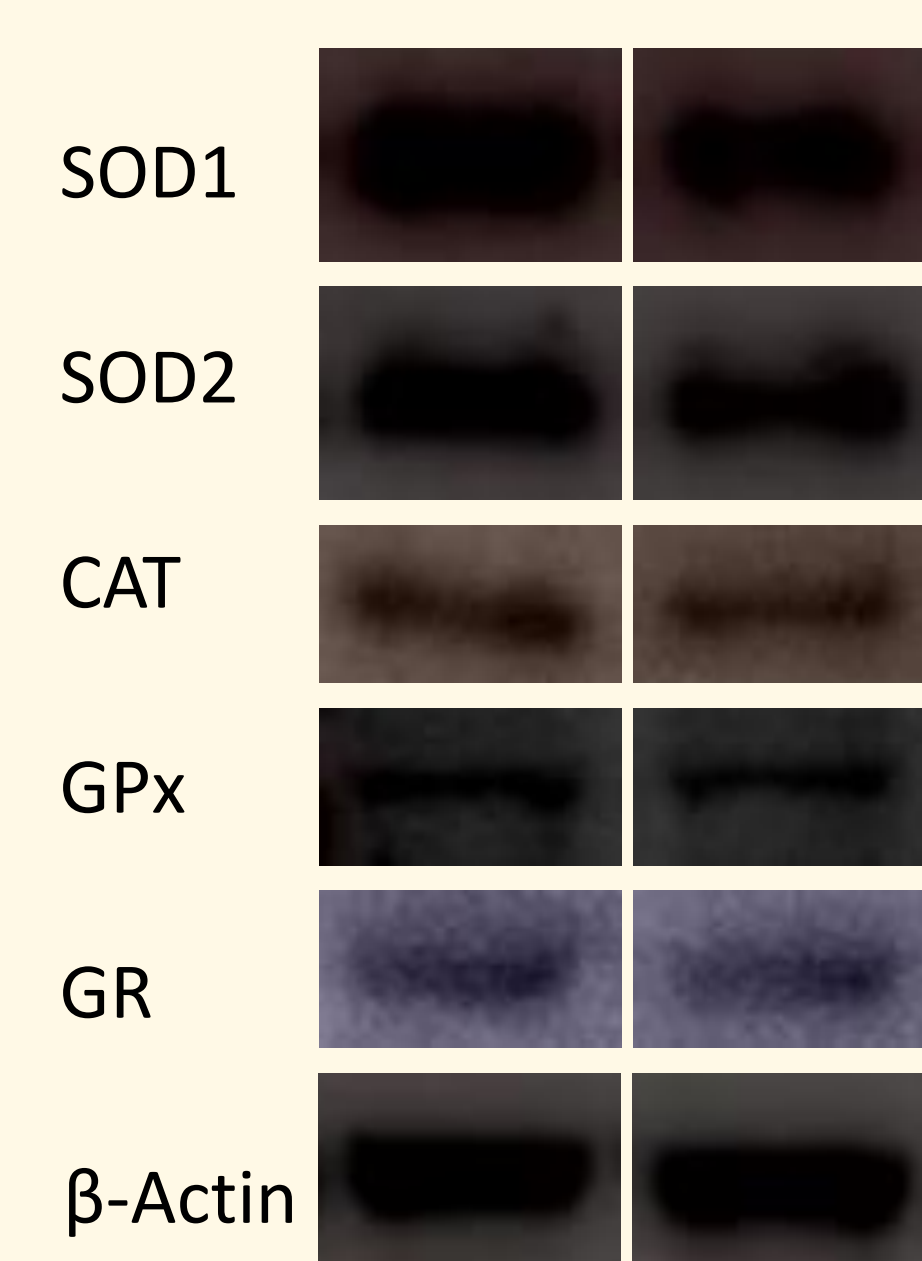
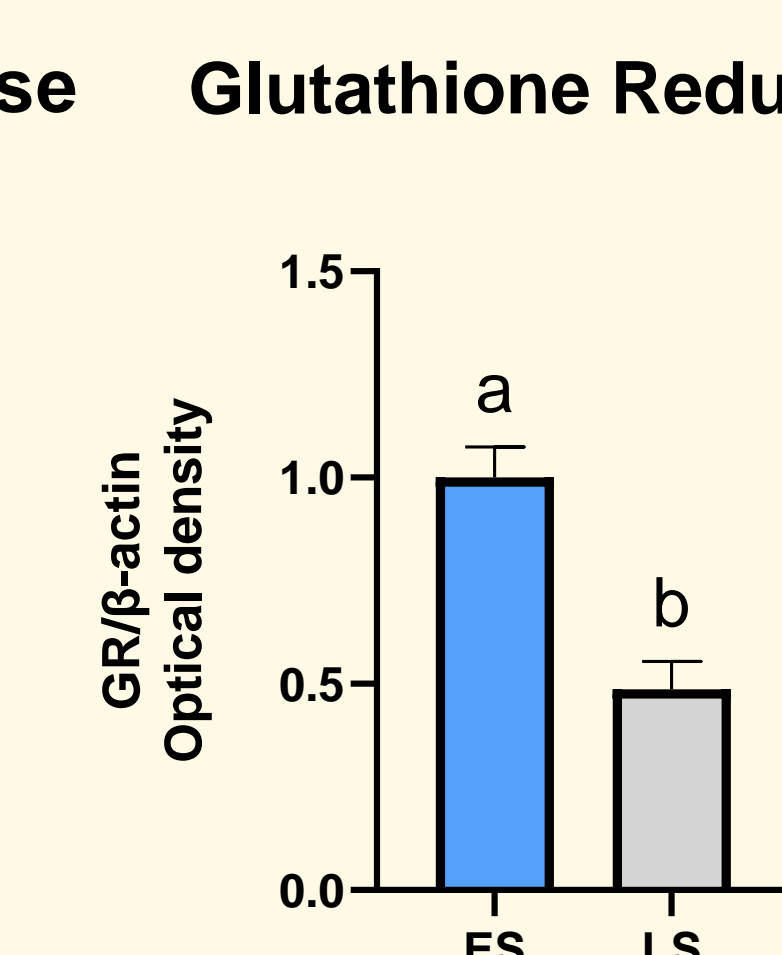
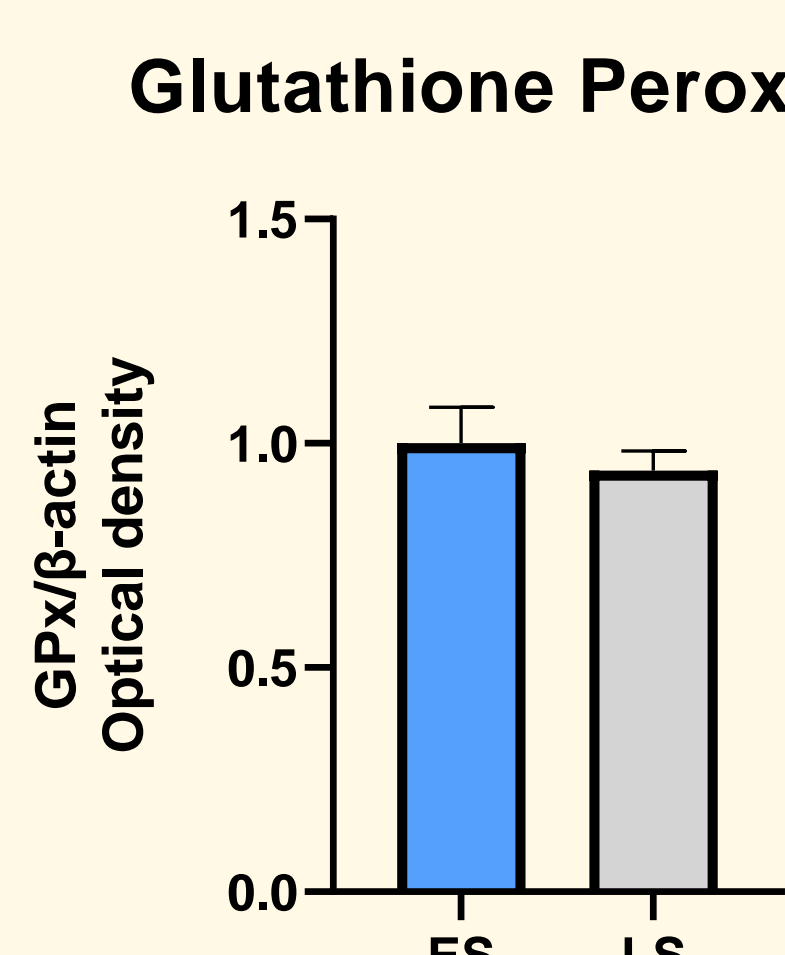
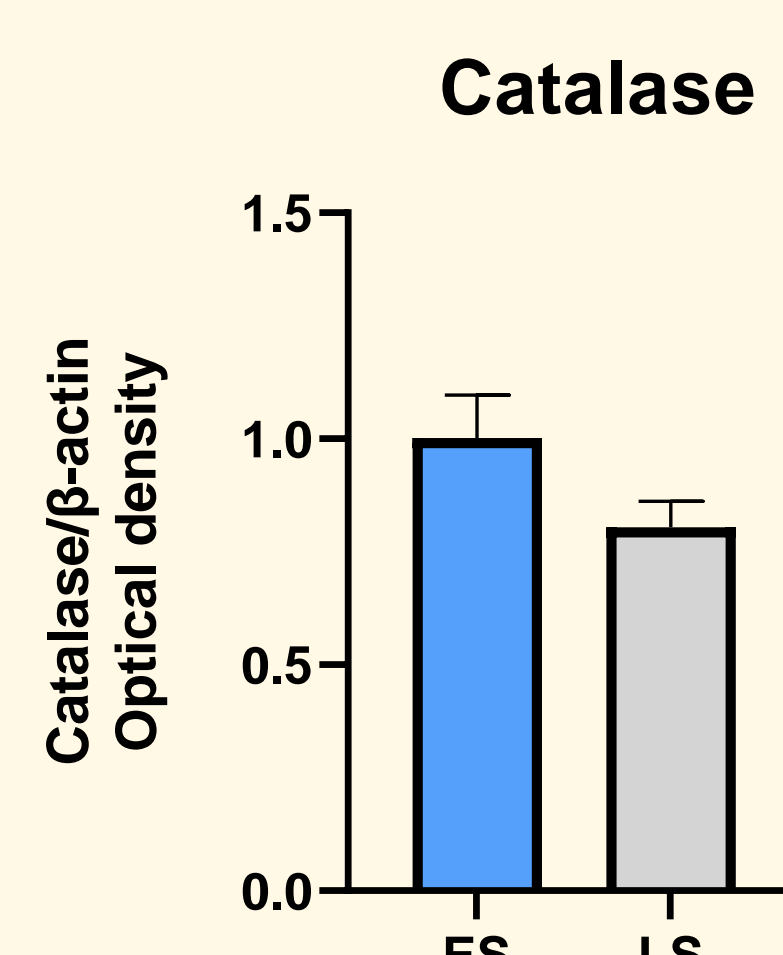
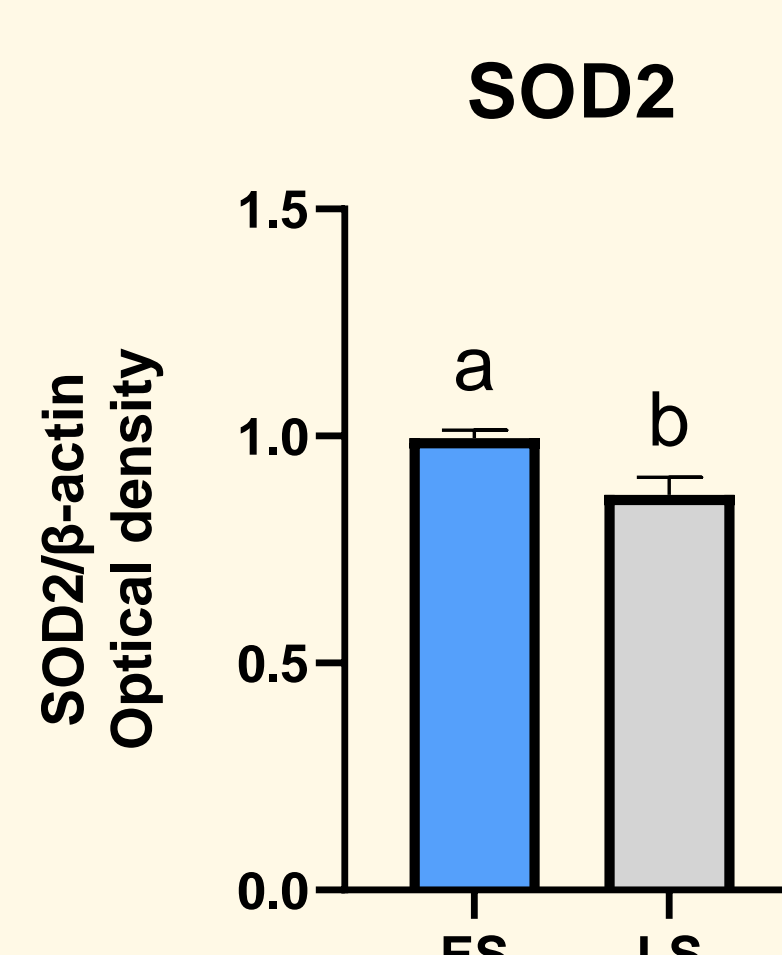
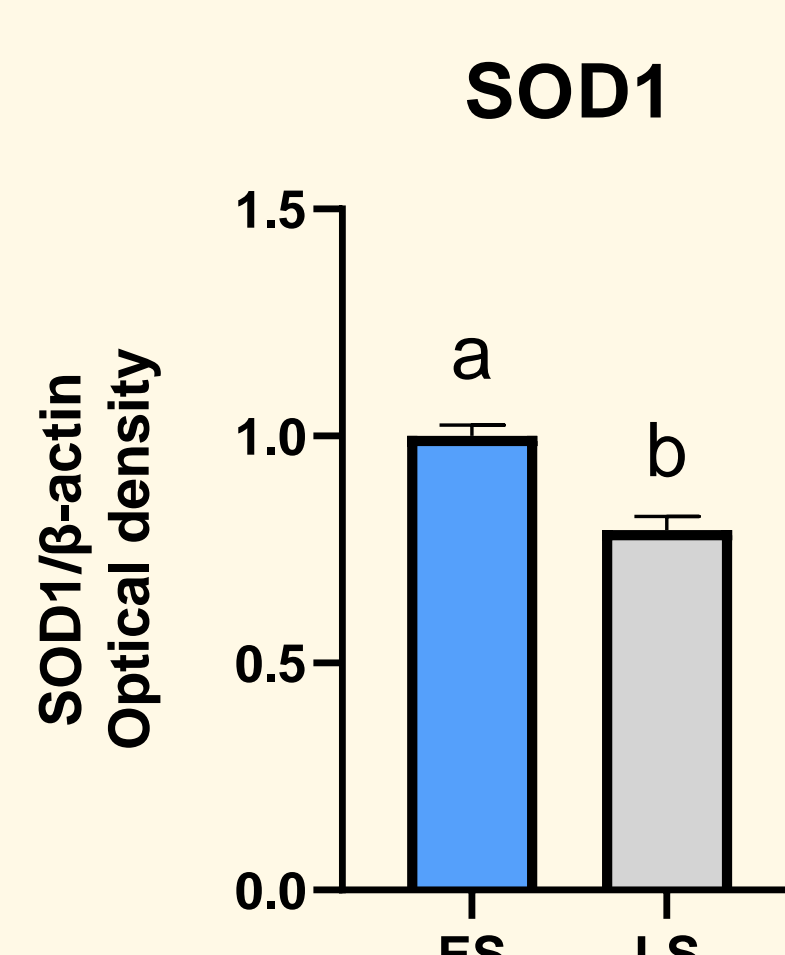
Insulin-signaling pathway



Glucose uptake



Antioxidant enzymes

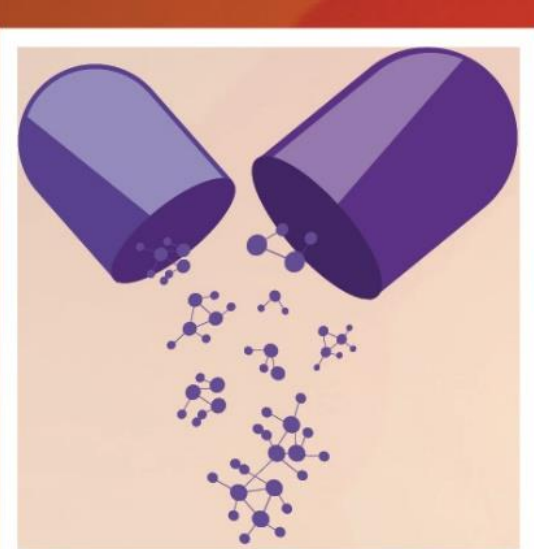


CONCLUSIONS

- ❖ The late-stage T2DM model induced by HFCD and a STZ-NAD injection represents an appropriate experimental tool to study the progression of brain insulin resistance in T3DM.
- ❖ This model could be useful to evaluate the efficacy of potential neuroprotective drugs.

ACKNOWLEDGEMENT

This study was supported by the Spanish Project PID2019-103872RB-I00.



The 7th International Electronic Conference on Medicinal Chemistry

01-30 NOVEMBER 2021 | ONLINE