

Effects of steviol on cytotoxicity, adipogenesis, ROS concentration and gene expression in 3T3-L1 Cell Line

Jakub Kurek¹, Joanna Zielińska-Wasielica², Katarzyna Kowalska², Anna Olejnik², Zbigniew Krejpcio¹

¹Institute of Human Nutrition and Dietetics, Poznań University of Life Sciences, Poznań, Poland

²Department of Biotechnology and Food Microbiology, Poznań University of Life Sciences, Poznań, Poland

The presented work is an integral part of the research project (National Science Centre, Poland, NCN 2017/27/B/NZ29/00677)



Background

The increasing prevalence of **diabetes mellitus has become one of the world's most serious public health problems**. This reflects an increase in associated risk factors such as being overweight or obese. Currently available therapies are associated with a number of adverse effects, so there is a need to search for new substances that can support the treatment.

Stevia rebaudiana Bertoni is a plant famous for its sweetness due to the content of steviol glycosides that are derivatives of diterpene – steviol. Stevia and its glycosides are becoming popular in the world of science due to their **anti-diabetic potential**. However, the mechanisms responsible for its action are not fully understood and require further investigation.

Study aim

The aim of this study was to evaluate the effects of steviol on **adipogenesis, intracellular ROS generation and the gene expression, as well as cytotoxicity in murine 3T3-L1 cell line**.

Methods

Cell viability, adipocyte differentiation, intracellular ROS generation and the influence on gene expression in the presence of steviol (in concentrations of 1 – 100 μM) was determined using MTT assay, Oil Red O staining, NBT methods and RT-PCR. Murine cell line 3T3-L1 was obtained from the American Type Culture Collection (ATCC® CL-173™, USA).

Results

It was found that **steviol in concentrations up to 100 μM did not affect proliferation** of 3T3-L1 cells. Also, there were **no effects on lipid accumulation and intracellular ROS generation** with examined concentrations (0 - 100 μM).

However, the **agent affected the expression of genes** regulating adipogenesis and lipogenesis processes:

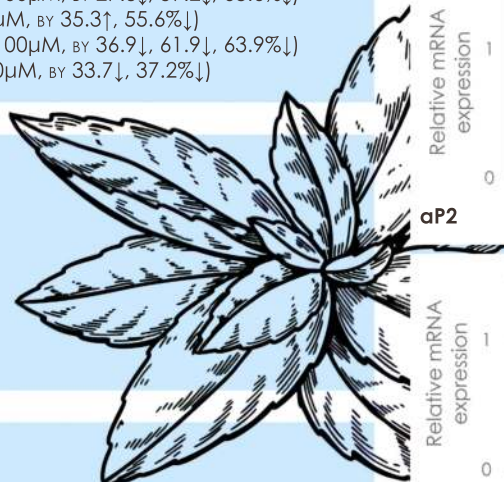
PPAR γ	(AT CONCENTRATIONS OF 10 AND 100μM, BY 38.1↓, 45.3%↓)
C/EBP α	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 36.9↓, 61.9↓, 45.3%↓)
SREBP-1	(AT CONCENTRATIONS OF 10 AND 100μM, BY 49.8↓, 37.2%↓)
LPL	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 45↓, 33.7↓, 30.8%↓)
AP2	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 36.9↓, 23.8↓, 52.4%↓)
FAS	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 55.4↓, 22.4↓, 55.6%↓)
LEP	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 27.5↓, 59.2↓, 68.6%↓)
ADIPOQ	(AT CONCENTRATIONS OF 1 AND 100μM, BY 35.3↑, 55.6%↓)
RSTN	(AT CONCENTRATIONS OF 1, 10 AND 100μM, BY 36.9↓, 61.9↓, 63.9%↓)
GLUT4	(AT CONCENTRATIONS OF 10 AND 100μM, BY 33.7↓, 37.2%↓)

Conclusion

The results of this study seem to confirm that **the mechanisms of anti-diabetic effects of steviol may be related to inhibition of expression of genes responsible for adipogenesis, lipogenesis and glucose transport in cells**.

References

- Belloir, C., Neiers, F., & Briand, L. (2017). Sweeteners and sweetness enhancers. *Current Opinion in Clinical Nutrition and Metabolic Care*, 20(4), 279–285.
- Bugaj, B., Leszczyńska, T., Pysz, M. et al. (2013). Charakterystyka i prozdrowotne właściwości *Stevia rebaudiana* Bertoni. *ŻYWNOSC. Nauka. Technologia. Jakość*, 3(88), 27–38.
- Gupta, E., Purwar, S., Sundaram, S., & Rai, G. K. (2013). Nutritional and therapeutic values of *Stevia rebaudiana*: A review. *Journal of Medicinal Plants Research*, 7(46), 3343–3353.
- Kroyer, G. (2010). Steviol and Stevia-sweetener in food: application, stability and interaction with food ingredients. *Journal für Verbraucherschutz und Lebensmittelsicherheit*, 5(2): 225–229.
- Perrier, J. D., Mihalov, J. J., & Carlson, S. J. (2018). FDA regulatory approach to steviol glycosides. *Food and Chemical Toxicology*, 4.6 Safety.



Graphs

