Diabetes, β cell apoptosis and the role of SIRT2 and Rel/A NFkB subunit

Abdemalek Rezgui 1 (PhD c), Recha Amel Tachour 2(PhD c), Rania Derguine 3(PhD c), Anfel Benmesseur 2(PhD c), Fatma Zahra Hab 2(PhD c) , Rym Agred 1 (PhD), Hakim Rahmoune 4 (MD), Nada Boutrid 4. (MD, PhD), Nadia Baiben 4 (MD c), Amira Baiben 4 (MD c), Widad Sobhi 1(PhD)

1: Laboratory of Plant Biotechnology and Ethnobotany, Faculty of Nature and Life Sciences, University of Bejaia, 06000, Bejaia, Algeria

2: Research Center of Biotechnology (CRBT), Ali Mendjli, 25000 Constantine, Algeria

3: Laboratory of Applied Biochemistry, Faculty of Nature & Life Sciences, Ferhat Abbas Setif 1 University, Algeria

4: Faculty of Medicine, University of Setif-1, Algeria

INTRODUCTION



Streptozotocin is a chemical compound once used as an antibiotic, but its toxicity for the β cells of the islets of Langerhans has dramatically limited its use.

This molecule is currently used in an experimental context for the study of type 1 diabetes, due to the latter adverse effects on β cells. In fact, the experimental model of STZ-induced diabetes helps to understand the mechanisms involved in β -cell death.

We performed this murine study to examine new mechanisms by which STZ induces β -cells apoptosis. Two cellular factors SIRT₂ and NF-kB were specifically targeted to unveil their potential roles in β -cell survival.

MATERIAL & METHODS

A culture of a NOD mouse in β -cell line, NIT₁, was exposed to STZ. In the cell lysate, we determined the inhibition of the two factors studied: SIRT₂ and NF-kB.

RESULTS & DISCUSSION

Streptozotocin (STZ) is a glucosamine-nitrosourea compound derived from Streptomyces achromogenes, which is used clinically as a chemotherapeutic

agent in the treatment of pancreatic β cell carcinoma. STZ-induced diabetic models are widely used in research for assessing the pathological

consequences of diabetes and screening potential therapies for the treatment of this condition

In our murine model, STZ did induce β -cell death with an IC 50 value = 4 μ g/mL. In addition, we confirmed that STZ inhibits the phosphorylation of the RelA p65, a NF-kB subunit, leading to the apoptotic process. Also, STZ does significantly inhibit SIRT₂.

CONCLUSION

STZ induces the death of β -cell trough the inhibition of SIRT₂ and NF-KBp65. It is not yet clear whether the two inhibitions are related or not.

Further studies are needed to reveal this. These results would also contribute to better understanding of the early stages of β-cell damages

and its role in the involving type 1 diabetes.