

Effects of EnduBerry™ Nu on Presenilin 2 and FOXP1 Gene Expression



S-980

Test institute: QIMA, 86160 Gençay, France

Sponsor: Mibelle AG Biochemistry, 5033 Buchs, Switzerland

INTRODUCTION

In this study, the effect of EnduBerry™ Nu on the expression of genes related mitochondrial function and cell protection (FOXP1, Presenilin 2) was investigated in aged human dermal fibroblasts isolated and compared to the corresponding untreated cells. FOXP1 is associated with increased proliferation in cardiomyocytes (1) and has been shown to suppress vascular inflammation (2). Further expression of FOXP1 has been shown to decrease with age in mesenchymal stem cells. It is thought that by controlling cellular senescence, FOXP1 can influence tendon and skeletal aging (3, 4). Therefore, increased FOXP1 may indicate a positive effect on cell fitness during aging. Presenilin 2 is a protease involved in the control of mitochondrial function. Dysfunction of presenilin 2 is associated disrupted mitochondrial bioenergetics and cellular oxidative stress (5, 6).

STUDY DESIGN

Cell cultivation and treatment

Normal human dermal fibroblasts (NHDF) were isolated from old donors. Prior to experimental start, the cells were grown for 24 hours in culture medium (DMEM + 10 % serum) and for another 24 hours in assay medium (DMEM + 1 % serum). Then the cells were treated with or without (control) 0.1 % EnduBerry™ Nu for 24 hours. After exposure, cells were harvested in phosphate buffered saline (PBS) and frozen at -80 °C. All conditions were performed in n=4.

Gene expression analysis

Replicates were pooled (yielding a final n=2) and total RNA extracted from each condition using TriPure™ Isolation Reagent (Roche) according to the supplier's instructions. The cDNA was synthesized by reverse transcription (RT) using a standard RT kit with oligo(dT) primers (Roche). Finally, the samples were amplified by quantitative real-time PCR using the LightCycler system (Roche Molecular System Inc.), using SYBR Green I reagent and primers specifically targeting FOXP1 and Presenilin 2.

RESULTS

Treatment with EnduBerry™ Nu at 0.1 % resulted in an increase of FOXP1 expression in aged fibroblasts (Figure 1). As FOXP1 is involved in cardiomyocyte proliferation and suppression of vascular inflammation, this may indicate a vasculoprotective effect of EnduBerry™ Nu.

Further, EnduBerry™ Nu led to increased Presenilin 2 expression, which may indicate a beneficial effect on mitochondrial function and cellular energy metabolism (Figure 1).

The information contained in this publication is provided in good faith and is based on our current knowledge. No legally binding promise or warranty regarding the suitability of our products for any specific use is made. These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure or prevent any disease. Mibelle AG Biochemistry will not assume any expressed or implied liability in connection with any use of this information. No part of this publication may be reproduced in any matter without the prior written permission of Mibelle AG Biochemistry.

Together the results indicate that EnduBerry™ Nu may promote cellular fitness by improving mitochondrial function and proliferation. This may translate to increased tissue energy levels, which can be beneficial in boosting overall physical performance.

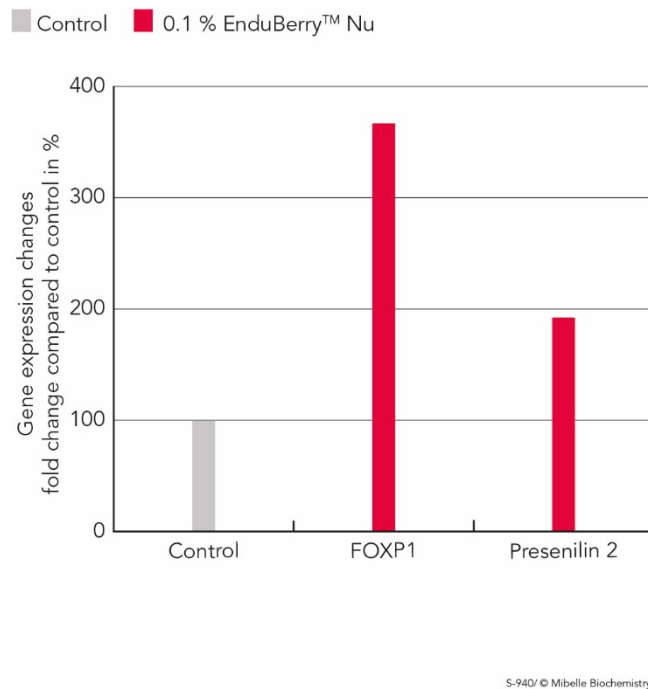


Figure 1. Increased expression of FOXP1 and Presenilin 2

REFERENCES

- 1) Liu, XM., Du, SL., Miao, R. et al. "Targeting the forkhead box protein P1 pathway as a novel therapeutic approach for cardiovascular diseases." *Heart Fail Rev* (2022) 27: 345–355.
- 2) Lim, G.B. "FOXP1 suppresses the endothelial NLRP3 inflammasome." *Nat Rev Cardiol* (2019) 16: 578–579.
- 3) Li, H., Liu, P., Xu, S. et al. "FOXP1 controls mesenchymal stem cell commitment and senescence during skeletal aging." *J Clin Invest.* (2017) 127(4): 1241-1253
- 4) Xu, H. & Liu, F. "Downregulation of FOXP1 correlates with tendon stem/progenitor cells aging." *Biochem Biophys Res Comm* (2018) 1(26): 96-102.
- 5) Filadi R., Greotti, E., Turacchio, G. et al. "Presenilin 2 modulates endoplasmic reticulum-mitochondria coupling by tuning the antagonistic effect of mitofusin 2" *Cell Reports* (2016) 15: 2226–2238
- 6) Contino, S., Porporato, PE., Bird, M. "Presenilin 2-dependent maintenance of mitochondrial oxidative capacity and morphology." *Front Physiol* (2017):

The information contained in this publication is provided in good faith and is based on our current knowledge. No legally binding promise or warranty regarding the suitability of our products for any specific use is made. These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure or prevent any disease. Mibelle AG Biochemistry will not assume any expressed or implied liability in connection with any use of this information. No part of this publication may be reproduced in any matter without the prior written permission of Mibelle AG Biochemistry.