

## 001. Cardioprotective effect of liraglutide is amplified with anti-inflammatory and decreased brain natriuretic peptide levels, in addition to glycemia and body weight reduction.

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Piljac A<sup>1</sup>, Jazbec A<sup>2</sup>, Duvnjak L<sup>1</sup>, Ljubic S<sup>1</sup>

1. Clinical Hospital Merkur, University Clinic Vuk Vrhovac, Zagreb, Croatia; 2. University of Zagreb, Zagreb, Croatia

### Purpose

Besides an impact on glycemic control and body weight, incretins emerged as important factors in cardiovascular (CV) protection in diabetes. Dipeptidyl peptidase-4 (DPP-4) inhibitors cleave multiple peptides, which in turn have direct effect on the heart and blood vessels. This distinguishes them in action when compared to GLP-1 agonists. The aim was to compare the impact of DPP-4 inhibitors GLP-1 agonist liraglutide on CV risk factors.

### Methods

A total of 442 type 2 diabetics were studied during a 6-month period and assigned into three study groups treated with DPP-4 inhibitors: linagliptin (Group [Gr] 1; n=158), vildagliptin (Gr 2; n=150) and with GLP-1 agonist liraglutide (Gr 3; n=134). Adiponectin (ApN), brain natriuretic peptide (BNP), high specific C-reactive protein (hsCRP), blood pressure (BP), glycated hemoglobin (HbA1c) and other CVR factors were determined at the beginning and at the end of the follow-up period. Differences for the analyzed variables between baseline values and values after 6 months were tested by t paired test.

### Results

Hs-CRP mean values at the beginning of the study were  $3.86 \pm 3.64$ ,  $2.67 \pm 2.52$  and  $5.31 \pm 2.37$  in the Gr 1, 2, 3 respectively, and were significantly reduced by 0.63 (95% CI: 0.1-1.15;  $p=0.018$ ), 1.35 (95% CI: -0.26-2.97;  $p=0.09$ ) and 1.71 (95% CI: 0.57-2.84;  $p=0.007$ ) on average in all three groups, with greater reduction in Gr 3 in comparison with Gr 2. HbA1c mean values at the beginning of the study were  $8.01 \pm 0.79$ ,  $7.36 \pm 0.87$  and  $8.01 \pm 0.95$  in the Gr 1, 2, 3 respectively, and were significantly reduced by 0.94 (95% CI: 0.73-1.15;  $p<0.01$ ), 0.69 (95% CI: 0.05-1.32;  $p=0.04$ ) and 1.15 (95% CI: 0.35-1.95;  $p<0.01$ ) on average, with no difference in reduction between groups. BNP and body mass index (BMI) were significantly reduced from baseline ( $30.5 \pm 14.6$  and  $39.3 \pm 4.5$ ) by 10.7 (95% CI: 4.73-16.61;  $p=0.002$ ) and 2.65 (95% CI: 1.35-3.94;  $p<0.01$ ) on average in Gr 3, whereas reduction in SBP was significant from baseline ( $137.5 \pm 16.9$ ) in Gr 2 by 9.0 (95% CI: -0.05-18.55). Postprandial C-peptide, gamma-glutamyl transpeptidase (GGT) and triglycerides were reduced in Gr 3 by -0.32 (95% CI: -0.65-0.01;  $p=0.058$ ), 8.42 (95% CI: -0.15-16.9;  $p=0.053$ ) and 0.67 (95% CI: -0.09-1.43;  $p=0.079$ ) on average but these reductions were not significant. Increase in amylase was not observed in studied groups.

### Conclusions

Except HbA1c and BMI reduction liraglutide proved more efficient in hs-CRP and BNP reduction in comparison with DPP-4 inhibitors. Treatment with liraglutide may exert cardioprotective benefits not only due to its glycemic control and body weight reduction but also through its pleiotropic effect.