

ESM Tab. 2. Clinical Study 1, details of type 2 diabetics subgroups

| | T2DM-N (N=39) | T2DM-NI (N=39) | T2DM-U (N=16) | p-value |
|---------------------------------------|--------------------------------|---------------------------------|--------------------------------|----------------|
| Age (years)* | 60.4 ± 7.9 | 68.3 ± 7.1 | 61.1 ± 10.0 | <0.001 |
| Gender (males)* | 30(77) | 34(87) | 12(75) | 0.417 |
| Smoking (n, %)** | 0(0) | 1(3) | 1(6) | 0.335 |
| Hypertension (n, %)** | 35(90) | 39(100) | 4(25) | <0.001 |
| Body Mass Index (kg/m ²)* | 30.6 ± 5.8 | 29.9 ± 5.1 | 30.7 ± 5.9 | 0.837 |
| Duration of diabetes (years)* | 12.3±12.5 | 15.9±9.9 | 7.4±9.5 | 0.009 |
| Laboratory test* | | | | |
| HbA _{1c} (%) (mmol/mol) | 8.1±1.8 (65±19.7) | 7.9±1.7 (63±18.6) | 6.8±1.2 (51±13.1) | 0.039 |
| Fasting glucose (mmol/l) | 8.7±2.8 | 8.2±2.9 | 7.9±3.2 | 0.485 |
| Creatinine (µmol/l) | 96.8±61.6 | 114.4±44 | 70.4±17.6 | <0.001 |
| Total cholesterol (mmol/l) | 4.45±1.13 | 4.29± 1.07 | 4.3±1.05 | 0.794 |
| LDL-cholesterol (mmol/l) | 2.45±0.92 | 2.39±1.03 | 2.67±0.81 | 0.707 |
| HDL-cholesterol (mmol/l) | 1.44±0.56 | 1.34±0.5 | 1.27±0.57 | 0.541 |
| Triglycerides (mmol/l) | 1.28±0.51 | 1.63± 0.72 | 1.31± 0.71 | 0.052 |
| Medications** | | | | |
| Insulin | 14(36) | 24(62) | 1(6) | 0.001 |
| Oral anti-diabetic drugs | 4(10) | 1(3) | 12(75) | <0.001 |
| Incretins | 4(10) | 0(0) | 0(0) | 0.053 |
| Statins | 19(49) | 27(69) | 8(50) | 0.150 |
| Anti-platelet therapy | 17(44) | 39(100) | 6(38) | <0.001 |

| Complications** | | | | |
|------------------------|--------|--------|------|--------|
| Retinopathy | 19(53) | 29(83) | 0(0) | <0.001 |
| Non recent MI/ictus | 13(33) | 29(76) | 1(6) | <0.001 |
| Nephropathy | 20(54) | 37(95) | 0(0) | <0.001 |

Distribution of clinical variables, laboratory tests, therapies and complications in type 2 diabetic patients affected by peripheral neuropathy (T2DM-N), or peripheral neuropathy and CLI (T2DM-NI) and uncomplicated patients (T2DM-U) enrolled in Study 1.

*Data presented as mean±SD; **Data presented as number and percentage.