

ONLINE APPENDIX TABLE 1: DIABHYCAR COHORT – STRATIFICATION OF INCIDENT CASES OF CAD ACCORDING TO CAD BASELINE STATUS

BASELINE	FOLLOW-UP								Total
	Non CAD	CABG	MI	SD	CABG + MI	CABG + SD	MI + SD	CABG + MI + SD	
Non CAD	2321	165	31	85	30	4	5	2	2643
Angina Pectoris	224	43	7	21	9	2	1	0	307
MI	66	20	2	9	3	2	1	0	103
Angina Pectoris + MI	47	13	4	4	0	1	0	0	69
Total	2658	241	44	119	42	9	7	2	3122

CAD: Coronary Artery Disease; CABG: Coronary Artery Bypass Graft; MI: Myocardial Infarction; SD: Sudden Death

ONLINE APPENDIX TABLE 2: DIABHYCAR COHORT – CHARACTERISTICS OF SUBJECTS AT BASELINE ACCORDING TO GENOTYPE

	UCP2 -866G>A			p
	GG	GA	AA	
Patients (n)	1275	1427	420	-
Male sex: (%)	72%	74%	72%	0.49
Age (years)	66 ± 8	66 ± 8	66 ± 9	0.83
Age at diagnosis of diabetes (years)	55 ± 9	55 ± 10	56 ± 10	0.67
Known duration of diabetes (years)	11 ± 8	10 ± 8	10 ± 8	0.15
Fasting plasma glucose (mM)	9.5 ± 3.0	9.6 ± 3.0	9.3 ± 3.3	0.20
HbA1c (%)	7.9 ± 1.8	7.8 ± 1.7	7.8 ± 1.8	0.29
BMI (kg/m ²)	29.3 ± 4.6	29.5 ± 4.7	29.4 ± 4.4	0.51
Ponderal Status: Lean / Overweight / Obese (%) *	12% / 46% / 32%	13% / 43% / 44%	12% / 42% / 45%	0.55
Triglycerides (mM)	2.23 ± 1.44	2.21 ± 1.41	2.21 ± 1.38	0.90
Total cholesterol (mM)	5.82 ± 1.03	5.77 ± 1.10	5.82 ± 1.10	0.28
LDL cholesterol (mM) †	3.57 ± 0.87	3.50 ± 0.89	3.50 ± 0.88	0.06 ‡
HDL cholesterol (mM)	1.31 ± 0.35	1.32 ± 0.36	1.32 ± 0.33	0.84

Creatinine clearance (ml/min)	85 ± 30	86 ± 28	86 ± 33	0.52
Urinary albumin excretion (mg/l)	234 ± 594	229 ± 689	217 ± 463	0.91
U.A.E.: µAlbuminuria / macroalbuminuria (%) §	76% / 24%	78% / 23%	78% / 23%	0.93
Arterial Hypertension (%)	56%	56%	58%	0.72

Data expressed as mean ± SD;. Statistics of quantitative parameters are Students t tests performed with log-transformed data.

* Ponderal Status. Lean: BMI <25 kg/m²; Overweight: 25 kg/m² ≤ BMI <30 kg/m²; Obese: BMI ≥30 kg/m²

† Data available for 1010 GG, 1153 GA and 335 AA carriers

‡ p=0.02 for a dominant model (AX vs GG carriers)

§ Urinary Albumin Excretion. µalbuminuria: 20 mg/l ≤ UAE <200 mg/l; macroalbuminuria: UAE ≥200 mg/l

|| Arterial hypertension: systolic blood pressure (SBP) >140 mmHg and/or diastolic blood pressure (DBP) >90 mmHg; or SBP and DBP below these values in the presence of antihypertensive medication and history of hypertension.

ONLINE APPENDIX TABLE 3. NCH COHORT - CHARACTERISTICS OF TYPE 2 DIABETIC MEN ACCORDING TO CORONARY ARTERY DISEASE STATUS

	With CAD	Without CAD	p
Patients (n)	174	161	-
Age (years)	66 ± 11	60 ± 10	<0.0001
Age at diagnosis of diabetes (years)	52 ± 12	50 ± 11	0.08
Known duration of diabetes (years)	14 ± 10	10 ± 9	0.006
Fasting plasma glucose (mM)	9.1 ± 4.2	9.0 ± 3.1	0.94
Hemoglobin A1c (%)	8.1 ± 1.8	7.9 ± 1.9	0.26
Treatment (Diet / OHA / Insulin)	15% / 50% / 35%	15% / 64% / 21%	0.01
BMI (kg/m ²)	28.3 ± 4.5	28.7 ± 4.6	0.41
Ponderal Status: Lean / Overweight / Obese (%) *	23% / 44% / 33%	23% / 43% / 34%	0.96
Triglycerides ≥1.70 mM (%) †	52%	55%	0.90
LDL cholesterol ≥3.35 mM (%) †	66%	52%	0.004
HDL cholesterol ≤1.02 mM (%) †	38%	27%	0.001
Creatinine clearance (ml/min)	73 ± 33	88 ± 32	<0.0001

U.A.E.: Normal / μ Albuminuria / Proteinuria (%) ‡	49% / 31% / 20%	65% / 22% / 13%	0.01
Arterial Hypertension (%) §	87%	70%	0.0002
Tobacco smoking (%)	69%	57%	0.04

Data expressed as mean \pm SD. Statistics of quantitative parameters are Students t tests performed with log-transformed data.

OHA: oral hypoglycemic agents

* Ponderal Status. Lean: BMI <25 kg/m²; Overweight: 25 kg/m² \leq BMI <30 kg/m²; Obese: BMI \geq 30 kg/m²

† present or pre-treatment (statins, fibrates) values: age-adjusted comparisons

‡ Urinary Albumin Excretion. Normal: UAE <20 mg/24h; μ albuminuria: 20 mg/24h \leq UAE <300 mg/24h; proteinuria: UAE \geq 300 mg/24h

§ Arterial hypertension: systolic blood pressure (SBP) >140 mmHg and/or diastolic blood pressure (DBP) >90 mmHg; or SBP and DBP below these values in the presence of antihypertensive medication and history of hypertension.

|| Present or past tobacco smoking