

**Table 2:** Changes in metabolic profile, liver and muscle enzymes of 648 study participants from baseline to last visit [Median follow-up: 7 years; 607 on a statin alone and 41 on a statin + chronic PPI. Parametric and non-parametric values are expressed as mean  $\pm$  standard deviation and median (interquartile range), respectively, unless percentages are shown].

		Baseline visit	Recent visit	P vs. baseline	Change, %
Glucose (fasting), mg/dL	Statin	95 (88-105)	97 (90-109)	<0.001	+2.1%
	Statin + PPI	97 (84-106)	96 (91-109)	0.183	-1.0%
BMI, kg/m <sup>2</sup>	Statin	27.5 $\pm$ 3.5	28.5 $\pm$ 3.8	<0.001	+3.6%
	Statin + PPI	27.6 $\pm$ 3.8	28.3 $\pm$ 4.9	0.173	+2.5%
Uric acid, mg/dL	Statin	5.1 $\pm$ 1.5	5.3 $\pm$ 1.5	0.002	+3.9%
	Statin + PPI	4.8 $\pm$ 1.5	5.2 $\pm$ 1.5	0.142	+8.3%
SBP, mmHg	Statin	140 (128-155)	129 (120-136)	<0.001	-7.8%
	Statin + PPI	148 (130-159)	132 (121-138)	<0.001	-10.8%
DBP, mmHg	Statin	88 (80-95)	79 (73-84)	<0.001	-10.2%
	Statin + PPI	90 (78-92)	74 (70-80)*	<0.001	-17.7%‡
eGFR, mL/min/1.73 m <sup>2</sup>	Statin	79 $\pm$ 14	74 $\pm$ 16	<0.001	-6.3%
	Statin + PPI	74 $\pm$ 12	67 $\pm$ 16	0.007	-9.4%
TCHOL, mg/dL	Statin	265 $\pm$ 46	174 $\pm$ 31	<0.001	-34.3%
	Statin + PPI	259 $\pm$ 43	162 $\pm$ 27*	<0.001	-37.5%‡
TG, md/dL	Statin	135 (97-190)	111 (85-148)	<0.001	-17.8%
	Statin + PPI	138 (101-169)	108 (93-134)	0.002	-21.7%
HDL-C, mg/dL	Statin	54 $\pm$ 14	55 $\pm$ 14	0.002	+1.9%
	Statin + PPI	59 $\pm$ 17	57 $\pm$ 15	0.568	-3.4%
LDL-C, mg/dL	Statin	181 $\pm$ 41	95 $\pm$ 25	<0.001	-47.5%
	Statin + PPI	172 $\pm$ 39	82 $\pm$ 25*	<0.001	-52.3%‡
AST, IU/L	Statin	21 (18-25)	23 (20-27)	<0.001	+9.5%
	Statin + PPI	21 (19-25)	22 (18-30)	0.260	+4.8%
ALT, IU/L	Statin	21 (17-28)	23 (18-29)	0.036	+9.5%
	Statin + PPI	20 (16-24)	19 (14-28)	0.993	-5.0%
$\gamma$ -GT, IU/L	Statin	18 (13-27)	19 (14-26)	0.227	+5.5%
	Statin + PPI	17 (11-26)	17 (12-38)	0.927	0%
ALP, IU/L	Statin	71 (57-94)	58 (47-73)	<0.001	-18.3%
	Statin + PPI	80 (58-111)	59 (43-75)	<0.001	-26.2%
CK, IU/L	Statin	94 (72-130)	105 (78-155)	<0.001	+11.7%
	Statin + PPI	111 (74-143)	111 (82-179)	0.178	0%

PPI: proton pump inhibitors, BMI: body mass index, SBP: systolic blood pressure, DBP: diastolic blood pressure, eGFR: estimated glomerular filtration rate, TCHOL: total cholesterol, TG: triglycerides, HDL-C: high-density lipoprotein cholesterol, LDL-C: low-density lipoprotein cholesterol, AST: aspartate aminotransferase, ALT: alanine aminotransferase,  $\gamma$ GT: gamma glutamyltranspetidase, ALP: alkaline phosphatase, CK: creatine phosphokinase, \*: p <0.05 after adjusting for baseline values. To convert from mg/dL to mmol/L multiply by 0.02586 for TCHOL, HDL-C and LDL-C, by 0.01129 for TG and by 0.06 for glucose.