

## S6 Supporting Information. Repeated measures analysis.

9/16/2018

### Repeated measures analysis

Gender, body mass index (BMI), age, and amount of smoking at randomization were analyzed in terms of their association with the level of the biomarkers and the longitudinal stability of these biomarkers. The repeated measures of log-transformed biomarkers during weeks 4-20 were analyzed for each factor (e.g., gender) using a linear mixed model with a fixed effect of the studied factor and a random intercept. The p-value of the fixed effect of the studied factor is reported.

Variable	Factor	Creatinine corrected	Non corrected	TNE corrected	Creatinine and TNE corrected
		Estimated fixed effect (95% CI), p-value	Estimated fixed effect (95% CI), p-value	Estimated fixed effect (95% CI), p-value	Estimated fixed effect (95% CI), p-value
Log(8-isoPGF <sub>2a</sub> )	Female	-0.01 (-0.13, 0.10) p = 0.81	-0.27 (-0.42, -0.12) <b>p &lt; 0.001</b>	-0.06 (-0.25, 0.14) p = 0.56	0.20 (-0.02, 0.42) p = 0.07
	Age (×10 years)	0.03 (-0.02, 0.07) p = 0.24	-0.10 (-0.16, -0.05) <b>p &lt; 0.001</b>	-0.12 (-0.19, -0.05) <b>p &lt; 0.001</b>	0.01 (-0.08, 0.09) p = 0.89
	BMI (kg/m <sup>2</sup> )	0.00 (-0.01, 0.01) p = 0.51	0.01 (-0.01, 0.02) p = 0.30	0.01 (-0.00, 0.03) p = 0.051	0.01 (-0.01, 0.03) p = 0.20
	CPD (×10)	0.16 (0.03, 0.30) <b>p = 0.02</b>	0.02 (-0.17, 0.20) p = 0.87	-0.59 (-0.81, -0.37) <b>p &lt; 0.0001</b>	-0.44 (-0.70, 0.19) <b>p &lt; 0.001</b>
Log(PGEM)	Female	-0.35 (-0.52, -0.18) <b>p &lt; 0.0001</b>	-0.62 (-0.82, -0.41) <b>p &lt; 0.0001</b>	-0.39 (-0.63, -0.14) <b>p &lt; 0.01</b>	-0.12 (-0.38, 0.13) p = 0.35
	Age (×10 years)	0.09 (0.02, 0.15) <b>p &lt; 0.01</b>	-0.04 (-0.13, 0.04) p = 0.31	-0.06 (-0.15, 0.03) p = 0.20	0.07 (-0.03, 0.16) p = 0.16
	BMI (kg/m <sup>2</sup> )	-0.01 (-0.02, 0.01) p = 0.31	-0.00 (-0.02, 0.01) p = 0.71	0.00 (-0.01, 0.02) p = 0.62	0.00 (-0.02, 0.02) p = 0.92
	CPD (×10)	0.04 (-0.16, 0.25) p = 0.69	-0.09 (-0.36, 0.17) p = 0.50	-0.71 (-1.00, -0.43) <b>p &lt; 0.0001</b>	-0.58 (-0.88, -0.28) <b>p &lt; 0.001</b>