

Supplementary Table 2. Statistical Power Related to Individual Primary Endpoints at the Planned Sample Size.

Based on a 5/8 Success Criterion. The planned sample size accounted for an anticipated 75% of the THS arm being predominant ($\geq 70\%$) THS product users. It was sufficient to attain an overall statistical power of more than 99% to show at least 5 out of 8 statistically significant effects on the primary endpoints using the Hailperin-Rüger approach, with test-wise power indicated in right column.

Primary Endpoint	Assumed Preserved Effect of Cessation (%)	Assumed Reduction from Smoking	Variability	Test-wise Power (%)[*]
HDL-C	80	3.3 mg/dL	10.0 mg/dL	99.6
WBC	70	-0.6 $10^9/L$	1.6 $10^9/L$	98.5
FEV ₁ % pred	75	1.6 %pred	6.0 %pred	82.2
sICAM-1	60	12% ^{***}	23% ^{****}	99.9
11-DTX-B ₂	80	18% ^{***}	27% ^{****}	99.9
8-epi-PGF _{2α}	50	16% ^{***}	36% ^{****}	99.9
COHb	80	65% ^{***}	39% ^{****}	99.9
Total NNAL	80	70% ^{***}	60% ^{****}	99.9

^{*}Type I error of 0.03125 (0.05 *5/8)

^{**}Difference (THS minus Smoking)

^{***}Relative reduction (Smoking minus THS)/Smoking

^{****}Geometric coefficient of variation