S5 Table. Population attributable fractions of deaths prior to age 75 given the distribution of covariates in the EPIC cohort, using waist-to-hip ratio to assess for obesity.

		All participants		Never-smokers		Current-smokers	
	$Covariate^1$	Overall <sup>2</sup>	Cumulative <sup>3</sup>	Overall <sup>2</sup>	Cumulative <sup>3</sup>	Overall <sup>2</sup>	Cumulative <sup>3</sup>
Women and Men	Smoking	0.30 [0.30,0.31]	0.30 [0.30,0.31]	-	-	0.55 [0.54,0.56]	0.55 [0.54,0.56]
	Diet	0.13 [0.11, 0.16]	$0.40 \ [0.38, 0.41]$	0.12 [0.10, 0.14]	0.12 [0.10, 0.14]	$0.14 \ [0.12, 0.17]$	0.62 [0.61, 0.63]
	Overweight and obesity (WHR)	0.09 [0.08, 0.11]	0.46 [0.44, 0.47]	0.09 [0.08, 0.11]	$0.20 \ [0.18, 0.23]$	0.09 [0.07, 0.11]	0.66 [0.65, 0.67]
	High blood pressure	0.09 [0.07, 0.10]	$0.51 \ [0.49, 0.52]$	0.09 [0.07, 0.11]	$0.27 \ [0.25, 0.30]$	0.08 [0.06, 0.09]	0.69 [0.68, 0.70]
	Physical inactivity	0.07 [0.05, 0.09]	$0.54 \ [0.52, 0.56]$	0.08 [0.06, 0.10]	$0.33 \ [0.30, 0.36]$	$0.06 \ [0.04, 0.08]$	0.71 [0.70, 0.72]
	Alcohol intake	$0.03 \ [0.03, 0.04]$	$0.56 \ [0.54, 0.58]$	$0.02 \ [0.01, 0.02]$	$0.34 \ [0.31, 0.37]$	$0.05 \ [0.04, 0.05]$	0.73 [0.71, 0.74]
	Combined	$0.56 \ [0.54, 0.58]$		$0.34 \ [0.31, 0.37]$		0.73 [0.71, 0.74]	
Women	Smoking	0.26 [0.25,0.26]	0.26 [0.25,0.26]	-	-	0.55 [0.54,0.56]	0.55 [0.54,0.56]
	Diet	0.13 [0.10, 0.16]	0.35 [0.33, 0.38]	0.12 [0.09, 0.15]	0.12 [0.09, 0.15]	0.15 [0.12, 0.18]	0.62 [0.60, 0.64]
	Overweight and obesity (WHR)	0.07 [0.04, 0.09]	0.40 [0.37, 0.42]	0.07 [0.04,0.09]	0.18 [0.15, 0.21]	0.07 [0.04, 0.09]	0.65 [0.63, 0.67]
	High blood pressure	0.09 [0.07,0.11]	0.46 [0.43, 0.48]	0.10 [0.08, 0.12]	0.26 [0.23, 0.30]	0.08 [0.06, 0.10]	0.68 [0.66, 0.70]
	Physical inactivity	0.06 [0.03, 0.09]	0.49 [0.46, 0.52]	0.06 [0.03, 0.10]	0.31 [0.27, 0.35]	0.05 [0.02, 0.08]	0.70 [0.68, 0.72]
	Alcohol intake	0.02 [0.01, 0.02]	0.50 [0.47, 0.52]	0.01 [0.01, 0.02]	0.32 [0.28, 0.35]	0.02 [0.01, 0.03]	0.71 [0.69, 0.72]
	Combined	0.50 [0.47,0.52]		0.32 [0.28,0.35]		0.71 [0.69,0.72]	
Men	Smoking	0.35 [0.33,0.36]	0.35 [0.33,0.36]	-	_	0.54 [0.53,0.56]	0.54 [0.53,0.56]
	Diet	0.13 [0.09, 0.16]	0.43 [0.41, 0.46]	0.13 [0.09, 0.16]	0.13 [0.09, 0.16]	0.13 [0.09, 0.17]	$0.61 \ [0.59, 0.63]$
	Overweight and obesity (WHR)	0.12 [0.10, 0.15]	0.51 [0.48, 0.54]	0.11 [0.09, 0.14]	0.22 [0.18, 0.26]	0.12 [0.10, 0.14]	0.67 [0.64, 0.69]
	High blood pressure	0.08 [0.04,0.11]	$0.55 \ [0.52, 0.58]$	0.08 [0.04,0.11]	0.29 [0.24, 0.33]	0.06 [0.03, 0.10]	0.69 [0.67, 0.71]
	Physical inactivity	0.07 [0.04,0.09]	0.59 [0.55, 0.61]	0.07 [0.04,0.09]	0.34 [0.29, 0.38]	0.06 [0.04, 0.09]	0.72 [0.69, 0.74]
	Alcohol intake	0.05 [0.04, 0.07]	0.61 [0.58, 0.64]	0.03 [0.02, 0.05]	0.36 [0.31, 0.40]	0.07 [0.05, 0.08]	0.74 [0.72, 0.76]
	Combined	$0.61 \ [0.58, 0.64]$		0.36 [0.31,0.40]		0.74 [0.72, 0.76]	

<sup>1</sup> Attributable fractions were calculated based on the difference in expected cumulative risk given the observed covariate distributions in EPIC and the expected cumulative risk under the following scenarios. Smoking: A population of never smokers. Diet: A population of people in the healthy category. Blood pressure: A population of people with normal blood pressure. High alcohol intake: A population who drink at most 1-2 drinks per day. Physical Activity: A population of people in the active category. Overweight and obesity: A population of people with WHR below the lowest sex-specific quintile. These attributable risks thus represent a best case, in that they are calculated based on a hypothetical reference population with risk factors removed entirely.

<sup>2</sup> Estimated using predictions from a model mutually adjusted for all listed covariates as well as age at baseline. Attributable fractions are based on modifying one covariate at a time, with the distribution of the

remaining covariates left as observed in EPIC.

<sup>&</sup>lt;sup>3</sup> The cumulative attributable fraction after the sequential addition of each covariate.