

Table S1 The association gene expression levels of AHRR and EXOC3 and gas exchange

Outcome	mRNA expression*	Model1		Model2	
		β	95% CI	β	95% CI
DLCO	<i>AHRR</i>	1.96	-1.13;5.05	3.36	0.49;6.24
	<i>EXOC3</i>	4.77	-0.53;10.07	7.06	2.26;11.85
DLCO/VA	<i>AHRR</i>	0.22	-0.26;0.70	0.4	-0.70;0.87
	<i>EXOC3</i>	0.76	-0.06;1.58	1.05	0.28;1.82

*mRNA expression levels are used in the log scale.

Model 1: adjusted for age, sex, smoking status, white blood cell counts and batch effects.

Model2: adjusted for age, sex, smoking status, weight, height, white blood cell counts and batch effects.

Bold indicates statistically significant association.

Table S2 Sensitivity analyses of smoking probes

Lead association	Probe Chr	Gene	Conditioned for	Discovery cohort (model2)				
				B	SE	P-value	FDR	
cg05575921	5	AHRR	-	0.065	0.013	3.60e-07	0.047	
	6	-	cg06126421	0.046	0.011	1.72e-05		
	2	ALPG ^a	cg05951221	0.028	0.010	8.11e-03		
			cg06126421 + cg05951221	0.025	0.0097	1.06e-02		
cg05951221	2	ALPG^a	-	0.049	0.0096	4.37e-07	0.047	
	6	-	cg06126421	0.038	0.0086	8.97e-06		
	5	AHRR	cg05575921	0.022	0.0079	5.40e-03		
			cg06126421 + cg05575921	0.022	0.0078	5.28e-03		

^aClosest gene

Table S3 Sensitivity analysis in non-smokers

Lead association	Study sample	Discovery cohort (model2)				
		B	SE	P-value	FDR	
cg05575921	Total population (N=659)	0.065	0.013	3.60e-07	0.047	
	In non-smokers (N=207)	0.023	0.011	3.65e-02		
cg05951221	Total population (N=659)	0.049	0.0096	4.37e-07	0.047	
	In non-smokers (N=207)	0.037	0.015	1.6e-02		

Table S4 Sensitivity analysis with additional adjustment for FEV₁/FVC

Lead association	Probe Chr	Gene	Conditioned for	Discovery cohort (model2)			
				B	SE	P-value	FDR
cg05575921	5	AHRR	-	0.065	0.013	3.60e-07	0.047
			FEV ₁ /FVC	0.047	0.013	2.61e-04	
cg05951221	2	ALPG ^a	-	0.049	0.0096	4.37e-07	0.047
			FEV ₁ /FVC	0.039	0.0098	7.82e-05	

^aClosest gene