

Supplemental Material

Plasma microRNA Expression and Micronuclei Frequency in Workers Exposed to Polycyclic Aromatic Hydrocarbons

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Table S1. Partial correlation coefficients^a between 10 urinary OH-PAHs, Σ OH-PAHs, and plasma BPDE-Alb adducts in 391 healthy male coke oven workers.

PAH internal exposure biomarkers ^b	1-hydroxy-naphthalene	2-hydroxy-naphthalene	2-hydroxy-fluorene	9-hydroxy-fluorene	1-hydroxy-phenanthrene	2-hydroxy-phenanthrene	3-hydroxy-phenanthrene	4-hydroxy-phenanthrene	9-hydroxy-phenanthrene	1-hydroxy-pyrene	Σ OH-PAHs	BPDE-Alb	Σ Partial r^c
1-hydroxynaphthalene		0.834**	0.473**	0.332**	0.541**	0.596**	0.480**	0.189**	0.614**	0.686**	0.862**	0.156*	5.763
2-hydroxynaphthalene	0.834**		0.442**	0.255**	0.497**	0.477**	0.408**	0.140*	0.519**	0.597**	0.793**	0.107*	5.069
2-hydroxyfluorene	0.473**	0.442**		0.364**	0.263**	0.621**	0.575**	0.132*	0.498**	0.407**	0.579**	-0.004	4.350
9-hydroxyfluorene	0.332**	0.255**	0.364**		0.253**	0.267**	0.078	0.330**	0.366**	0.282**	0.485**	0.065	3.077
1-hydroxyphenanthrene	0.541**	0.497**	0.263**	0.253**		0.436**	0.213**	0.130*	0.564**	0.653**	0.695**	0.074	4.319
2-hydroxyphenanthrene	0.596**	0.477**	0.621**	0.267**	0.436**		0.718**	0.053	0.690**	0.644**	0.672**	0.057	5.231
3-hydroxyphenanthrene	0.480**	0.408**	0.575**	0.078	0.213**	0.718**		0.017	0.464**	0.404**	0.483**	0.068	3.908
4-hydroxyphenanthrene	0.189**	0.140*	0.132*	0.330**	0.130*	0.053	0.017		0.126*	0.179**	0.305**	0.092	1.693
9-hydroxyphenanthrene	0.614**	0.519**	0.498**	0.366**	0.564**	0.690**	0.464**	0.126*		0.673**	0.736**	0.108*	5.358
1-hydroxypyrene	0.686**	0.597**	0.407**	0.282**	0.653**	0.644**	0.404**	0.179**	0.673**		0.852**	0.246**	5.623
Σ OH-PAHs	0.862**	0.793**	0.579**	0.485**	0.695**	0.672**	0.483**	0.305**	0.736**	0.852**		0.201**	6.663
BPDE-Alb	0.156*	0.107*	-0.004	0.065	0.074	0.057	0.068	0.092	0.108*	0.246**	0.201**		1.170

^aPartial correlation analysis with adjustment for age, smoking status, pack-years of smoking, drinking status, working years, workplace, and BMI. ^bLn-transformed. ^cThe sum of the partial correlation coefficients for each PAH internal exposure biomarker. * $P < 0.05$; ** $P < 0.001$.

Table S2. LOQ and percentage of quantifiable samples of each biomarker.

PAH internal exposure biomarkers^a	LOQ	n	Q%
1-hydroxynaphthalene	0.9	364	99.73
2-hydroxynaphthalene	0.9	364	99.73
2-hydroxyfluorene	0.1	359	98.36
9-hydroxyfluorene	0.2	360	98.63
1-hydroxyphenanthrene	0.3	363	99.45
2-hydroxyphenanthrene	0.1	363	99.45
3-hydroxyphenanthrene	0.2	360	98.63
4-hydroxyphenanthrene	0.1	356	97.53
9-hydroxyphenanthrene	0.2	362	99.18
1-hydroxypyrene	0.5	365	100
6-hydroxychrysene	1.4	0	0
3-hydroxybenzo[a]pyrene	1.0	0	0
BPDE-Alb	1.0	343	93.97

^aThe unit for OH-PAHs is $\mu\text{mol}/\text{mmol}$ creatine, and the unit for BPDE-Alb adducts is ng/mg albumin.

Table S3. Associations between PAH exposure levels and MN frequency (as the dependent variable) in the validation stage.

PAH internal exposure biomarkers^a	FR (95% CI)	P^b
1-hydroxynaphthalene	1.035 (0.976, 1.099)	0.253
2-hydroxynaphthalene	1.022 (0.959, 1.090)	0.500
2-hydroxyfluorene	1.060 (1.001, 1.125)	0.049
9-hydroxyfluorene	1.064 (1.005, 1.128)	0.034
1-hydroxyphenanthrene	1.036 (0.977, 1.100)	0.237
2-hydroxyphenanthrene	1.033 (0.977, 1.094)	0.258
3-hydroxyphenanthrene	1.020 (0.965, 1.078)	0.491
4-hydroxyphenanthrene	1.085 (1.025, 1.150)	0.006
9-hydroxyphenanthrene	1.020 (0.965, 1.078)	0.491
1-hydroxypyrene	1.051 (0.992, 1.113)	0.091
ΣOH-PAHs	1.072 (1.012, 1.135)	0.017
BPDE-Alb	1.134 (1.078, 1.192)	8.41×10 ⁻⁷

Abbreviations: FR: frequency ratio.

^aLn-transformed. ^bPoisson regression analysis with adjustment for age, smoking status, pack-years of smoking, drinking status, working years, workplace, and BMI.

Table S4. Associations of the expression levels of five PAH-associated miRNAs (as the dependent variable) with smoking status, age, and drinking status in the validation stage [standardized β (95% CI)].

miRNA ^a	Drinking status	<i>P</i> ^b	Smoking status	<i>P</i> ^c	Age	<i>P</i> ^d
miR-24-3p	-0.273 (-1.043, 0.497)	0.486	0.429 (-0.407, 1.266)	0.313	-0.223 (-1.175, 0.709)	0.627
miR-27a-3p	-0.236 (-0.803, 0.330)	0.412	0.500 (-0.120, 1.119)	0.114	-0.162 (-0.855, 0.531)	0.646
miR-142-5p	-0.393 (-1.009, 0.224)	0.211	0.579 (-0.094, 1.252)	0.091	0.129 (-0.627, 0.886)	0.737
miR-28-5p	0.000 (-0.634, 0.633)	1.000	0.162 (-0.532, 0.855)	0.647	-0.049 (-0.824, 0.726)	0.900
miR-150-5p	0.224 (-0.147, 0.595)	0.236	0.037 (-0.368, 0.441)	0.859	-0.051 (-0.504, 0.401)	0.824

^aLog₂-transformed. ^bMultivariable linear regression analysis with adjustment for age, smoking status, pack-years of smoking, working years, workplace, BMI, Σ OH-PAHs, and BPDE-Alb adducts. ^cMultivariable linear regression analysis with adjustment for age, drinking status, working years, workplace, BMI, Σ OH-PAHs, and BPDE-Alb adducts. ^dMultivariable linear regression analysis with adjustment for smoking status, pack-years of smoking, drinking status, working years, workplace, BMI, Σ OH-PAHs, and BPDE-Alb adducts.

Table S5. Associations between miRNA expression and MN frequency (as the dependent variable) in workers with different smoking status or age groups [FR (95% CI)].

miRNAs ^a	Nonsmokers (n=113)	<i>P</i> ^b	Smokers (n=252)	<i>P</i> ^b	<i>P</i> _{interaction} ^c	20-40 years old (n=164)	<i>P</i> ^d	41-60 years old (n=201)	<i>P</i> ^d	<i>P</i> _{interaction} ^e
miR-24-3p	1.066 (0.966, 1.177)	0.204	1.190 (1.106, 1.282)	3.51×10 ^{-6*}	0.098	1.111 (1.013, 1.219)	0.025	1.174 (1.085, 1.271)	7.56×10 ^{-5*}	0.684
miR-27a-3p	1.011 (0.919, 1.111)	0.826	1.142 (1.062, 1.229)	3.80×10 ^{-4*}	0.049	1.070 (0.977, 1.172)	0.145	1.097 (1.017, 1.185)	0.018	0.875
miR-142-5p	1.118 (1.009, 1.240)	0.034	1.094 (1.021, 1.173)	0.011	0.814	1.088 (0.993, 1.191)	0.071	1.092 (1.013, 1.178)	0.023	0.870
miR-28-5p	1.070 (0.962, 1.192)	0.214	1.173 (1.094, 1.258)	7.38×10 ^{-6*}	0.187	1.111 (1.015, 1.217)	0.022	1.167 (1.078, 1.264)	1.49×10 ^{-4*}	0.737
miR-150-5p	1.068 (0.960, 1.188)	0.224	1.099 (1.028, 1.174)	0.005 [*]	0.684	1.063 (0.977, 1.155)	0.155	1.101 (1.019, 1.188)	0.014	0.860

Abbreviations: FR: frequency ratio.

^aLog2-transformed. ^bPoisson regression analysis with adjustment for age, drinking status, working years, workplace, BMI, ΣOH-PAHs, and BPDE-Alb adducts.

^c*P*_{interaction} was calculated by entering the interaction term (miRNA*smoking status) into Poisson regression models, with adjustment for age, drinking status, working years, workplace, BMI, ΣOH-PAHs, and BPDE-Alb adducts. ^dPoisson regression analysis with adjustment for smoking status, pack-years of smoking, drinking status, working years, workplace, BMI, ΣOH-PAHs, and BPDE-Alb adducts. ^e*P*_{interaction} was calculated by entering the interaction term (miRNA*age group) into Poisson regression models, with adjustment for smoking status, pack-years of smoking, drinking status, working years, workplace, BMI, ΣOH-PAHs, and BPDE-Alb adducts.