# THE LANCET Planetary Health

## Supplementary appendix

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

Supplement to: Balakrishnan K, Steenland K, Clasen T, et al. Exposure–response relationships for personal exposure to fine particulate matter (PM<sub>2-5</sub>), carbon monoxide, and black carbon and birthweight: an observational analysis of the multicountry Household Air Pollution Intervention Network (HAPIN) trial. *Lancet Planet Health* 2023; **7:** e387–96.

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Pollutant	Model	Estimate (95% CI)	p-value	AIC
PM <sub>2·5</sub>	Quartile 1	reference		40218
	Quartile 2	-24.9 (-68.9, 19.2)	0.27	
	Quartile 3	-5.2 (-50.1, 39.7)	0.82	
	Quartile 4**	-24.5 (-70.4, 21.4)	0.30	
BC	Quartile 1	reference		37885
	Quartile 2	-29.9 (-75.3, 15.5)	0.20	
	Quartile 3	-18.3 (-64.9, 28.3)	0.44	
	Quartile 4**	-39.2 (-86.1, 7.7)	0.10	
СО	Quartile 1	reference		
	Quartile 2	44.0 (1.7, 86.2)	0.04	
	Quartile 3	47.4 (4.9, 89.9)	0.03	
	Quartile 4**	16.1 (-27.3, 59.5)	0.47	41015

Table S1: Change in birthweight for by quartiles of exposure\*

\*All models adjusted for mother's education, baseline BMI, nulliparity, diet diversity, food insecurity score, second-hand smoke, baseline hemoglobin, age, as well as infant sex, and 10 randomization strata. \*\*Cutpoints for quartiles for PM<sub>2.5</sub>: 40·8, 69·5, and 115·3cutpoints for quartiles for BC: 5·4, 8·6, and 12·7, cutpoints for quartiles for CO: 0·55, 1·13, and 2·23.

#### Table S2: Change in z-scores by quartiles of exposure\*

Pollutant	Model	Estimate (95% CI)	p-value	AIC
PM <sub>2.5</sub>	Quartile 1	reference		7024
	Quartile 2	-0.094 (-0.191, 0.002)	0.06	
	Quartile 3	-0.045 (-0.144, 0.054)	0.37	
	Quartile 4**	-0.102 (-0.202, 0.001)	0.05	
BC	Quartile 1	reference		6595
	Quartile 2	-0.063 (-0.163, 0.037)	0.21	
	Quartile 3	-0.076 (-0.179, 0.023)	0.14	
	Quartile 4**	-0.137 (-0.240, -0.034)	0.01	
СО	Quartile 1	reference		7212
	Quartile 2	0.013 (0.038, 0.225)	0.01	
	Quartile 3	0.097 (0.003, 0.192)	0.04	
	Quartile 4**	0.060 (-0.037, 0.156)	0.23	

\*All models adjusted for mother's education, baseline BMI, nulliparity, diet diversity, food insecurity score, second-hand smoke, baseline hemoglobin, age, infant sex, and 10 randomization strata.

\*\*Cutpoints for quartiles for PM2.5: 40.8, 69.5, and 115.3 cutpoints for quartiles for BC: 5.4, 8.6, and 12.7, cutpoints for quartiles for CO: 0.55, 1.13, and 2.23.

Outcome	Exposure	Model	AIC
birthweight	$PM_{2\cdot 5}$	linear	40211.7
		Log linear	40215.0
		quadratic	40206.2
		2 piece linear	40207.4
		Cubic spline	40212.4
birthweight	BC	linear	37876.6
		Log linear	37880.8
		quadratic	37878.2
		2 piece linear	37877.5
		Cubic spline	37878.2
z-score	PM <sub>2·5</sub>	linear	7021.6
		Log linear	7023-2
		quadratic	7023.5
		2 piece linear	7022.9
		Cubic spline	7023.6
z-score	BC	linear	6590.7
		Log linear	6592.8
		quadratic	6592.5
		2 piece linear	6592.3
		Cubic spline	6592.5

Table S3. AICs for different models for birthweight and z-score, in relation to PM<sub>2.5</sub> and BC

#### Table S4. Change in outcome by increase in pollutant IQR for full term births

Outcome	Pollutant and model	Change in outcome (g)	LCL change	UCL change
birthweight	PM <sub>2.5</sub> linear	-13.315	-27.017	0.395
	BC linear	-19.701	-34.757	-4.645
	CO linear	-3.183	-11.570	5.204
z-score	PM <sub>2.5</sub> linear	-0.045	-0.075	-0.015
	BC linear	-0.054	-0.089	-0.018
	CO linear	-0.006	-0.025	0.014

Table S5. Change in outcome by increase in pollutant IQR by infant sex

Outcome	Pollutant and model	Change in outcome (g)	LCL change	UCL change
birthweight (male)	PM <sub>2.5</sub> linear	-2.776	-23.415	17.863
	BC linear	-7.871	-31.385	15.643
	CO linear	-6.144	-19.729	7.441
birthweight (female)	PM <sub>2.5</sub> linear	-26.674	-46.009	-7.339
	BC linear	-34.745	-56.388	-13.103
	CO linear	-0.081	-12.063	11.901
z-score (male)	PM <sub>2.5</sub> linear	-0.032	-0.076	0.012
	BC linear	-0.040	-0.090	0.010
	CO linear	-0.010	-0.039	0.020
z-score (female)	PM <sub>2.5</sub> linear	-0.031	-0.075	0.012
	BC linear	-0.059	-0.108	-0.010
	CO linear	0.002	-0.026	0.029

	Birthwe	eight <mark>(gr</mark>	ams)		PM <sub>2·5</sub>	(μg/m <sup>3</sup> )			BC (	ug/m <sup>3</sup> )			CO	(ppm)	
	Mean	SD	Ν	Mean	SD	Median	Ν	Mean	SD	Median	Ν	Mean	SD	Median	N
India	2591	394	773	94.29	90.90	69.32	710	10.44	8.49	8.24	698	1.49	2.21	0.87	735
Rwanda	3022	439	749	93.50	72.45	75.42	695	10.42	6.38	9.19	618	1.94	2.66	1.04	710
Guatemala	2861	428	750	112.88	83.89	88.62	703	10.90	6.40	9.94	677	1.56	1.64	1.08	727
Peru	3180	410	730	64.37	79.85	38.99	609	7.96	7.71	5.75	567	3.09	4.41	1.76	600
Study-wide	2909	471	2560	92.19	83.85	69.45	2717	10.01	7.39	8.57	2560	1.97	2.90	1.13	2772

#### Table S6: Birthweights and pollutant exposures by country and overall

Pollutant and model	Country	Exposure-response coefficient	p-value
$PM_{2\cdot 5}$ linear model	Guatemala	-0.11	0.55
	India	-0.09	0.57
	Peru	-0.16	0.44
	Rwanda	-0.56	0.01
	Combined	-0.20	0.04
BC linear model	Guatemala	-4.5	0.07
	India	-2.6	0.13
	Peru	-0.9	0.69
	Rwanda	-3.5	0.22
	Combined	-3.0	0.01
CO linear model	Guatemala	-10.6	0.26
	India	3.1	0.61
	Peru	-5.6	0.13
	Rwanda	10.9	0.24
	Combined	-1.8	0.49

Table S7. Linear models for birthweight and three pollutants in each country

Table S8. Linear models for weight-for-gestational age z-scores and three pollutants in each country

Pollutant and model	Country	Exposure-response coefficient	p-value
PM <sub>2.5</sub> linear model	Guatemala	-0.0005	0.29
	India	-0.0004	0.25
	Peru	-0.0002	0.61
	Rwanda	-0.0006	0.23
	Combined	-0.0004	0.04
BC linear model	Guatemala	-0.125	0.03
	India	-0.006	0.09
	Peru	-0.001	0.76
	Rwanda	-0.005	0.37
	Combined	-0.007	0.01
CO linear model	Guatemala	-0.006	0.79
	India	0.011	0.40
	Peru	-0.010	0.23
	Rwanda	0.009	0.49
	Combined	-0.002	0.78

	Coefficient	t-statistic	p-value					
Birthweight	Birthweight							
BC overall avg	-2.9996	7.42	0.0065					
BC BL*	-0.856	1.06	0.3044					
BC P1*	-1.8725	3.38	0.0661					
BC P2*	-1.8086	3.62	0.0571					
PM <sub>2.5</sub> overall avg	-0.1983	4.3	0.0381					
PM <sub>2.5</sub> BL	-0.07	1.15	0.2843					
PM <sub>2.5</sub> P1	-0.0033	0	0.9728					
PM <sub>2.5</sub> P2	-0.0645	0.51	0.4761					
CO overall avg	-1.8209	0.46	0.4999					
CO BL	2.3535	1.59	0.2076					
CO P1	2.1364	0.61	0.4357					
CO P2	-3.7919	2.13	0.1448					
Weight-for-gestation	nal age z-scores							
BC overall avg	-0.0067	7.62	0.0058					
BC BL*	-0.0028	2.41	0.1202					
BC P1*	-0.004	3.09	0.0786					
BC P2*	-0.0043	4.17	0.0411					
PM <sub>2.5</sub> overall avg	-0.0004	4.32	0.0377					
PM <sub>2.5</sub> BL	-0.0002	2.61	0.1063					
PM <sub>2.5</sub> P1	-0.0001	0.19	0.6659					
PM <sub>2.5</sub> P2	-0.0002	0.88	0.3475					
CO overall avg	-0.0017	0.08	0.7787					
CO BL	0.0064	2.38	0.1226					
CO P1	0.0043	0.5	0.4794					
CO P2	-0.0073	1.58	0.2089					
*BC = black carbon	$\frac{1}{CO}$ – carbon mono	vide						

Table S9. Exposure-response coefficient using exposure as measured in early, middle, and late gestation

\*BC = black carbon, CO = carbon monoxide \*\*BL = baseline measurement, P1 and P2 are 2<sup>nd</sup> and 3<sup>rd</sup> measurements

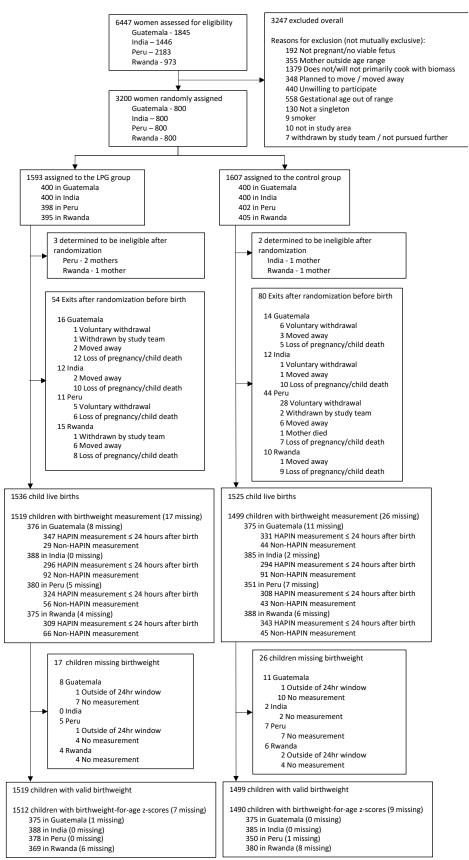
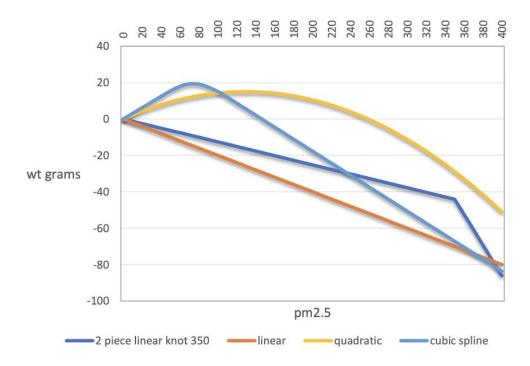
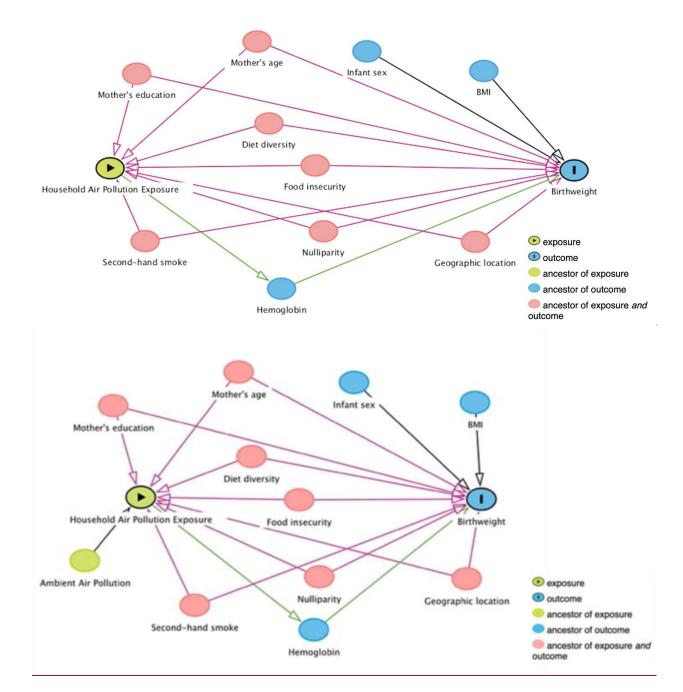


Figure S1. CONSORT diagram



**Figure S2.** Linear, quadratic, 2-piece linear and cubic spline models for PM<sub>2-5</sub> and birthweight Linear model AIC 40212; 2-piece spline model knot 350, AIC 40207; Quadratic AIC 40206. Cubic spline model AIC 40212.



**Figure S3.** Directed acyclic graph (DAG) of the a priori covariates adjusted in the exposure-response models for the associations between household air pollution exposure and birthweight. Causal path (green line); Biasing path (red line).

#### Appendix. Potential impact of decrease in HAP on increased birthweight and infant mortality

We found a coefficient of -0.15 birthweight change per unit of  $PM_{2.5}$ , or 1.5 gm decrease per 10 µg/m<sup>3</sup> increase in  $PM_{2.5}$ . This translates to about a 20 gm increased in birthweight for a lowering of 1 IQR of  $PM_{2.5}$  in our study. Another recent paper (Balakrishnan et al. 2018) of HAP and birthweight found a decrease of 4 gm per 10 µg/m<sup>3</sup>

increase in PM<sub>2.5</sub>. Let us then assume that the true effect is about a 2.7 gm decrease in birthweight per 10  $\mu$ g/m<sup>3</sup>, an average of our study and Balakrishnan et al's. 2018 finding. Let us further assume that the average personal exposure to PM<sub>2.5</sub> among biomass users worldwide is about 270  $\mu$ g/m<sup>3</sup> (based on reviews by Rehfuess et al. 2014, and Balakrishnan et al. 2014), higher than our study which had a mean of about 150 ug/m<sup>3</sup> (Johnson et al 2022). Let us further assume a gas intervention would drop this to 35  $\mu$ g/m<sup>3</sup>, the WHO interim target 1 goal (WHO 2021) and about what we found in our intervention group (Johnson et al 2022). That would mean that the birthweight among biomass users should increase by about 66gms (=24\*2.75), with a decrease of PM<sub>2.5</sub> from 270  $\mu$ g/m<sup>3</sup> to 35  $\mu$ g/m<sup>3</sup>. While such an increase may not be of clinical significance for individuals, shifting birthweight this much would be expected to decrease infant mortality on a population basis. If we assume that there is a drop of one unit (per 1000) in infant mortality for each 30 gm of birthweight (Wilcox, 2001), and an infant mortality rate of 30/1000 in LMICs (World Bank 2017). an improvement of 66 gm would translate to an approximate drop from 30 to 28/1000 in infant mortality. WHO estimates that there are 2.4 million infant deaths per year (in the first month of life, when most infant deaths occur), the great majority in LMICs (WHO 2020), a 7% decrease translates then, to a decrease of about 168,000 infant deaths per year.

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