Supplementary data

Title

Maternal HIV infection is associated with distinct systemic cytokine profiles throughout

pregnancy in South African women

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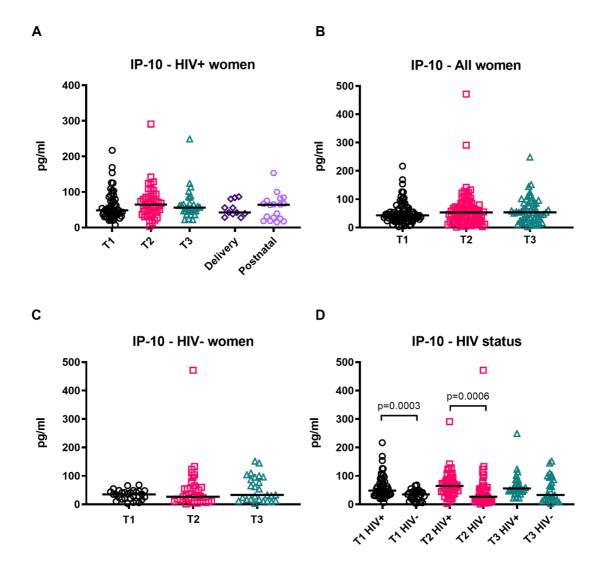
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Supplementary Figure 1. Systemic IP-10 concentrations across pregnancy. IP-10 concentration was measured in maternal plasma collected during the first (T1), second (T2) and third (T3) trimester in women living with HIV (HIV+ women) and HIV-negative women (HIV- women), as well as at delivery and six weeks postnatal for n=11 and n=18 HIV+ women, respectively. IP-10 measured in (A) HIV+ women, (B) all women (both HIV positive and HIV negative), (C) HIV- women and (D) HIV+ versus HIV- women. The Mann-Whitney U-test was used to compare median values between two groups within a trimester. The Kruskal-Wallis test was used to compare median values of more than two groups followed by Dunn's multiple comparisons test to determine differences between two groups.

			Ti	rimester							
	-	1		2		3	D	elivery	P	ostnatal	P
	n		n		n		n		n		
All Women											
IL-1β	23_a	18.9%	16 _a	16.2%	9 _a	13.2%					0.625
TNF-α	1 _a	0.8%	4a	4.0%	2 _a	2.9%					0.261
IP-10	83a	68.0%	82 _b	82.8%	57 _{a,b}	83.8%					0.01
IL-8	4 a	3.3%	6a	6.1%	2 _a	2.9%					0.559
IFNα2	6a	4.9%	7 _a	7.1%	7 _a	10.3%					0.374
IFNβ	36a	29.5%	26a	26.3%	15a	22.1%					0.528
IFNλ1	12 _a	9.8%	33_b	33.3%	25 _b	36.8%					0.00
IFNλ2/3	35a	28.7%	33 _a	33.3%	16a	23.5%					0.398
IFNγ	3_{a}	2.5%	4 _a	4.0%	3_a	4.4%					0.772
IL-2	19 _a	15.7%	11 _a	11.1%	10 _a	14.7%					0.642
L-12p70	3 _a	2.5%	4 _a	4.0%	3_a	4.4%					0.772
L-4	22 _a	18.2%	11 _a	11.1%	14 _a	20.6%					0.198
L-5	11 _a	9.1%	11 _a	11.1%	13 _a	19.1%					0.115
L-13	22 _a	18.2%	16a	16.2%	11 _a	16.2%					0.920
L-17A	19 _a	15.7%	7 _a	7.1%	9 _a	13.2%					0.151
L-17F	36a	29.8%	24 _a	24.2%	17 _a	25.0%					0.636
IL-21	13 _a	10.7%	10a	10.1%	5a	7.4%					0.768
L-22	11 _a	10.9%	11 _a	12.8%	8 _a	12.9%					0.909
L-6	4 a	3.3%	5a	5.1%	2 _a	2.9%					0.732
L-9	15 _a	12.4%	12a	12.1%	13 _a	19.1%					0.373
GM-CSF	3_a	2.5%	6 _a	6.1%	2 _a	2.9%					0.387
IL-10	7 _a	5.7%	10a	10.1%	5a	7.4%					0.454
IIV+ wome	n										
IL-1β	2 _a	3.7%	3_a	6.7%	3_a	9.7%	O ¹	0.0%	O ¹	0.0%	0.617
TNF-α	1 _a	1.9%	3 _a	6.7%	2 _a	6.5%	O ¹	0.0%	0 ¹	0.0%	0.613
IP-10	54 ¹	100.0%	45 ¹	100.0%	31 ¹	100.0%	11 ¹	100.0%	18¹	100.0%	
IL-8	2 _a	3.7%	3 _a	6.7%	0 ¹	0.0%	O ¹	0.0%	0 ¹	0.0%	0.603
IFNα2	3 _a	5.6%	3 _a	6.7%	5a	16.1%	2 _a	18.2%	0 ¹	0.0%	0.162
FNβ	7 _a	13.0%	6 _a	13.3%	3 _a	9.7%	1 _a	9.1%	O ¹	0.0%	0.614
IFNλ1	5a	9.3%	21 _b	46.7%	17 _b	54.8%	1 _{a,b}	9.1%	0 ¹	0.0%	0.000
IFNλ2/3	4 _a	7.4%	7 _a	15.6%	2 _a	6.5%	3 _a	27.3%	0 ¹	0.0%	0.097
IFNγ	O ¹	0.0%	1 _a	2.2%	1 _a	3.2%	O ¹	0.0%	O ¹	0.0%	0.559
IL-2	17 _a	31.5%	9 _a	20.0%	7 _a	22.6%	4 a	36.4%	2 _a	11.1%	0.345
IL-12p70	2 _a	3.7%	4 _a	8.9%	3 _a	9.7%	1 _a	9.1%	0 ¹	0.0%	0.489
IL-4	11 _a	20.4%	6 _a	13.3%	8 _a	25.8%	2 _a	18.2%	O ¹	0.0%	0.130
IL-5	9 _a	16.7%	10a	22.2%	10a	32.3%	3 _a	27.3%	1 _a	5.6%	0.190
IL-13	9 _a	16.7%	8a	17.8%	6a	19.4%	2 _a	18.2%	1 _a	5.6%	0.772
IL-17A	16 _a	29.6%	6 _a	13.3%	6 _a	19.4%	5 _a	45.5%	4 _a	22.2%	0.132
L-17F	20a	37.0%	13 _a	28.9%	8a	25.8%	5a	45.5%	3 _a	16.7%	0.377
L-21	3 _a	5.6%	3 _a	6.7%	2 _a	6.5%	2 _a	18.2%	O ¹	0.0%	0.443
L-22	8a	14.8%	7 _a	15.6%	4 _a	12.9%	4 _a	36.4%	2 _a	11.1%	0.460
L-6	4 _a	7.4%	4 _a	8.9%	2 _a	6.5%	1 _a	9.1%	1 _a	5.6%	1.000
L-9	11 _a	20.4%	6a	13.3%	7 _a	22.6%	2 _a	18.2%	2a	11.1%	0.771
GM-CSF	2a	3.7%	2 _a	4.4%	2 _a	6.5%	O ¹	0.0%	0 ¹	0.0%	0.924
IL-10	7 _a	13.0%	7 _a	15.6%	5a	16.1%	3 _a	27.3%	1 _a	5.6%	0.584
IIV- womer											
IL-1β	21 _a	30.9%	13 _a	24.1%	6a	16.2%					0.245

TNF-α	0 ¹	0.0%	1 _a	1.9%	O ¹	0.0%	0.572
IP-10	29a	42.6%	37 _b	68.5%	26 _b	70.3%	0.003
IL-8	2 _a	2.9%	3_a	5.6%	2 _a	5.4%	0.693
IFNα2	3 _a	4.4%	4 _a	7.4%	2 _a	5.4%	0.905
IFNβ	29a	42.6%	20a	37.0%	12a	32.4%	0.567
IFNλ1	7 _a	10.3%	12 _a	22.2%	8 a	21.6%	0.152
IFNλ2/3	31 _a	45.6%	26a	48.1%	14a	37.8%	0.647
IFNγ	3 _a	4.4%	3 a	5.6%	2 _a	5.4%	1.000
IL-2	2 _a	3.0%	2 a	3.7%	3 _a	8.1%	0.483
IL-12p70	1 _a	1.5%	O ¹	0.0%	O ¹	0.0%	1.000
IL-4	11 _a	16.4%	5a	9.3%	6a	16.2%	0.472
IL-5	2 _a	3.0%	1 _a	1.9%	3 _a	8.1%	0.321
IL-13	13 _a	19.4%	8 a	14.8%	5a	13.5%	0.733
IL-17A	3 _a	4.5%	1 _a	1.9%	3 _a	8.1%	0.426
IL-17F	16 _a	23.9%	11 _a	20.4%	9 _a	24.3%	0.885
IL-21	10a	14.9%	7 _a	13.0%	3 _a	8.1%	0.620
IL-22	3 _a	6.4%	4 a	9.8%	4 a	12.9%	0.567
IL-6	O ¹	0.0%	1 _a	1.9%	0 ¹	0.0%	0.572
IL-9	4 a	6.0%	6a	11.1%	6a	16.2%	0.256
GM-CSF	1 _a	1.5%	4 a	7.4%	O ¹	0.0%	0.127
IL-10	O ¹	0.0%	3 _a	5.6%	O ¹	0.0%	0.050

Supplementary Table 1. Systemic cytokines across pregnancy. Cytokines and chemokines were measured in maternal plasma collected during the first, second and third trimester in all women studied (both women living with HIV (HIV+ women) and HIV-negative women (HIV- women)), as well as at delivery and six weeks postnatal for 11 and 18 women living with HIV, respectively. Data presented are the number and percentage of samples in which cytokines were detected by multiplex cytokine assay. The Chi-square test or Fisher's exact test was used for statistical comparisons as appropriate. Values in the same row not sharing the same subscript are significantly different in the two-sided test of equality for column proportions. Cells with no subscript are not included in the test. ¹This category is not used in comparisons because its column proportion is equal to zero or one.

			HIV S	tatus	;			Preterm	Stat	us			SGA Status			
		Positive		Ne	gative	-	Р	reterm	7	Гerm	-	SGA		no	SGA	<u>-</u>
		n		n		P	n		n		P	n		n		P
IL-1β	T1	2	3.7%	21	30.9%	0.000	12	25.5%	11	14.7%	0.158	6	20.7%	17	18.3%	0.789
·	T2	3	6.7%	13	24.1%	0.027	6	19.4%	10	14.7%	0.770	5	21.7%	11	14.5%	0.518
	Т3	3	9.7%	6	16.2%	0.494	0	0.0%	9	13.6%	1.000	4	22.2%	5	10.0%	0.231
TNF-α	T1	1	1.9%	0	0.0%	0.443	0	0.0%	1	1.3%	1.000	1	3.4%	0	0.0%	0.238
	T2	3	6.7%	1	1.9%	0.327	1	3.2%	3	4.4%	1.000	2	8.7%	2	2.6%	0.230
	T3	2	6.5%	0	0.0%	0.204	0	0.0%	2	3.0%	1.000	1	5.6%	1	2.0%	0.462
IP-10	T1	54	100.0%	29	42.6%	0.000	28	59.6%	55	73.3%	0.162	22	75.9%	61	65.6%	0.366
	T2	45	100.0%	37	68.5%	0.000	25	80.6%	57	83.8%	0.776	20	87.0%	62	81.6%	0.755
	Т3	31	100.0%	26	70.3%	0.001	2	100.0%	55	83.3%	1.000	14	77.8%	43	86.0%	0.464
IL-8	T1	2	3.7%	2	2.9%	1.000	1	2.1%	3	4.0%	1.000	2	6.9%	2	2.2%	0.239
	T2	3	6.7%	3	5.6%	1.000	3	9.7%	3	4.4%	0.374	2	8.7%	4	5.3%	0.621
	T3	0	0.0%	2	5.4%	0.496	0	0.0%	2	3.0%	1.000	0	0.0%	2	4.0%	1.000
IFNα2	T1	3	5.6%	3	4.4%	1.000	0	0.0%	6	8.0%	0.081	1	3.4%	5	5.4%	1.000
	T2	3	6.7%	4	7.4%	1.000	2	6.5%	5	7.4%	1.000	3	13.0%	4	5.3%	0.348
IENIO	T3	5	16.1%	2	5.4%	0.233	1	50.0%	6	9.1%	0.197	2	11.1%	5	10.0%	1.000
IFNβ	T1	7	13.0%	29	42.6%	0.001	19	40.4%	17	22.7%	0.043	6	20.7%	30	32.3%	0.233
	T2	6	13.3%	20	37.0% 32.4%	0.011	9	29.0%	17 15	25.0%	0.806 1.000	6	26.1%	20	26.3%	1.000 1.000
IFNλ1	T3 T1	3 5	9.7%	12 7	10.3%	1.000	2	0.0% 4.3%	15 10	22.7% 13.3%	0.127	3	22.2%	11 9	9.7%	1.000
ILINVI	T2	21	9.5% 46.7%	, 12	22.2%	0.018	∠ 11	4.5% 35.5%	22	32.4%	0.127	3 10	43.5%	23	30.3%	0.313
	T3	17	54.8%	8	21.6%	0.016	1	50.0%	24	36.4%	1.000	8	44.4%	23 17	34.0%	0.570
IFNλ2/3	T1	4	7.4%	31	45.6%	0.000	14	29.8%	21	28.0%	0.840	8	27.6%	27	29.0%	1.000
IF NAZ/3	T2	7	7.4 <i>%</i> 15.6%	26	48.1%	0.000	9	29.0%	24	35.3%	0.648	7	30.4%	26	34.2%	0.805
	T3	2	6.5%	14	37.8%	0.001	0	0.0%	16	24.2%	1.000	5	27.8%	11	22.0%	0.747
IFNγ	T1	0	0.0%	3	4.4%	0.254	1	2.1%	2	2.7%	1.000	0	0.0%	3	3.2%	1.000
11 144	T2	1	2.2%	3	5.6%	0.624	1	3.2%	3	4.4%	1.000	1	4.3%	3	3.9%	1.000
	T3	1	3.2%	2	5.4%	1.000	0	0.0%	3	4.5%	1.000	1	5.6%	2	4.0%	1.000
IL-2	T1	17	31.5%	2	3.0%	0.000	8	17.4%	11	14.7%	0.798	8	27.6%	<u>-</u> 11	12.0%	0.075
	T2	9	20.0%	2	3.7%	0.021	2	6.5%	9	13.2%	0.494	5	21.7%	6	7.9%	0.121
	Т3	7	22.6%	3	8.1%	0.167	0	0.0%	10	15.2%	1.000	3	16.7%	7	14.0%	0.717
IL-12p70	T1	2	3.7%	1	1.5%	0.583	0	0.0%	3	4.0%	0.284	1	3.4%	2	2.2%	0.560
-	T2	4	8.9%	0	0.0%	0.040	2	6.5%	2	2.9%	0.587	2	8.7%	2	2.6%	0.230
	Т3	3	9.7%	0	0.0%	0.090	0	0.0%	3	4.5%	1.000	1	5.6%	2	4.0%	1.000
IL-4	T1	11	20.4%	11	16.4%	0.639	8	17.4%	14	18.7%	1.000	9	31.0%	13	14.1%	0.053
	T2	6	13.3%	5	9.3%	0.750	4	12.9%	7	10.3%	0.736	4	17.4%	7	9.2%	0.275
	Т3	8	25.8%	6	16.2%	0.378	0	0.0%	14	21.2%	1.000	3	16.7%	11	22.0%	0.745
IL-5	T1	9	16.7%	2	3.0%	0.012	3	6.5%	8	10.7%	0.530	5	17.2%	6	6.5%	0.131
	T2	10	22.2%	1	1.9%	0.002	5	16.1%	6	8.8%	0.313	5	21.7%	6	7.9%	0.121
	Т3	10	32.3%	3	8.1%	0.015	0	0.0%	13	19.7%	1.000	6	33.3%	7	14.0%	0.090
IL-13	T1	9	16.7%	13	19.4%	0.814	8	17.4%	14	18.7%	1.000	10	34.5%	12	13.0%	0.014
	T2	8	17.8%	8	14.8%	0.787	5	16.1%	11	16.2%	1.000	5	21.7%	11	14.5%	0.518
	Т3	6	19.4%	5	13.5%	0.742	0	0.0%	11	16.7%	1.000	5	27.8%	6	12.0%	0.144

IL-17A	T1	16	29.6%	3	4.5%	0.000	10	21.7%	9	12.0%	0.199	7	24.1%	12	13.0%	0.157
	T2	6	13.3%	1	1.9%	0.044	1	3.2%	6	8.8%	0.429	3	13.0%	4	5.3%	0.348
	Т3	6	19.4%	3	8.1%	0.282	0	0.0%	9	13.6%	1.000	1	5.6%	8	16.0%	0.427
IL-17F	T1	20	37.0%	16	23.9%	0.161	17	37.0%	19	25.3%	0.220	14	48.3%	22	23.9%	0.019
	T2	13	28.9%	11	20.4%	0.355	8	25.8%	16	23.5%	1.000	9	39.1%	15	19.7%	0.093
	Т3	8	25.8%	9	24.3%	1.000	0	0.0%	17	25.8%	1.000	7	38.9%	10	20.0%	0.126
IL-21	T1	3	5.6%	10	14.9%	0.141	5	10.9%	8	10.7%	1.000	4	13.8%	9	9.8%	0.509
	T2	3	6.7%	7	13.0%	0.340	5	16.1%	5	7.4%	0.279	3	13.0%	7	9.2%	0.694
	Т3	2	6.5%	3	8.1%	1.000	0	0.0%	5	7.6%	1.000	2	11.1%	3	6.0%	0.602
IL-22	T1	8	14.8%	3	6.4%	0.213	5	14.3%	6	9.1%	0.507	4	16.7%	7	9.1%	0.286
	T2	7	15.6%	4	9.8%	0.526	0	0.0%	11	18.3%	0.030	4	20.0%	7	10.6%	0.272
	Т3	4	12.9%	4	12.9%	1.000	0	0.0%	8	13.3%	1.000	2	13.3%	6	12.8%	1.000
IL-6	T1	4	7.4%	0	0.0%	0.036	0	0.0%	4	5.3%	0.159	1	3.4%	3	3.2%	1.000
	T2	4	8.9%	1	1.9%	0.174	2	6.5%	3	4.4%	0.647	3	13.0%	2	2.6%	0.08
	Т3	2	6.5%	0	0.0%	0.204	0	0.0%	2	3.0%	1.000	1	5.6%	1	2.0%	0.462
IL-9	T1	11	20.4%	4	6.0%	0.025	3	6.5%	12	16.0%	0.161	7	24.1%	8	8.7%	0.048
	T2	6	13.3%	6	11.1%	0.766	2	6.5%	10	14.7%	0.331	3	13.0%	9	11.8%	1.000
	Т3	7	22.6%	6	16.2%	0.549	0	0.0%	13	19.7%	1.000	3	16.7%	10	20.0%	1.000
GM-CSF	T1	2	3.7%	1	1.5%	0.583	2	4.3%	1	1.3%	0.558	0	0.0%	3	3.2%	1.000
	T2	2	4.4%	4	7.4%	0.686	1	3.2%	5	7.4%	0.662	2	8.7%	4	5.3%	0.621
	Т3	2	6.5%	0	0.0%	0.204	0	0.0%	2	3.0%	1.000	0	0.0%	2	4.0%	1.000
IL-10	T1	7	13.0%	0	0.0%	0.003	1	2.1%	6	8.0%	0.247	3	10.3%	4	4.3%	0.355
	T2	7	15.6%	3	5.6%	0.178	4	12.9%	6	8.8%	0.720	3	13.0%	7	9.2%	0.694
	Т3	5	16.1%	0	0.0%	0.016	0	0.0%	5	7.6%	1.000	1	5.6%	4	8.0%	1.000

Supplementary Table 2. Systemic cytokines across pregnancy according to HIV status, preterm birth status and small-for-gestational-age (SGA) status. Cytokines and chemokines were measured in maternal plasma collected during the first, second and third trimester. Data presented are the number and percentage of samples in which cytokines were detected by multiplex cytokine assay. The Chi-square test or Fisher's exact test was used for statistical comparisons as appropriate.

			Tı	rimester							
		1		2		3	_ [Delivery	Р	ostnatal	P
	n		n		n		n		n		
All Women											
Pro-inflammatory	101	82.8%	92	92.9%	61	89.7%					0.062
Anti-viral	46	37.7%	55	55.6%	98	56.7%					0.009
Th1	22	18.0%	17	17.2%	13	19.1%					0.950
Th2	31	25.6%	28	28.8%	23	33.8%					0.486
Th17	47	38.8%	37	37.4%	21	30.9%					0.537
Pleiotropic	20	16.4%	19	19.2%	14	20.6%					0.746
HIV+ women											
Pro-inflammatory	54	100%	45	100%	31	100%	11	100%	18	100%	
Anti-viral	13	24.1%	25	55.6%	22	71.0%	3	27.3%	0	0.0%	<0.0001
Th1	17	31.5%	12	26.7%	8	25.8%	4	36.4%	2	11.1%	0.499
Th2	13	24.1%	16	35.6%	14	45.2%	3	27.3%	1	5.6%	0.037
Th17	22	40.7%	15	33.3%	9	29.0%	5	45.5%	6	33.3%	0.772
Pleiotropic	15	27.8%	10	22.2%	8	25.8%	3	27.3%	3	16.7%	0.893
HIV- women											
Pro-inflammatory	47	69.1%	47	87.0%	30	81.1%					0.052
Anti-viral	33	48.5%	30	55.6%	16	44.4%					0.557
Th1	5	7.4%	5	9.3%	5	13.5%					0.587
Th2	18	26.9%	12	22.2%	9	24.3%					0.839
Th17	25	37.3%	22	40.7%	12	32.4%					0.723
Pleiotropic	5	7.4%	9	16.7%	6	16.2%					0.228

Supplementary Table 3. Systemic cytokine subgroups across pregnancy. Cytokines and chemokines were measured in maternal plasma collected during the first, second and third trimester in all women studied (both women living with HIV (HIV+ women) and HIV-negative women (HIV- women)), as well as at delivery and six weeks postnatal for 11 and 18 women living with HIV, respectively. Data presented are the number and percentage of samples in which subgroups of cytokines were detected, i.e. pro-inflammatory (IL-1 β , TNF- α , IP-10, IL-8), anti-viral (IFN α 2, IFN β , IFN α 1, IFN α 2, IFN α 3, Th1 (IFN α 7, IL-12, IL-12p70), Th2 (IL-4, IL-5, IL-13), Th17 (IL-17A, IL-17F, IL-21, IL-22) and pleiotropic (IL-6, IL-9, GM-CSF, IL-10) cytokines. The Chi-square test or Fisher's exact test was used for statistical comparisons as appropriate.

		HIV Status						Preter	m Sta	atus			SGA	Statu		
		F	ositive	N	egative	_	P	reterm		Term	_	- ;	SGA	no S	SGA	
		n		n		P	n		n		P	n		n		P
Pro-	T1	54	100%	47	69.1%	<0.0001	38	80.9%	63	84.0%	0.654	26	89.7%	60	76.9%	0.178
inflammatory	T2	45	100%	47	87.0%	0.015	29	93.6%	63	92.7%	>0.99	22	95.7%	56	90.3%	0.668
	Т3	31	100%	30	81.1%	0.013	2	100%	59	89.4%	>0.99	16	88.9%	35	87.5%	>0.99
Anti-viral	T1	13	24.1%	33	48.5%	0.006	20	42.6%	26	34.7%	0.382	9	31.0%	28	33.3%	0.820
	T2	25	55.6%	24	55.6%	>0.99	19	61.3%	36	52.9%	0.438	14	60.9%	33	48.5%	0.306
	Т3	22	71.0%	16	44.4%	0.029	2	100%	36	55.4%	0.502	12	66.7%	20	46.5%	0.151
Th1	T1	17	31.5%	5	7.4%	0.0006	9	19.2%	13	17.3	0.800	8	27.6%	12	13.2%	0.070
	T2	12	26.7%	5	9.3%	0.022	5	16.1%	12	17.7%	0.853	6	26.1%	10	13.3%	0.148
	Т3	8	25.8%	5	13.5%	0.199	0	0.0%	13	19.7%	>0.9 9	4	22.2%	7	14.6%	0.474
Th2	T1	13	24.1%	18	26.9%	0.727	12	26.1%	19	25.3%	0.927	12	41.4%	16	18.0%	0.010
	T2	16	35.6%	12	22.2%	0.143	10	32.3%	18	26.5%	0.553	10	43.5%	13	18.3%	0.015
	Т3	14	45.2%	9	24.3%	0.071	0	0.0%	23	34.9%	0.546	8	44.4%	12	25.5%	0.139
Th17	T1	22	40.7%	25	37.3%	0.701	23	50.0%	24	32.0%	0.049	16	55.2%	24	28.2%	0.009
	T2	15	33.3%	22	40.7%	0.448	13	41.9%	24	35.3%	0.526	11	47.8%	19	27.5%	0.072
	Т3	9	42.9%	12	57.1%	0.763	0	0.0%	21	31.8%	>0.9 9	8	44.4%	11	22.9%	0.085
Pleiotropic	T1	15	27.8%	5	7.4%	0.003	5	10.6%	15	20.0%	0.174	7	24.1%	10	11.1%	0.081
	T2	10	22.2%	9	16.7%	0.485	5	16.1%	14	20.6%	0.601	6	26.1%	8	11.3%	0.083
	Т3	8	25.8%	6	16.2%	0.330	0	0.0%	14	21.2%	>0.9 9	3	16.7%	8	17.0%	>0.99

Supplementary Table 4. Systemic cytokine subgroups across pregnancy according to HIV status, preterm birth status and small-for-gestational-age (SGA) status. Cytokines and chemokines were measured in maternal plasma collected during the first, second and third trimester. Data presented are the number and percentage of samples in which subgroups of cytokines were detected, i.e. pro-inflammatory (IL-1 β , TNF- α , IP-10, IL-8), anti-viral (IFN α 2, IFN β , IFN λ 1, IFN λ 2/3), Th1 (IFN γ , IL-2, IL-12p70), Th2 (IL-4, IL-5, IL-13), Th17 (IL-17A, IL-17F, IL-21, IL-22) and pleiotropic (IL-6, IL-9, GM-CSF, IL-10) cytokines. The Chi-square test or Fisher's exact test was used for statistical comparisons as appropriate.