SUPPLEMENTARY MATERIAL

IgG subclass-specific responses against *Plasmodium falciparum* merozoite antigens are associated with control of parasitemia and protection from symptomatic illness

Danielle I. Stanisic, Jack S. Richards, Fiona J. McCallum, Pascal Michon, Christopher L. King, Sonja Schoepflin, Paul R. Gilson, Vincent J. Murphy, Robin F. Anders, Ivo Mueller, James G. Beeson.

Table S1: Characteristics of antibody responses to *P. falciparum* merozoite antigens among the longitudinal cohort of children.

	n	Proportion Positive (%) ¹	Median ²	IQR ²	T33 ²	T67 ²
AMA-1						
IgG	206	99.5	0.85	[0.46, 1.10]	0.60	1.01
IgG1	206	94.2	0.97	[0.45, 1.26]	0.70	1.17
IgG3	206	76.2	0.09	[0.02, 1.43]	0.02	0.30
MSP1-19						
IgG	206	96.6	1.19	[0.43, 1.68]	0.76	1.51
IgG1	206	69.9	0.17	[0.03, 0.77]	0.06	0.52
IgG3	206	63.6	0.03	[0.01, 0.11]	0.01	0.06
MSP2 3D7						
IgG	206	85.9	0.54	[0.25, 0.79]	0.36	0.69
IgG1	206	41.3	0.01	[0.00, 0.02]	0.00	0.02
IgG3	203	84.2	0.74	[0.20, 1.18]	0.39	1.09
MSP2 FC27						
IgG	206	83.0	0.30	[0.13, 0.60]	0.19	0.49
IgG1	206	51.5	0.02	[0.01, 0.11]	0.01	0.05
IgG3	202	80.7	0.70	[0.13, 1.11]	0.33	0.98

Notes:

- 1. Represents the proportion of samples that were classified as positive to each antigen (defined as optical density > mean + 3SD of the non-exposed control samples).
- 2. Values represent optical density measured by ELISA
 Abbreviations: IQR, Inter-quartile range; T33: 33.33% tertile, T67: 66.67% tertile; AMA1, Apical Membrane Antigen 1; MSP1-19, 19kDa C-terminal region of Merozoite
 Surface Protein 1; MSP2, Merozoite Surface Protein 2 (3D7 and FC27 alleles).

Table S2: The prevalence of IgG1 and IgG3 to merozoite antigens of *P. falciparum* among the longitudinal cohort of children in relation to age and infection.

Antigens	Prevalence ¹ <=9 yrs (%)	Prevalence >9 yrs (%)	p- value ³	Prevalence Pf pos (%) ²	Prevalence Pf neg (%)	p- value
	n = 91	n = 115	varac	n = 139	n = 67	
AMA-1						
IgG1	87.9	99.1	0.001	97.8	86.6	0.001
IgG3	62.6	87.0	< 0.001	83.5	61.2	< 0.001
MSP1- 19 IgG1 IgG3	59.3 51.7	78.3 73.0	0.003 0.002	75.5 72.7	58.2 44.8	0.011 <0.001
MSP2 3D7						
IgG1	42.9	40.0	0.679	48.2	26.9	0.004
IgG3	73.0	93.0	< 0.001	87.6	77.3	0.059
MSP2 FC27						
IgG1	45.1	56.5	0.102	59.7	34.3	0.001
IgG3	68.9	90.2	< 0.001	82.6	76.6	0.311

Notes:

- 1. Samples were classified as positive if the optical density was greater than the mean + 3SD of the non-exposed control samples. Children were divided into 2 age groups: >9 and <9 years to determine the effect of age on the IgG subclass response.
- 2. P. falciparum infection was detected by PCR.
- 3. P values determined by Chi square test.

Abbreviations: Pf: *Plasmodium falciparum*; pos, positive; neg, negative; AMA-1, Apical Membrane Antigen 1; MSP1-19, 19kDa C-terminal region of Merozoite Surface Protein 1; MSP2, Merozoite Surface Protein 2 (3D7 and FC27 alleles).

FIGURE LEGENDS FOR SUPPLEMENTARY FIGURES

- Figure S1: The effect of exposure on the IgG subclass responses to merozoite antigens in Papua New Guinean children. IgG1: unshaded bars; IgG3: light gray bars. Median ODs for IgG1 and IgG3; y error bars indicate lower and upper quartiles. Exposure was determined by antibody responsiveness to *P. falciparum* schizont extract; these responses were divided into quartiles (Q1-4, where Q1 is the lowest quartile).
- Figure S2: Changes to IgG subclass antibody responses to different merozoite antigens in association with age for individuals 0-3 years (n=50), 4-6 years (n=48), 7-9 years (n=50) and >10 years (n=59). IgG1: unshaded bars; IgG3: light gray bars. Median ODs for IgG1 and IgG3; y error bars indicate lower and upper quartiles.



