

Supplemental Digital Content 5:

Hemagglutination inhibition geometric mean titers and geometric mean fold rise for each influenza strain in the fever/no fever groups and antipyretic/no antipyretic groups of the according-to-protocol subset of Pavia-Ruz (2013).¹⁰ Data from subjects in all study groups in study. Individuals were classified in the Fever Y group if vaccine-related fever $\geq 38^{\circ}\text{C}$ occurred during day 0-3 after any vaccine dose. Antipyretic use was administration of antipyretic medication during day 0-3 after any vaccine dose. Figures in bold indicate statistical significance.

Strain/Analysis	Fever N (n=2900)	Fever Y (n=165)	P value	Antipyretic N (n=2543)	Antipyretic Y (n=522)	P value
Geometric mean titers (GMT)						
A/Brisbane/59/07 (H1N1)	146.7	180.3	0.16	160.3	101.6	<0.0001
A/Uruguay/716/07 (H3N2)	177.1	190.8	0.55	184.7	148.0	0.005
B/Florida/4/06	146.6	188.8	0.03	152.5	131.1	0.03
B/Brisbane/3/07	168.5	211.5	0.04	175.8	146.9	0.009
Geometric mean fold rise (GMFR)						
A/Brisbane/59/07 (H1N1)	13.9	16.5	0.16	14.6	11.2	0.0004
A/Uruguay/716/07 (H3N2)	15.2	16.8	0.34	15.9	12.6	0.0001
B/Florida/4/06	17.2	21.4	0.04	17.6	16.9	0.53
B/Brisbane/3/07	18.7	23.4	0.03	19.1	18.1	0.38