### **Supplemental figures**

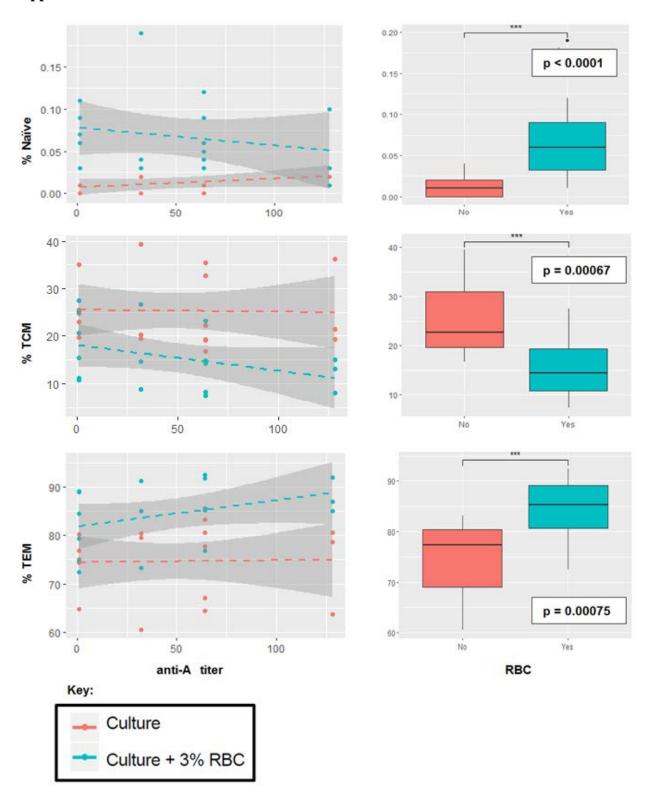
Appendix S1. Additional phenotypic variables by anti-A titer. Left column shows scatterplots of % naïve, %  $T_{CM}$ , %  $T_{EM}$  by anti-A titer with a best fit line by presence/absence RBC contamination. Anti-A titer revealed no appreciable effect on naïve,  $T_{CM}$ , or  $T_{EM}$  percentage. Right column shows boxplots of % naïve, %  $T_{CM}$ , %  $T_{EM}$  by presence/absence of RBC contamination. P-values are shown within inset boxes on each respective boxplot.

Appendix S2. Most upregulated and downregulated transcripts/pathways in cultures with RBC contamination

Appendix S3. Cytokine and chemokine expression in cultures with and without RBC contamination. Hierarchical clustering showing 10 differently expressed analytes in cultures with and without RBC contamination. Violet and green lines on the tree represent cultures with and without 3% RBC spike, respectively.

Appendix S4. RBC agglutination in cultures grown in the presence of high-titer O plasma.

A) Macroscopic RBC agglutination is visible within the culture bag on day 7 in a culture supplemented with plasma containing an anti-A titer of 128. B) Microscopically, agglutination of intact RBCs (clustered, red, biconcave cells) is seen; lymphocytes (clear, round cells) continue to expand evenly across the field.



Upregulated

Phagosome Maturation

Activation of IRF by Cytosolic Pattern Recognition Receptors

Endoplasmic Reticulum Stress Pathway

Interferon Signaling

Assembly of RNA Polymerase II Complex

Estrogen Receptor Signaling Th1 and Th2 Activation Pathway

Phosphatidylethanolamine Biosynthesis II

EIF2 Signaling Eicosanoid Signaling

Polyamine Regulation in Colon Cancer

Th2 Pathway

Clathrin-mediated Endocytosis Signaling

Natural Killer Cell Signaling Mitochondrial Dysfunction Complement System Huntington's Disease Signaling

Autophagy

Assembly of RNA Polymerase I Complex

Protein Ubiquitination Pathway

Rac Signaling

Tumoricidal Function of Hepatic Natural Killer Cells

Glucocorticoid Receptor Signaling

Hematopoiesis from Multipotent Progenitor Cells

PI3K Signaling in B Lymphocytes B Cell Receptor Signaling Death Receptor Signaling

Lipid Antigen Presentation by CD1

Glycoaminoglycan-protein Linkage Region Biosynthesis

NF-κB Activation by Viruses

Downregulated

Cell Cycle Control of Chromosomal Replication

Mismatch Repair in Eukaryotes Hereditary Breast Cancer Signaling

tRNA Charging

Role of BRCA1 in DNA Damage Response

Cell Cycle: G2/M DNA Damage Checkpoint Regulation

Superpathway of Cholesterol Biosynthesis

Small Cell Lung Cancer Signaling

Parkinson's Signaling

Superpathway of Serine and Glycine Biosynthesis I Pyrimidine Deoxyribonucleotides De Novo Biosynthesis I

Cholesterol Biosynthesis I

Cholesterol Biosynthesis II (via 24,25-dihydrolanosterol)

Cholesterol Biosynthesis III (via Desmosterol) DNA damage-induced 14-3-3σ Signaling

Folate Transformations I

Antiproliferative Role of TOB in T Cell Signaling

Serine Biosynthesis

dTMP De Novo Biosynthesis

p53 Signaling ATM Signaling

Epoxysqualene Biosynthesis

Cysteine Biosynthesis/Homocysteine Degradation

**IL-4 Signaling** 

Role of Tissue Factor in Cancer Mitotic Roles of Polo-Like Kinase Ovarian Cancer Signaling

Pancreatic Adenocarcinoma Signaling Docosahexaenoic Acid (DHA) Signaling

EGF Signaling

