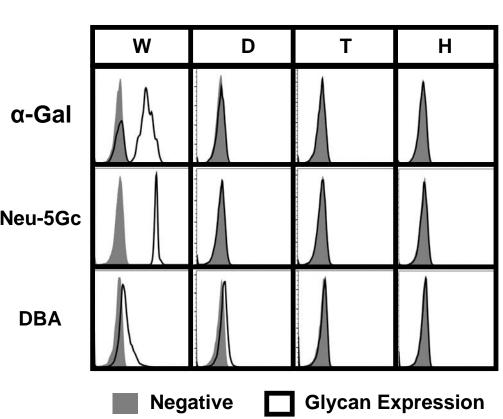
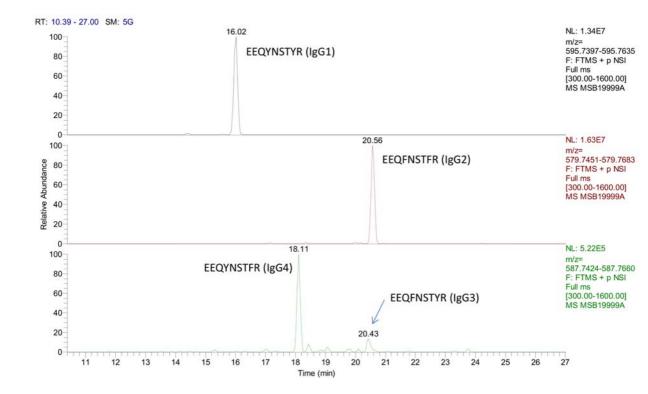
Supplementary Figure 1: RBC Phenotypes



| | AUC per Isotype | | | | Total (AUC) | | %laC1 | %laC2 | %lqG3 | %lgG4 |
|--------------------|-----------------|------------|---------|-----------|-------------|----------|-------|-------|---------|--------|
| | lgG1 | lgG2 | lgG3 | lgG4 | lgG | IgM | %lgG1 | %lgG2 | /olg@3 | %IGG4 |
| Number of values | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| Minimum | 253000000 | 1370000000 | 669000 | 40100000 | 1660000000 | 42500000 | 15 | 72 | 0 | 2.41 |
| 25% Percentile | 253000000 | 1370000000 | 669000 | 40100000 | 1660000000 | 42500000 | 15 | 72 | 0 | 2.41 |
| Median | 377000000 | 1510000000 | 975000 | 65000000 | 1950000000 | 42700000 | 19 | 77 | 0 | 3 |
| 75% Percentile | 843000000 | 2620000000 | 1290000 | 176000000 | 3640000000 | 63400000 | 23 | 82 | 0.06 | 5 |
| Maximum | 843000000 | 2620000000 | 1290000 | 176000000 | 3640000000 | 63400000 | 23 | 82 | 0.06 | 5 |
| Mean | 491000000 | 1833000000 | 978000 | 93700000 | 2417000000 | 49530000 | 19 | 77 | 0.02 | 3.47 |
| Std. Deviation | 311100000 | 684900000 | 310511 | 72350000 | 1069000000 | 12010000 | 4 | 5 | 0.03464 | 1.357 |
| Std. Error of Mean | 179600000 | 395400000 | 179274 | 41770000 | 617400000 | 6934000 | 2.309 | 2.887 | 0.02 | 0.7837 |

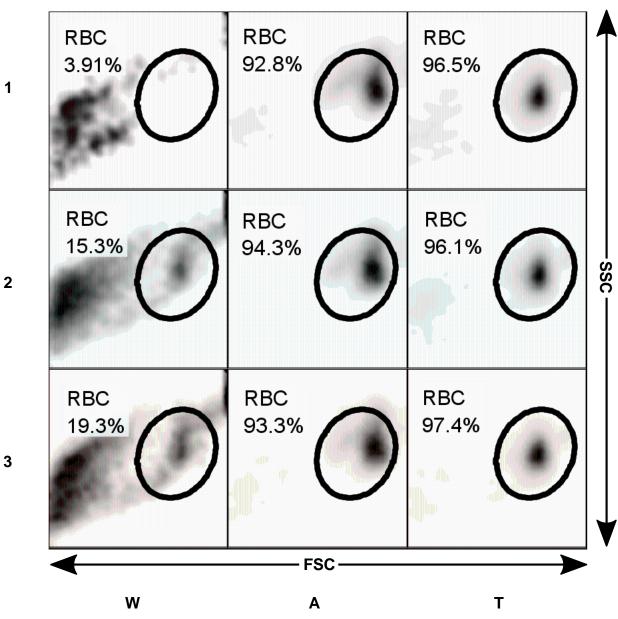
Supplementary Figure 2: Serum from a single donor and RBC from a single triple knockout pig were incubated together. After washing antibodies were eluted and relative bound amounts determined by mass spectroscopy. Three identical experiments were performed.

Supplementary Figure 3: Representative Mass Spectroscopy Chromatogram of Immunoglobulin-Derived Peptides



Supplementary Figure 4: Gating and Fate of RBC Incubated with Different Sera

Serum



Three human sera were incubated with RBC from wild type pigs (W), autologous human RBC (A), and RBC from pigs lacking the GGTA1/CMAH/B4GalNT2 genes (T). After incubating with fluorescent secondary antibodies to report bound human immunoglobulin, cells were analyzed by flow cytometry. Forward scatter (FSC, x-axis) and side scatter (SSC, y-axis) were used to identify RBC. Black ovals represent gates used to select RBC for analysis. The percentages shown next to each gate represents the fraction of total events that resided within the gate. Human sera disrupted wild type swine RBC as seen by an increase in debris and the reduction in RBC falling in the gated regions.