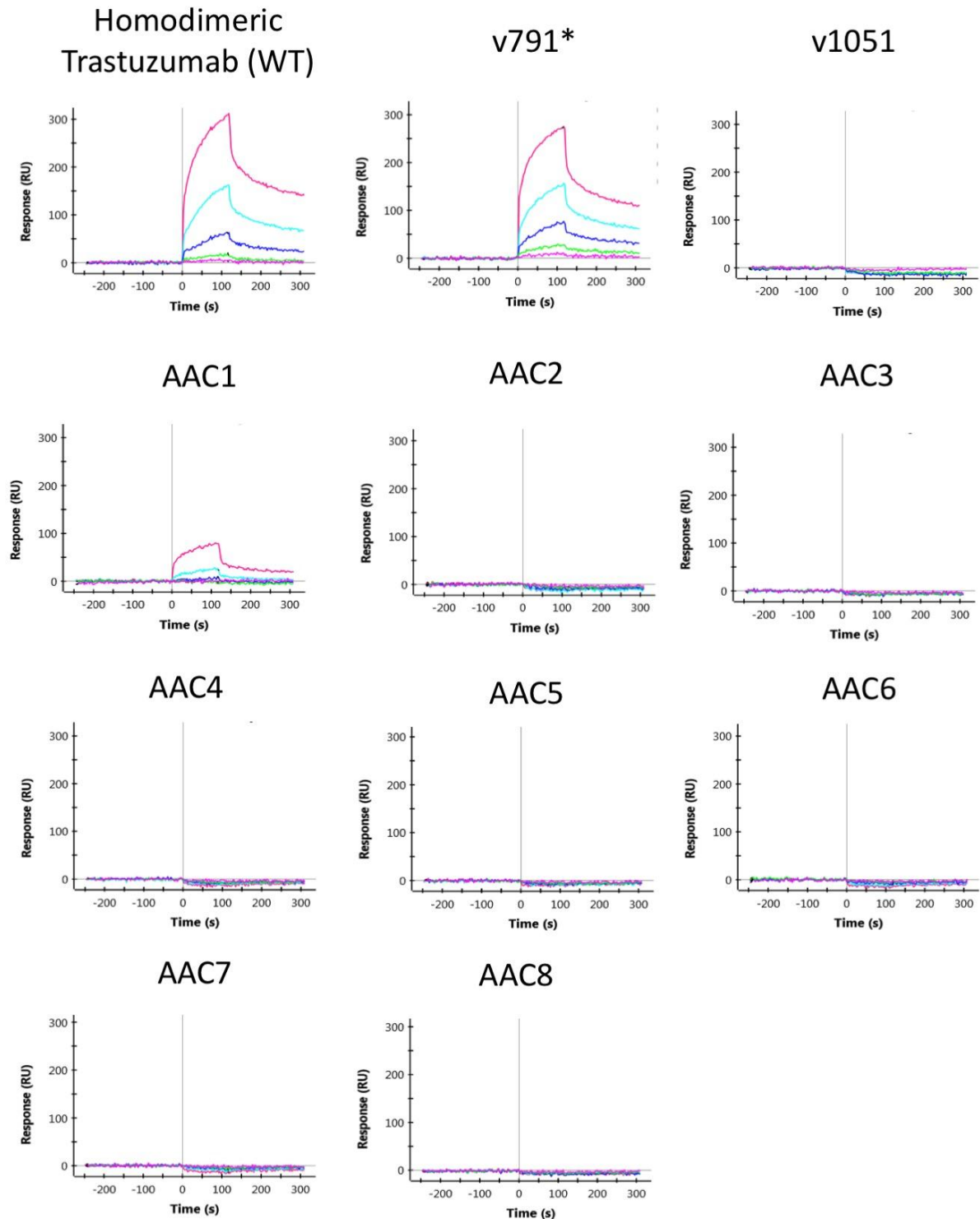


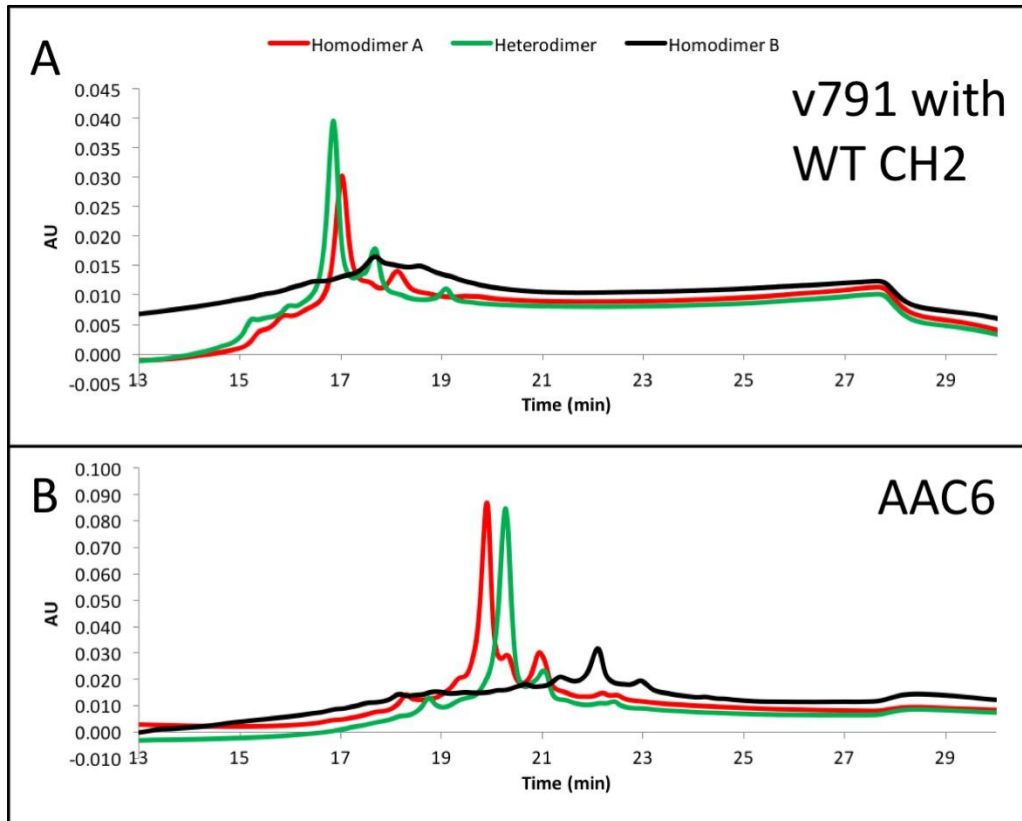
Supplementary Figure S1

Sensograms for the asymmetric antibody constructs binding to C1q (see methods). Due to the avidity of the hexameric C1q, only qualitative data was obtained, where the binding was classified as observed (yes, e.g. WT), diminished compared to the WT control (partial, e.g. AAC1) or not detected (NB) as summarized in Table 1. The different traces correspond to 30 nM top-nominal 3-fold dilutions: 30 nM (red), 10 nM (cyan), 3.33nM (blue), 1.11nM (green), 0.37 nM (magenta). v791* had C-terminal His and mRFP tags.



Supplementary Figure S2

Exemplary result showing the resolution by ion exchange chromatography of homodimers and heterodimers of asymmetric antibody constructs independently expressed and purified for v791 (A) and AAC6 (B) using a pH gradient (from low to high) on a weak cation exchange column.



Supplementary Table S1 – Derived parameters from the interpolated symmetrical sigmoidal shape shown in Figure 4. The standard equation $Y = \text{Bottom} + (\text{Top} - \text{Bottom}) / (1 + 10^{((\text{LogEC50} - X) * \text{HillSlope}))}$ was used using the software Prism.

Parameter	Trastuzumab	v1051	AAC6
Span (Top-Bottom)	47%	25%	Poor fit
EC50 (µg/mL)	2.2E-3	9.9E-2	
HillSlope	1.1	0.84	
R ²	0.95	0.91	

Supplementary Table S2 – Derived parameters from the interpolated symmetrical sigmoidal shape shown in Figure 5. The standard equation $Y = \text{Bottom} + (\text{Top} - \text{Bottom}) / (1 + 10^{((\text{LogEC50} - X) * \text{HillSlope}))}$ was used using the software Prism.

ADCC

Parameter	Commercial Rit	Control WT Rit	AAC9
Span (Top-Bottom)	62	65	Poor fit
EC50 (nM)	1.4E-2	1.2E-1	
HillSlope	1.0	1.2	
R ²	0.98	0.99	

CDC

Parameter	Commercial Rit	Control WT Rit	AAC9
Span (Top-Bottom)	88.6	96.3	Plateau not reached
EC50 (nM)	1.8	2.9	
HillSlope	2.1	2.1	
R ²	0.98	0.99	