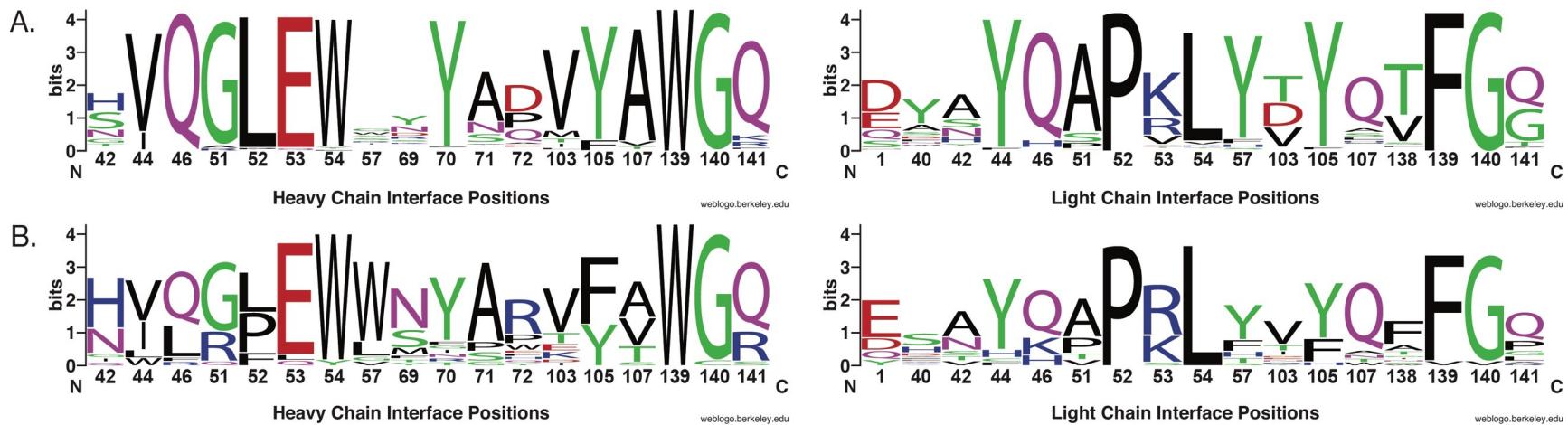


**Figure S1. Distribution of human antibody/antigen co-crystals.** Human antibody/antigen co-crystals were analyzed with Abangle. The resulting angle distributions for the HL(purple), HC1(teal), LC1(Blue), HC2(orange), LC2(green), and dc(grey) values were produced using Prism.



**Figure S2. Sequence variation at the HC/LC interface.** (A) Weblogos11 showing frequency of amino acids in the HC/LC interface for antibodies that fail to bind to the CD4 binding site (CD4BS) epitope. (B) Weblogos showing the frequency of amino acids in the HC/LC interface for CD4BS antibodies.