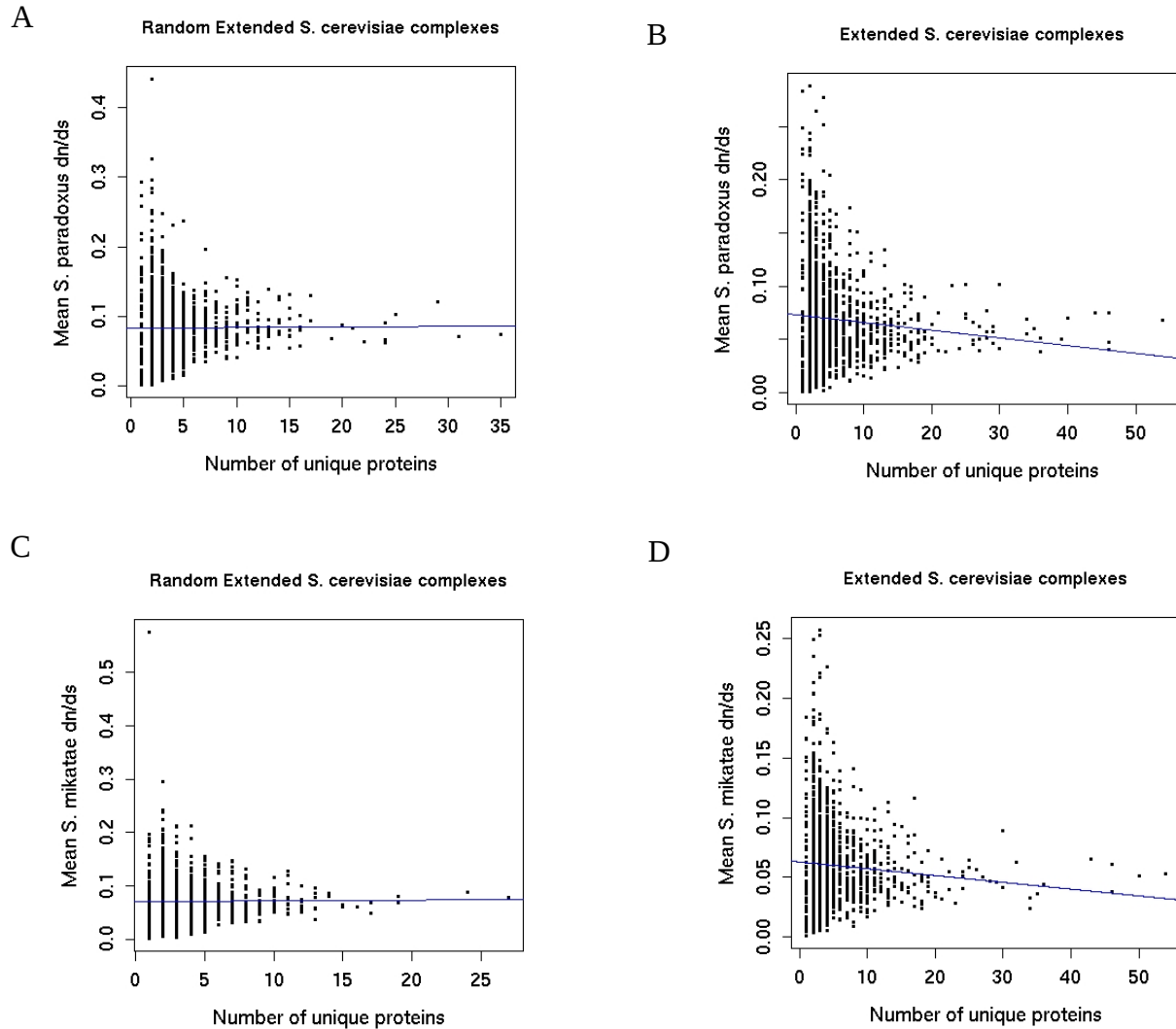


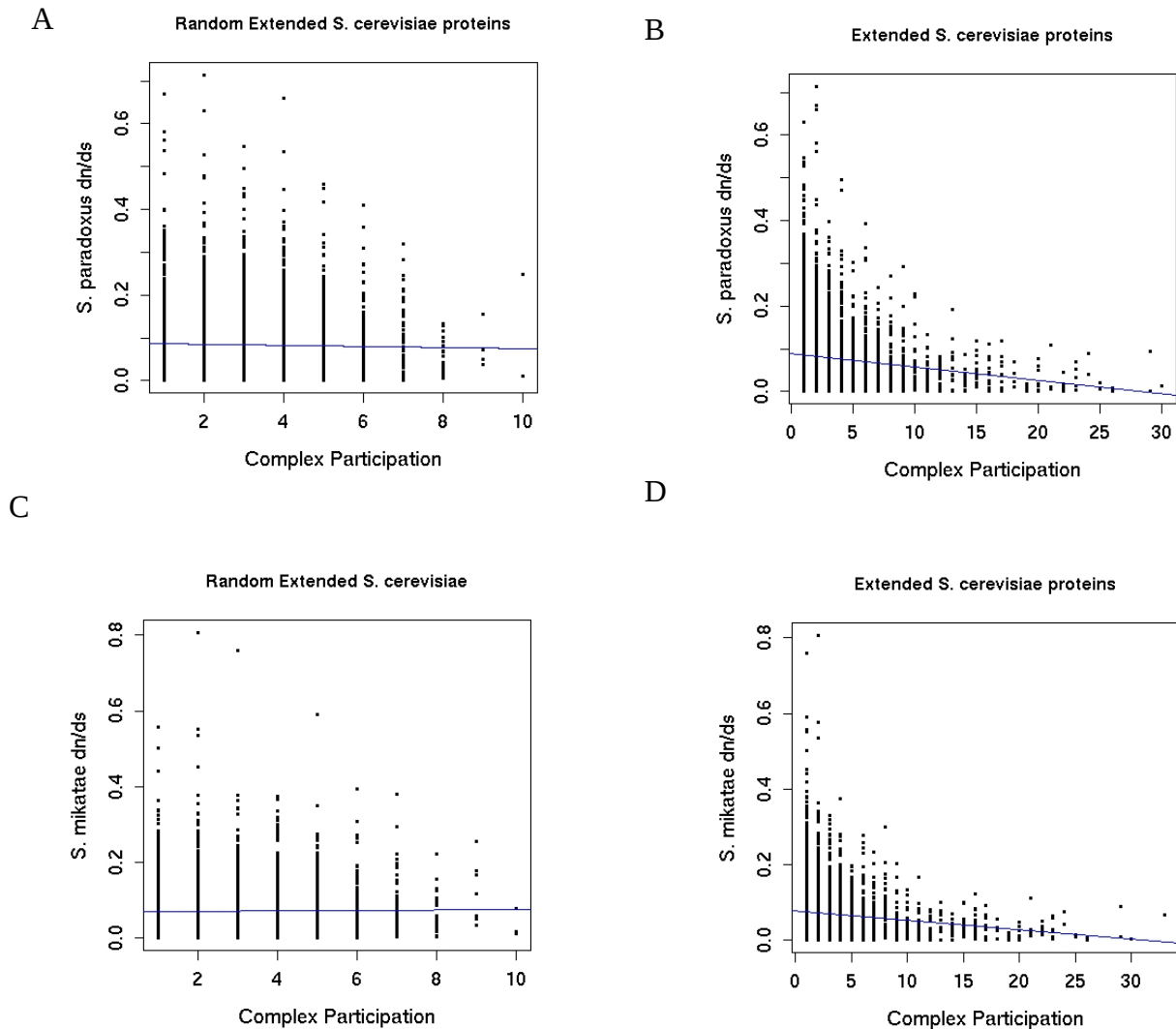
## Additional file 12: Extension and subsets of the protein complex data

### Extended Yeast Complex complexity and dN/dS



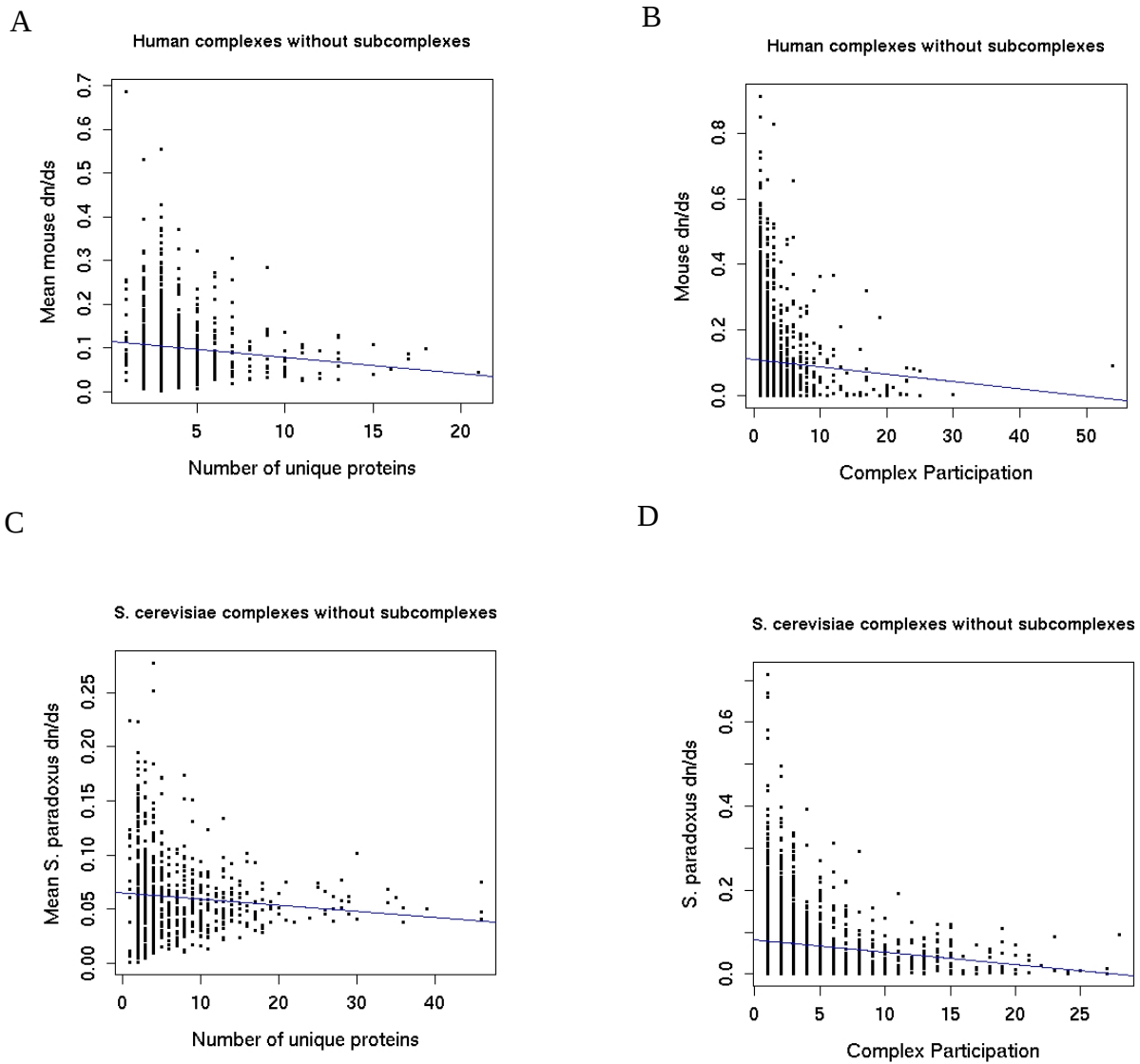
**Figure S12 - Complexes and Selection on Yeast Genes.** B) The mean dN/dS ratio for *S. cerevisiae*-*S. paradoxus* orthologs is plotted against the number of unique proteins for each extended yeast complex. Complexes with more unique proteins tend to have significantly smaller mean dN/dS ratios than those with less unique proteins (t-test:  $P < 1.5 \times 10^{-5}$ ). D) The trend remains present when *S. cerevisiae*-*S. mikatae* orthologs were used to compute the dN/dS ratio (t-test:  $P < 9.3 \times 10^{-5}$ ). A,C) Such trends were not observed in model 1 random complexes.

## Extended Yeast Complex participation and dN/dS



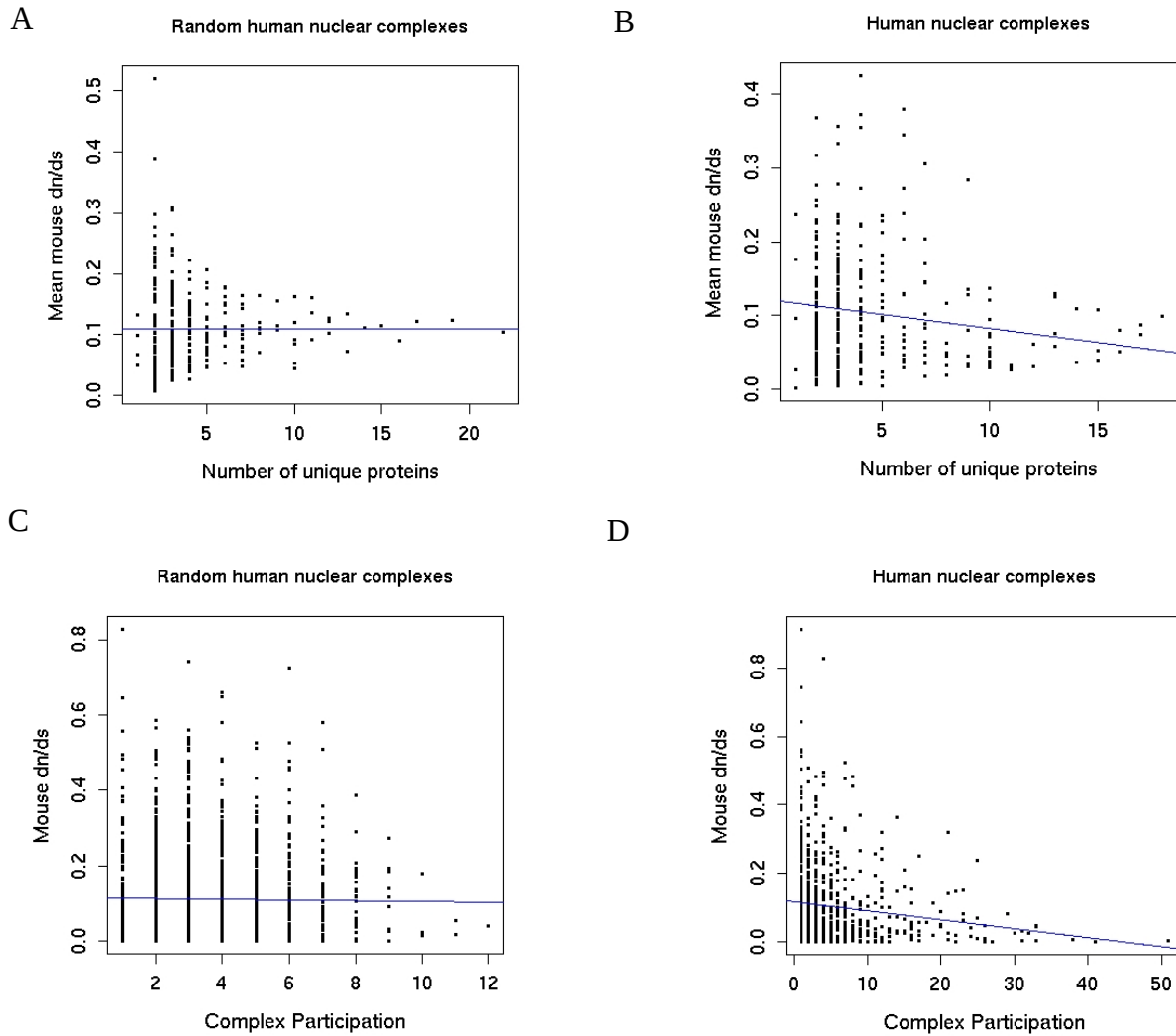
**Figure S12b - Complex participation and Selection on Yeast Genes.** B) The mean dN/dS ratio from aligning *S. cerevisiae* - *S. paradoxus* orthologs is plotted against the number of extended complexes the *S. cerevisiae* protein was found in. Proteins participating in more complexes tend to have significantly smaller dN/dS ratios than those participating in less complexes (t-test:  $P < 2 \times 10^{-16}$ ). D) We found a similar trend when *S. cerevisiae*-*S. mikatae* orthologs were used to calculate dN/dS (t-test:  $P < 2 \times 10^{-16}$ ). A,C) Such trends were not observed in model 1 random complexes.

## Analysis without subcomplexes



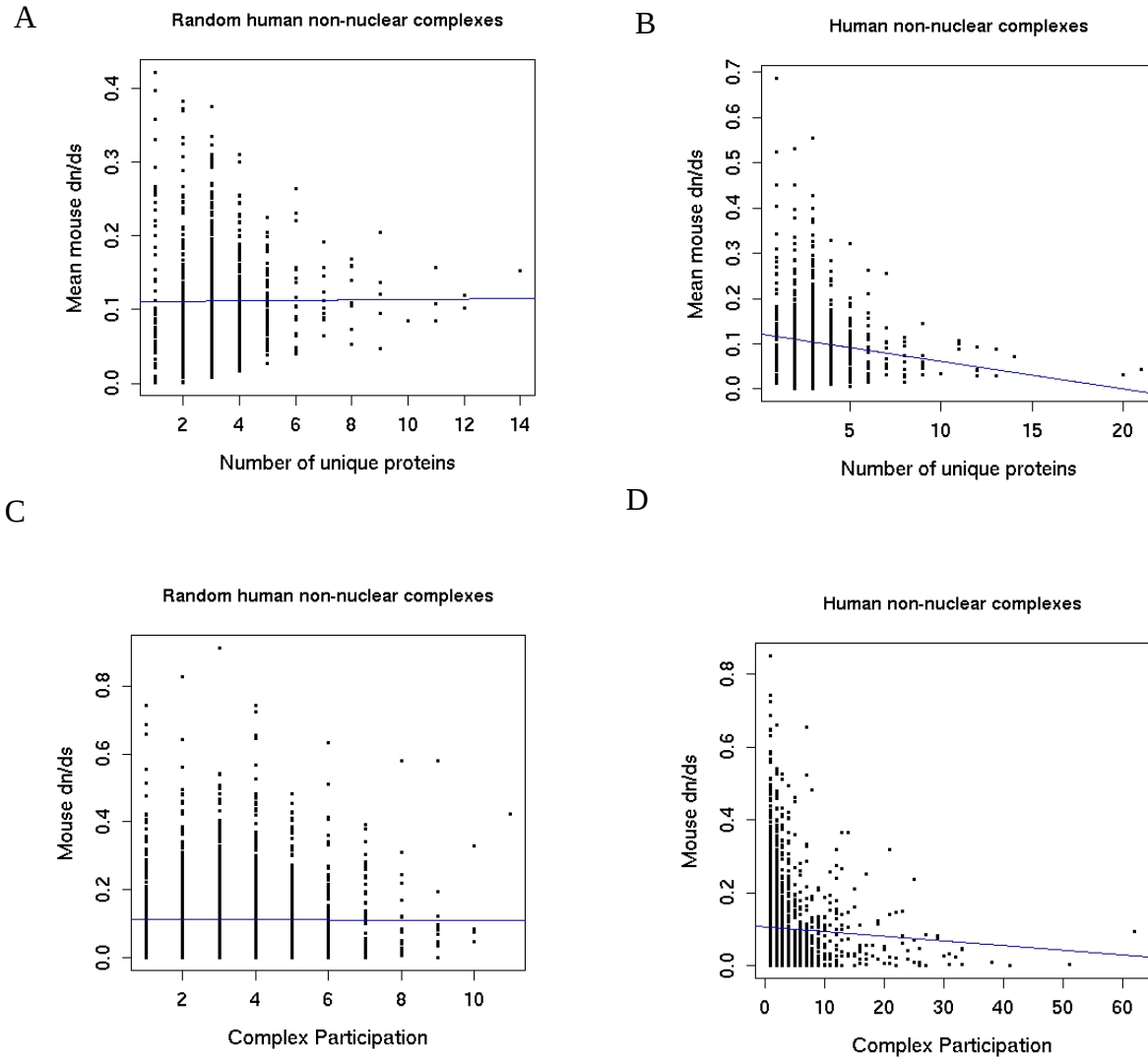
**Figure S12c - Analysis without subcomplexes.** We re-analyzed the human (A,B) and yeast (C,D) complex data with subcomplexes of complexes removed. A,C) The mean dn/ds ratio tends to decrease with increasing complex complexity (t-test:  $P < 0.0002$ ). B,D) The dn/ds ratio of proteins tends to decrease with increasing complex participation (t-test:  $P < 0.0004$ ).

# Nuclear Complexes and dN/dS



**Figure S12d - Nuclear Complexes and Selection on Human Genes.** B) The mean dN/dS ratio for human-mouse orthologs is plotted against the number of unique proteins for each human nuclear complex. Complexes with more unique proteins tend to have significantly smaller mean dN/dS ratios than those with less unique proteins (t-test:  $P < 0.003$ ). D) The dN/dS ratio for human-mouse orthologs is plotted against the number of nuclear complexes they participate in. Proteins participating in more complexes have significantly lower dN/dS ratios than those with less complex participation (t-test:  $P < 1.9 \times 10^{-6}$ ). A,C) Such trends were seldomly observed for model 1 randomly-generated complexes.

## Non-Nuclear Complexes and dN/dS



**Figure S12e - Non-Nuclear Complexes and Selection on Human Genes.** B) The mean dN/dS ratio for human-mouse orthologs is plotted against the number of unique proteins for each human non-nuclear complex. Complexes with more unique proteins tend to have significantly smaller mean dN/dS ratios than those with less unique proteins (t-test:  $P < 4.1 \times 10^{-6}$ ). D) The dN/dS ratio for human-mouse orthologs is plotted against the number of non-nuclear complexes they participate in. Proteins participating in more complexes have significantly lower dN/dS ratios than those with less complex participation (t-test:  $P < 1.3 \times 10^{-8}$ ). A,C) Such trends were seldomly observed for model 1 randomly-generated complexes.