Supplementary Information for:

Synthesis and Evaluation of a Series of Conjugatable 1,2,4,5-Tetrazines for Bioorthogonal Conjugation

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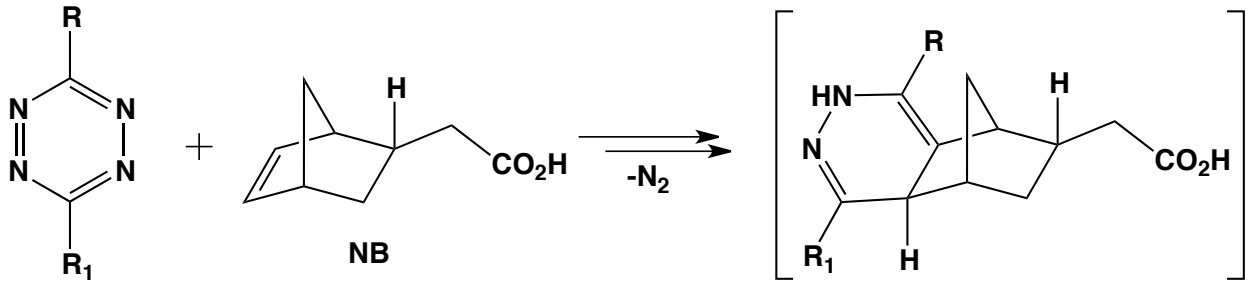

Scheme S1. General cycloaddition reaction of (1S,2S,4S)-bicyclo[2.2.1]hept-5-en-2-ylacetic acid (Norbornene, NB) with a generic 1,2,4,5-tetrazine showing one representative reaction product isomer.

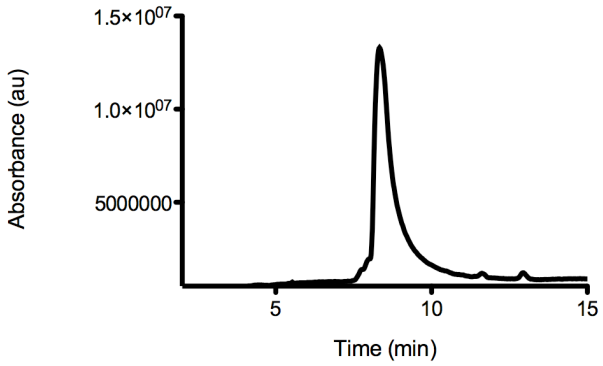

Figure S1. HPLC trace of purified AF750-**6** using a 0-100% buffer B gradient over 15 min.

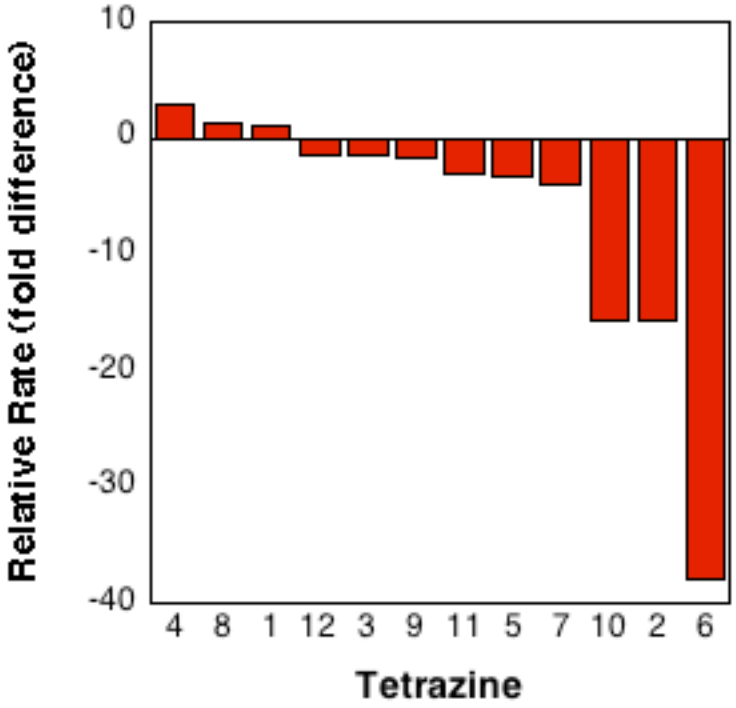

Figure S2. Relative second order [4+2] cycloaddition rate constants to **1** (4-(1,2,4,5-tetrazin-3-yl)phenyl)methanamine (k_2 = 1) of tetrazines **1-12** with NB in PBS at 37°C.

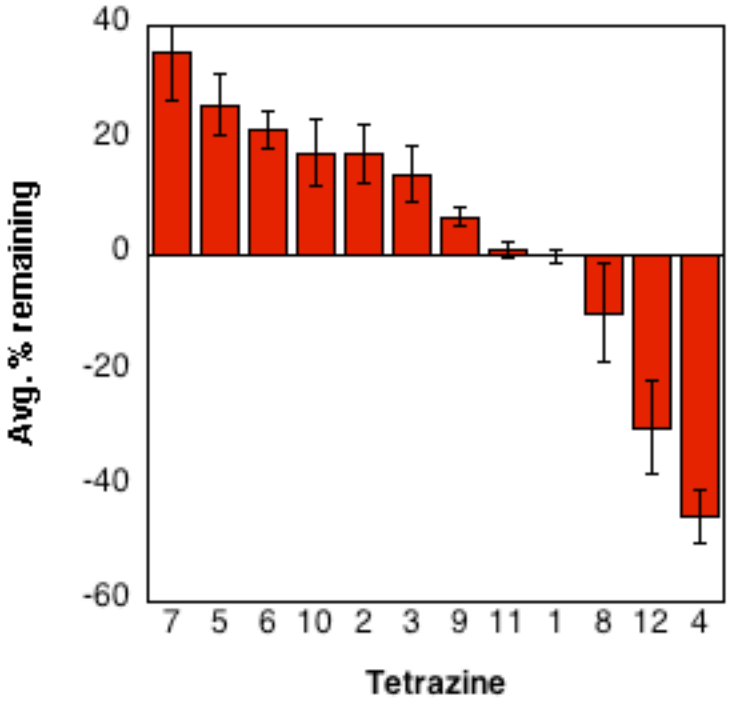

Figure S3. Relative tetrazine stability in relation to **1** in PBS after 14 h at 37°C.