

**Supplementary Figure 1.** Extracted ion chromatograms (XICs) showing the relative abundance of unlabeled (top) and SATA-labeled (bottom) peptide pairs from BSA. The labeled BSA was digested by trypsin and peptides were analyzed by LC-LTQ MS. All possible unlabeled forms of the peptides (top) were observed with <1% or <2% based on the peak area comparison between top and bottom panels.

**Supplementary Figure 2.** The guanidination of lysine  $\varepsilon$ -amine. The pKa of lysine is ~10.5. At pH ~10.5, lysine  $\varepsilon$ -amine serves as a nucleophile to react with *O*-methylisourea.