

## Supporting Information

# Cysteine [2,4] Disulfide Bond as a New Modifiable Site of $\alpha$ -Conotoxin TxIB

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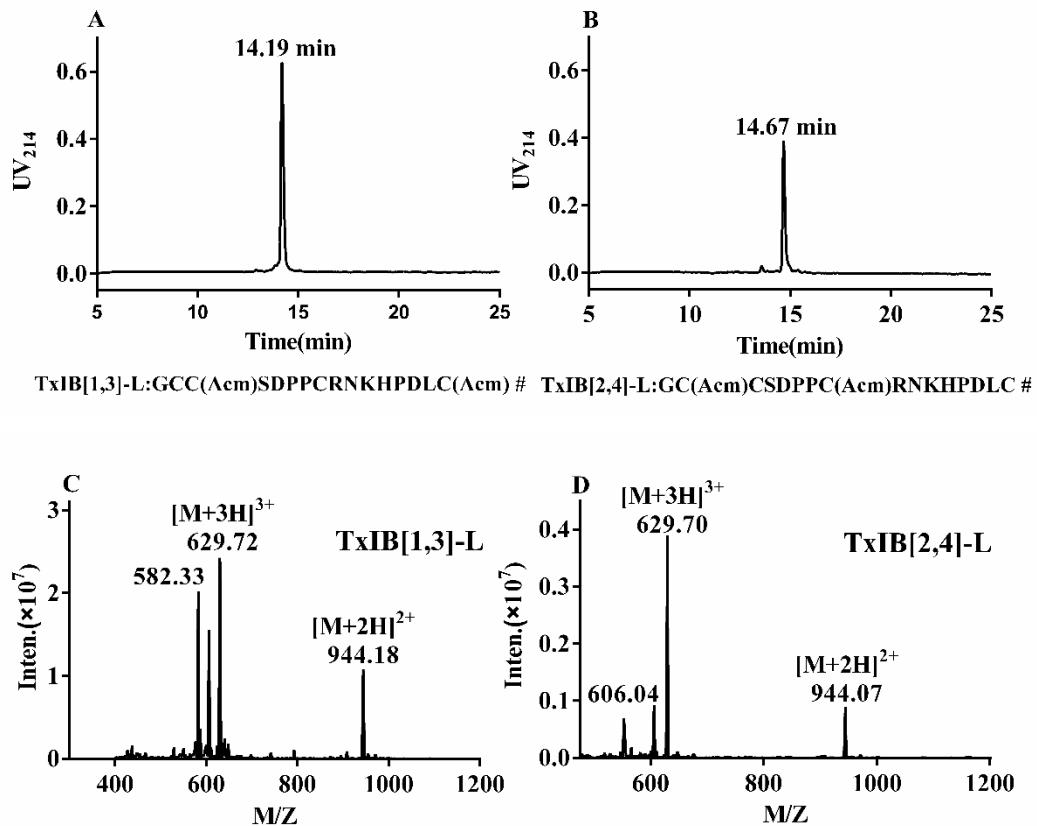
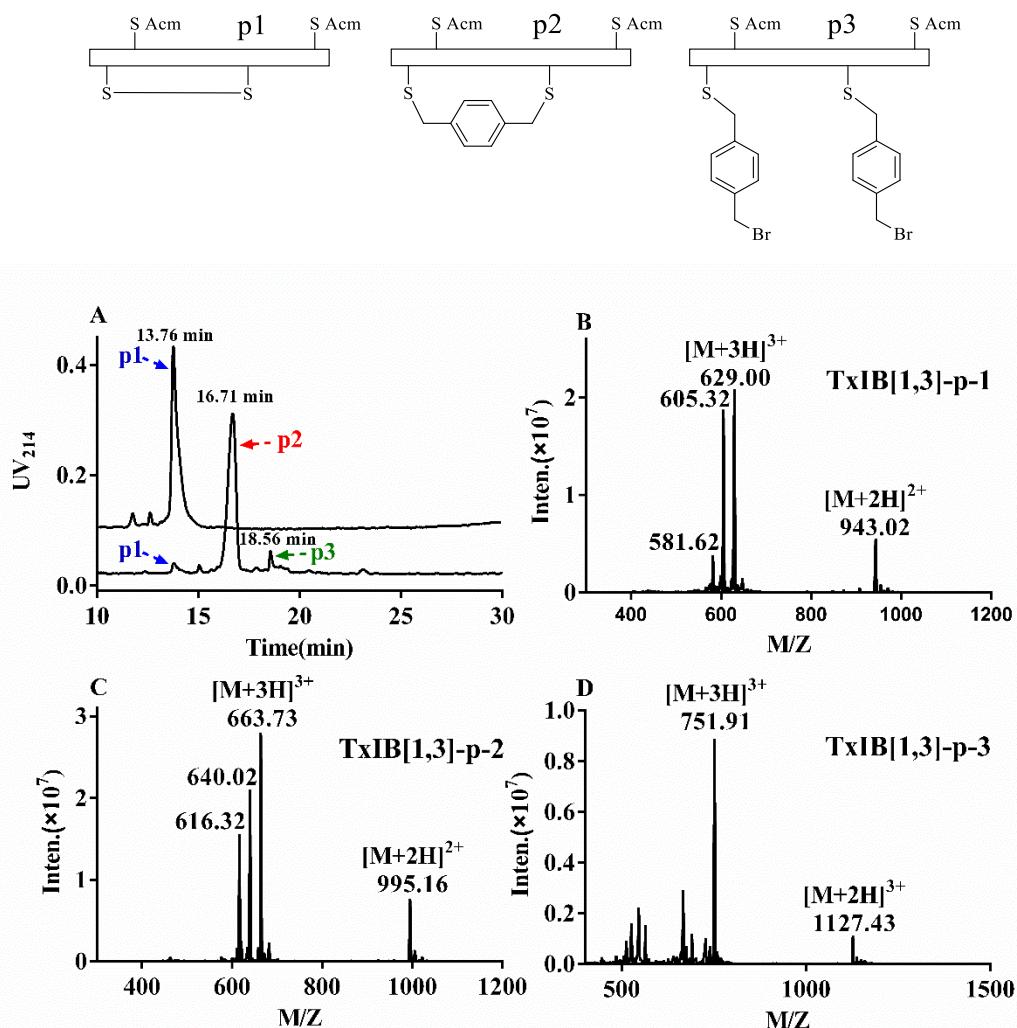
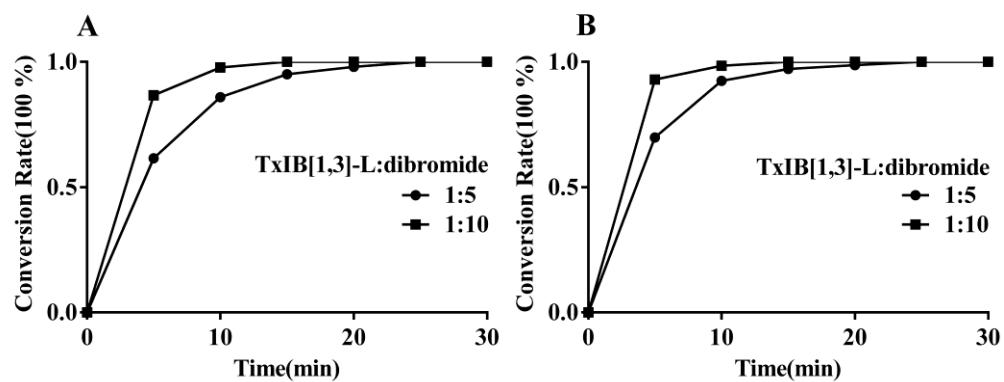


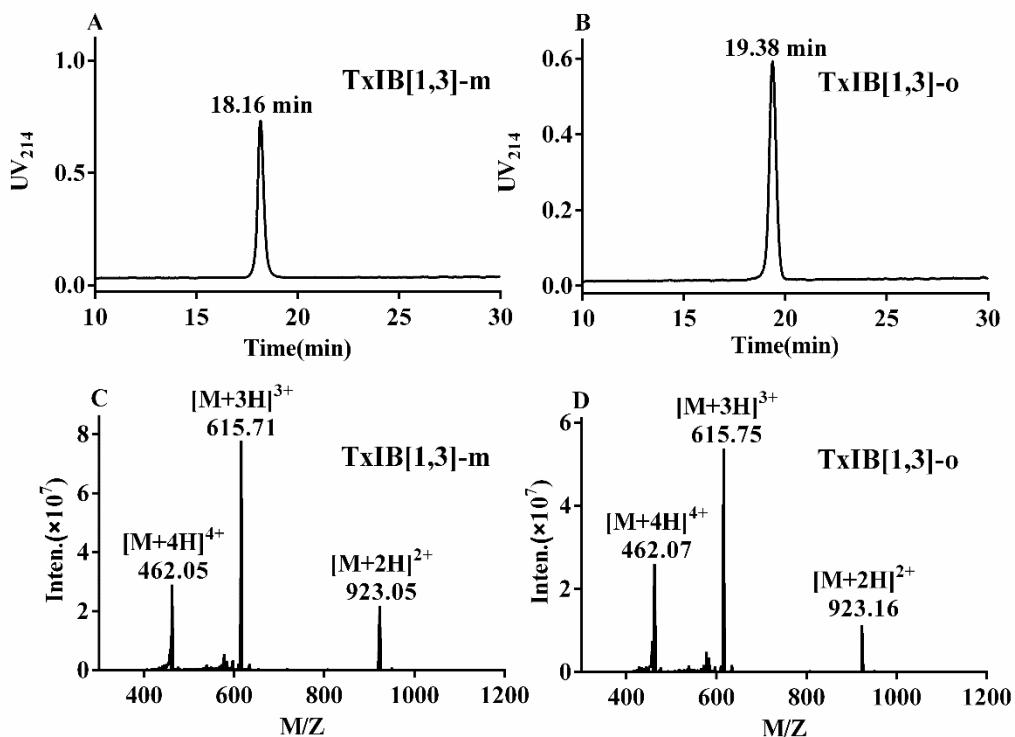
Figure S1. HPLC chromatogram and mass spectra of the linear peptides TxIB[1,3] and TxIB[2,4].



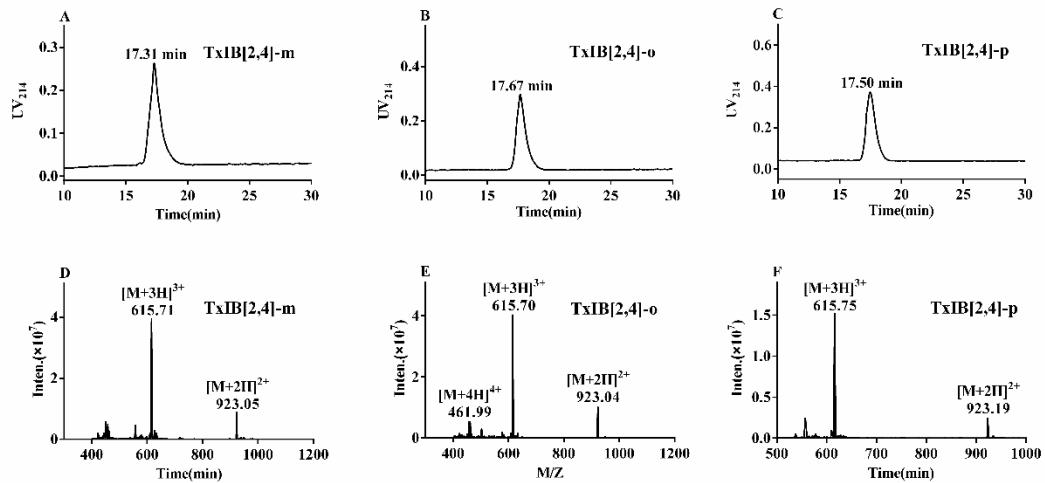
**Figure S2.** HPLC chromatogram and MS spectra of the linear peptide TxIB[1,3] reaction with p-xylene dibromide.



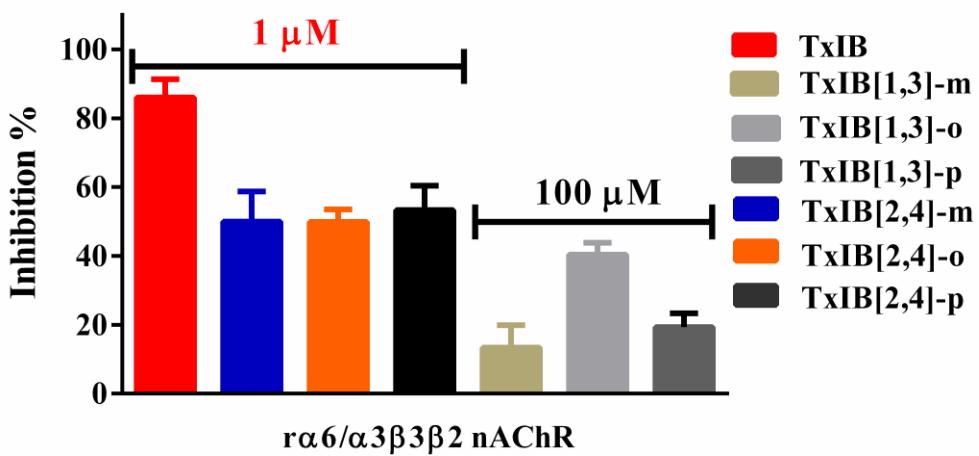
**Figure S3.** A. The conversion rate of TxIB[1,3]-L with m-xylene dibromide. B. The conversion rate of TxIB[1,3]-L with o-xylene dibromide.



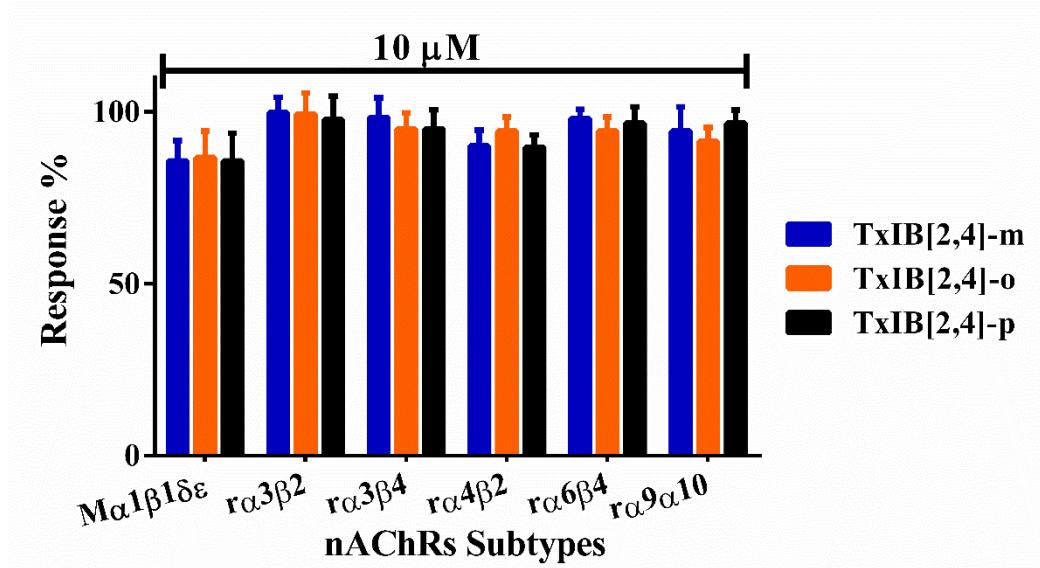
**Figure S4.** HPLC chromatogram and MS spectra of TxIB[1,3]-m and TxIB[1,3]-o.



**Figure S5.** HPLC chromatogram and MS spectra of TxIB[2,4]-m, o, p.



**Figure S6.** The inhibition of TxIB and its analogs on nAChR subtypes expressed in *Xenopus laevis* oocytes. TxIB and [2,4] modified analogs were determined at the concentration of 1  $\mu$ M and [1,3] modified products were determined at the concentration of 100  $\mu$ M on  $\alpha$ 6/α3β3β2 nAChR. All data represent mean  $\pm$  S.E.M, n = 3–6.



**Figure S7.** The response histogram of TxIB[1,3]-m, o, p on M $\alpha$ 1 $\beta$ 1 $\delta$  $\varepsilon$ , α3β2, α3β4, α4β2, α6/α3β4 and α9α10 nAChR subtypes. All data represent mean  $\pm$  S.E.M, n = 3–6.