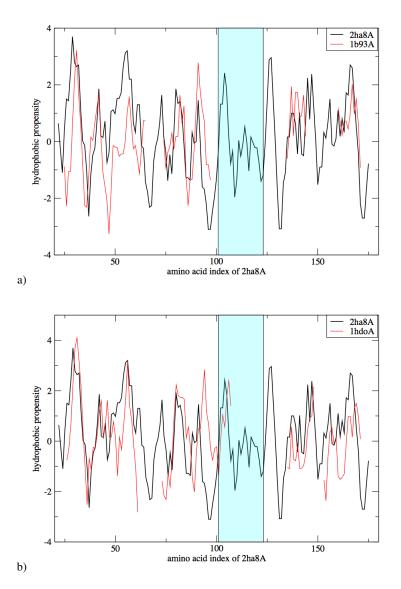
## Supporting Information - figure S2 "Knotted vs. unknotted proteins: evidence of knot-promoting loops"

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Hydrophobicity profiles for the knotted protein 2ha8A and the MISTRAL structurally-matching amino acids of the unknotted protein 1b93A (panel a) and 1hdoA (panel b). The knot-promoting segment (101–123) is highlighted by the light blue box. The hydrophobicity was calculated using the Kyte and Doolittle scale and an averaging window of 5 amino acids. Notice that, at variance with the case in panel a, the MISTRAL alignment of 2ha8A and 1hdoA, panel b, is non-sequential (see diagram in Fig. 3 of our paper) and shows two gaps, one of which is the knot-promoting segment.