

Foldamers: expanding the chemical space

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Our group has developed helical foldamers – oligomers that adopt stable helical folded conformations – derived from aromatic amino acids.^[1] Some of these folded objects have shown unprecedented conformational stability,^[2] and constitute convenient building blocks to elaborate synthetic, very large (protein-sized) folded architectures (Figure 1).^[3] They possess a high propensity to assemble into double, triple and quadruple helices.^[4] Cavities can be designed within such synthetic molecules that enable them to act as artificial receptors^[5] including for chiral guests. Water soluble analogues of these foldamers show promise in nucleic acid recognition.^[6]

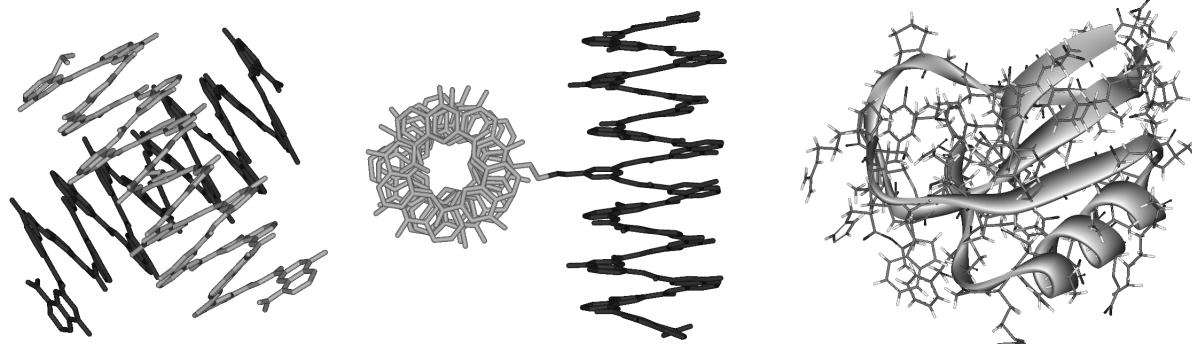


Figure 1. Crystal structure of a large foldamer comprised of two helices of opposite handedness at a 90° angle. The protein crystal structure on the right is shown as the same scale for size comparison.

References

1. S. Hecht, I. Huc (Eds), *Foldamers: Structure, Properties, and Applications*, **2007**, Wiley-VCH, Weinheim, ISBN: 978-3-527-31563-5.
2. H. Jiang, J.-M. Léger, I. Huc, *J. Am. Chem. Soc.* **2003**, *125*, 3448; N. Delsuc, T. Kawanami, J. Lefeuvre, A. Shundo, H. Ihara, M. Takafuji, I. Huc *ChemPhysChem* **2008**, *9*, 1882.
3. C. Dolain, J.-M. Léger, N. Delsuc, H. Gornitzka, I. Huc *Proc. Natl. Acad. Sci. USA* **2005**, *102*, 16146; N. Delsuc, J.-M. Léger, S. Massip, I. Huc *Angew. Chem. Int. Ed.* **2007**, *46*, 214; D. Sánchez-García, B. Kauffmann, T. Kawanami, H. Ihara, M. Takafuji, M.-H. Delville, I. Huc, *J. Am. Chem. Soc.* **2009**, *131*, 8642.
4. Q. Gan, C. Bao, B. Kauffmann, A. Grélard, J. Xiang, S. Liu, I. Huc, H. Jiang, *Angew. Chem. Int. Ed.* **2008**, *47*, 1715; D. Haldar, H. Jiang, J.-M. Léger, I. Huc, *Angew. Chem. Int. Ed.* **2006**, *45*, 5483; Y. Ferrand, A. Kendhale, J. Garric, B. Kauffmann, I. Huc, *Angew. Chem. Int. Ed.* **2010**, *49*, 1718.
5. Bao, B. Kauffmann, Q. Gan, K. Srinivas, H. Jiang; I. Huc *Angew. Chem. Int. Ed.* **2008**, *47*, 4153; Y. Ferrand, A. M. Kendhale, B. Kauffmann, A. Grélard, C. Marie, V. Blot, M. Pipelier, D. Dubreuil, I. Huc, *J. Am. Chem. Soc.* **2010**, *132*, 7858.
6. P. S. Shirude, E. R. Gillies, S. Ladame, F. Godde, K. Shin-ya, I. Huc, S. Balasubramanian, *J. Am. Chem. Soc.* **2007**, *129*, 11890.