Origin of Regular Chiral Fluctuation or Symmetry Breaking Unique to Preferential Enrichment

Rui Tamura, Japan, Kyoto University

Preferential enrichment (PE) is the first spontaneous enantiomeric resolution phenomenon that originates from regular chiral fluctuation or symmetry breaking and can apply to racemic crystals.^{1,2} The mechanism of PE has been interpreted in terms of a nonlinear complexity phenomenon including two unique processes: i) a solvent-assisted solid-to-solid polymorphic transition^{1,3} and ii) subsequent selective redissolution of the excess one enantiomer from the transformed disordered crystals into the mother liquor.^{1,4} Furthermore, recently we have demonstrated that PE is applicable to common racemic crystals if the four requirements based on the mechanism are satisfied.^{1,5}

Here I talk about the concept, properties, mechanism and requirements with respect to PE, particularly focusing on (1) the *in situ* XRD and microscopic observation of polymorphic transition,³ (2) the kinetic and thermodynamic origin of chiral symmetry breaking,⁴ (3) the ternary phase diagram consistent with the mechanism,⁴ and (3) the practical application of PE to the racemic crystals of common neutral or basic amino acids and chiral drugs, and their racemic cocrystals with achiral dicarboxylic acids, etc.⁵

References

1) Reviews: CrystEngComm 2011, 13, 5296; Symmetry 2010, 2, 112; Top. Curr. Chem. 2007, 269, 53.

2) Angew. Chem. Int. Ed. **1996**, *35*, 2372 & **1998**, *37*, 2876; J. Am. Chem. Soc. **2002**, *124*, 13139; Cryst. Growth Des. **2003**, *3*, 973; Chem. Eur. J. **2006**, *12*, 3515; Cryst. Growth Des. **2007**, *7*, 1643 & **2008**, *8*, 540.

3) Cryst. Growth Des. 2015, 15, 3052 & 2017, 17, 671.

4) Chem. Eur. J. 2016, 22, 11660.

5) Cryst. Growth Des. **2010**, *10*, 2668 & **2011**, *11*, 607; Chem. Commun. **2012**, *48*, 2791; Chem. Eur. J. **2014**, *20*, 10343; CrystEngComm **2014**, *16*, 5811; Chirality **2015**, *27*, 405.