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“Carbohydrates and Sulfur Compounds: Smart Tools in the Enantioselective Synthesis of Therapeutically Relevant Molecules”

Abstract:

The importance of enantiopure compounds in different areas such as agriculture, fragrance and medicine is currently well recognized. As illustrative data, at present more than 50% of the drugs in the market are chiral compounds, and about 80% of new drugs patented are single enantiomers.¹ Therefore there is a strong demand for the development of new and effective asymmetric synthesis methods. Catalysis can be considered as the most efficient method in asymmetric synthesis since it can be applied to a wider range of transformations than any other enantioselective synthesis method, and ligand design has played a central role for the development of efficient metal and organo-catalyzed processes. In this sense, sulfinyl-based ligands present indubitable advantages for their applications in asymmetric catalysis. They are air, oxygen and moisture stable, and nowadays a number of highly efficient approaches allow the rapid synthesis of both enantiomers of sulfinyl-based ligands and to easily modulate their structure.² In this sense, our research centers on the asymmetric synthesis of enantiopure sulfinyl derivatives of synthetical and biological interest, having developed a general and enantiodivergent approach for the synthesis of enantiopure sulfinyl derivatives, using diacetone-D-glucose as the unique source of chirality.³

With these bases, in the first part of the talk I will show our approach to the design and synthesis of a number of C1 and C2 symmetric bidentate sulfinyl-based ligands, and their applications in different enantioselective organic and organometallic catalytic processes.⁴

In the second part, I will present our approach to the development of new lead compounds with antitumor activities, based on the NK1 receptor⁵ and the Nrf2 activation factor as therapeutic targets.⁶

References:

- 1 a) L.A. Nguyen, H. He, Ch. Pham-Huy, Int. J. Biomed. Sci. 2006, 2, 85–100.
- 2 T. Jia, M. Wang, J. Liao, Top Curr Chem (Z) 2019, 377: 8. b) G. Sipos, E. E. Drinkel, R. Dorta, Chem. Soc. Rev. 2015, 44, 3834-3860. c) I. Fernandez, N. Khiar, 2008, Asymmetric catalysis using sulfoxides as ligands. In T. Toru and C. Bolm (Eds.) Organosulfur Chemistry in Asymmetric Synthesis, pp. 265-290. Weinheim, Wiley-VCH. d) I. Fernandez, N. Khiar, Chem. Rev., 2003, 103, 3651.

- 3 a) I. Fernandez, N. Khiar, J. M. Llera, F. Alcudia, *J. Org. Chem.*, 1992, 57, 6789. b) N. Khiar, F. Alcudia, J.L. Espartero, L. Rodríguez, I. Fernández, *J. Am. Chem. Soc.*, 2000, 122, 7598–7599.
4 a) N. Khiar, A. Salvador, V. Valdivia, A. Chelouan, A. Alcudia, E. Álvarez, I. Fernández, *J. Org. Chem.* 2013, 78, 6510–6521. b) N. Khiar, A. Salvador, A. Chelouan, A. Alcudia, I. Fernandez, *Org. Biomol. Chem.*, 2012, 10, 2366. c) N. Khiar, V. Valdivia, A. Salvador, A. Chelouan, A. Alcudia, I. Fernandez, *Adv. Synth. Catal.*, 2013, 355, 1303. d) V. Valdivia, I. Fernandez and N. Khiar, *Org. Biomol. Chem.*, 2014, 12, 1211. e) A. Chelouan, R. Recio, L.G. Borrego, E. Alvarez, N. Khiar, I. Fernández, *Org. Letters*, 2016, 18, 3258-3261. f) I. Fernández, A. Alcudia, B. Gori, V. Valdivia, R. Recio, N. Khiar, *Org. Biomol. Chem.* 2010, 8, 4388-4393. g) L.G. Borrego, R. Recio, M. Alcaranza, N. Khiar, I. Fernandez, *Adv. Synth. Catal.* 2018, 360, 1273-1279.
5 a) R. Recio, E. Vengut, B. Mouillac, H. Orcel, M. López Lázaro, E. Álvarez, N. Khiar, I. Fernández, *Eur. J. Med. Chem.* 2017, 138, 644-660. b) N. Khiar, I. Fernández, R. Recio. PCT/ES2013/070124, WO2013132124A1.
6 a) R. Recio, E. Elhalem, J.M. Benito, I. Fernandez, N. Khiar, *Carbohydr. Polym.* 2018, 187, 118-125. b) E. Elhalem, R. Recio, S. Werner, F. Lieder, J.M. Calderón, M. López-Lázaro, I. Fernández, N. Khiar, *Eur. J. Med. Chem.* 2014, 87, 552-563. c) N. Khiar, S. Werner, S. Mallouk, F. Lieder, A. Alcudia, I. Fernandez, *J. Org. Chem.* 2009, 74, 6002-6009. d) N. Khiar, I. Fernández, R. Recio, M. López-Lázaro, J. M. Calderón-Montaña. PCT/ES2016/070383, WO2016189179A1.