

Biosynthesis of Roussoellatide from the Marine-Derived Fungus *Roussoella* sp.

Everton L. F. Ferreira¹, Laura P. Ióca¹, Mario F. C. Santos¹, Antonio G. Ferreira², David E. Williams³, Raymond J. Andersen³, Brian O. Patrick³, Luciana M. Elias⁴, Simone P. Lira⁴, Michel Z. R. Passarini⁵, Lara D. Sette⁵, Roberto G. S. Berlinck¹

¹Instituto de Química de São Carlos, Universidade de São Paulo, CP 780, CEP 13560-970, São Carlos, SP Brazil; ²Universidade Federal de São Carlos, São Carlos, Brazil; ³University of British Columbia, Vancouver, Canada; ⁴Departamento de Ciências Exatas, Escola Superior de Agricultura Luiz de Queiroz, Universidade de São Paulo, Piracicaba – SP, Brazil; ⁵Divisão de Recursos Microbianos, Centro Pluridisciplinar de Pesquisas Químicas, Biológicas e Agrícolas, Universidade Estadual de Campinas, Paulínia, SP, Brazil e Departamento de Bioquímica e Microbiologia, Instituto de Biociências, Universidade Estadual Paulista “Júlio de Mesquita Filho”, Campus Rio Claro, Rio Claro, SP, Brazil;

evertonleandro28@hotmail.com

Abstract: During our program for the discovery of new bioactive and structurally unique metabolites from fungi obtained at various environments, roussoellatide, a novel bis-chlorinated polyketide has been isolated from cultures of the marine-derived fungus *Roussoella* sp. DLM33. The structure and absolute configuration of roussoellatide have been established by analysis of spectroscopic data and X-ray diffraction analysis. The use of experimental design under different growth conditions, followed by chemometric analysis allowed us to establish the optimal growth conditions for producing roussoellatide by *Roussoella* sp., in order to investigate its biosynthesis. The intriguing polyketide skeleton of roussoellatide had its biosynthesis investigated by feeding experiments with [1-¹³C]acetate, [1,2-¹³C]acetate, and [*methyl*-¹³C]methionine. The results indicated the involvement of Favorskii rearrangements, as well as of an intermolecular Diels-Alder reaction, leading to the unique structure of roussoellatide.

Acknowledgments: To FAPESP (2013/50228-8) and CAPES (ou CNPq) for the financial support.