

SEPTEMBER
23-27TH
2024



BRAZILIAN MEETING
ON ORGANIC SYNTHESIS
BENTO GONÇALVES, RS - BRAZIL

A one-pot C-H functionalization protocol for the synthesis of (hetero)chalcones

Franco J. Caires^{1*}, Thais R. Arroio¹, Isabela W. F. Pinto¹, Luiz Vinicius S. de Oliveira¹ and Giuliano C. Clososki¹

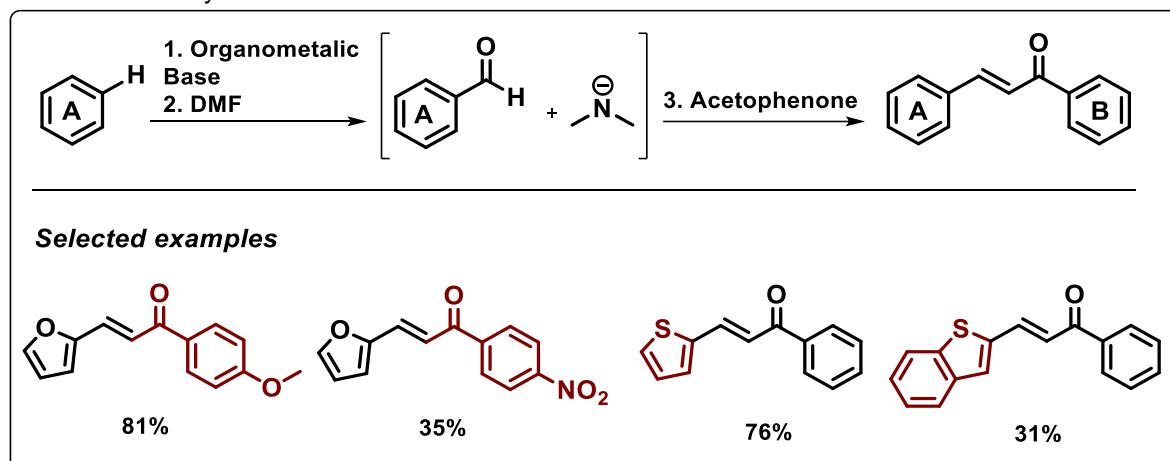
¹⁾ Department of Biomolecular Sciences, Faculty of Pharmaceutical Sciences of Ribeirão Preto, University of São Paulo, FCFRP-USP, 14040-903

*e-mail: fjcaires@usp.br

Keywords: metalation, chalcones, C-H functionalization.

ABSTRACT

Chalcones comprehend an interesting class of organic compounds which have been studied in several works.¹ They can be obtained from natural sources as important precursors of secondary metabolites and, not restricted to biosynthetic roles, chalcones can also be synthetized and employed in different fields such as optical devices² and medicinal chemistry, exhibiting a wide range of relevant biological activities.³ A variety of synthetic methodologies are available to provide chalcones like classical Claisen-Schmidt aldol condensation⁴ or more recent ones, using C-H activation.⁵ In our study, a number of (hetero)aromatic chalcones displaying interesting drug-like structural features could be synthesized in a one-pot approach, using a novel organometallic base-mediated protocol, that explores a direct C-H functionalization of readily available aromatic and heterocyclic substrates.



ACKNOWLEDGEMENTS

The authors gratefully acknowledge financial support for this work by the Brazilian foundations Fundação de Amparo à Pesquisa do Estado de São Paulo (FAPESP), Conselho Nacional de Desenvolvimento e Tecnológico (CNPq) and Coordenação de Aperfeiçoamento de Pessoal de Nível Superior (CAPES).

REFERENCES

1. Zhuang, C.; Zhang, W.; Sheng, C.; Zhang, W.; Xing, C.; Miao, Z. Chalcone: A Privileged Structure in Medicinal Chemistry. *Chem. Rev.* **2017**, *117*, 7762. <https://doi.org/10.1021/acs.chemrev.7b00020>
2. Custodio, J. M. F.; D'Oliveira, G. D. C.; Gotardo, F.; Cocca, L. H. Z.; De Boni, L.; Perez, C. N.; Maia, L. J. Q.; Valverde, C.; Osório, F. A. P.; Napolitano, H. B. Chalcone as Potential Nonlinear Optical Material: A Combined Theoretical, Structural, and Spectroscopic Study. *J. Phys. Chem. C* **2019**, *123*, 5931. <https://doi.org/10.1021/acs.jpcc.9b01063>.
3. Qin, H. L.; Zhang, Z. W.; Lekkala, R.; Alsulami, H.; Rakesh, K. P. Chalcone hybrids as privileged scaffolds in antimalarial drug discovery: A key review. *Eur. J. Med. Chem.* **2020**, *193*, 112215. DOI: 10.1016/j.ejmec.2020.112215
4. Yadav, G. D.; Wagh, D. P. Claisen-Schmidt Condensation using Green Catalytic Processes: A Critical Review. *ChemistrySelect*, **2020**, *5*, 9059. <https://doi.org/10.1002/slct.202001737>.
5. Shi, Y.; Xing, H.; Huang, T.; Liu, X.; Chen, J.; Guo, X.; Li, G. B.; Wu, Y. Divergent C-H activation synthesis of chalcones, quinolones and indoles. *Chem. Commun.* **2020**, *56*, 1585. <https://doi.org/10.1039/c9cc08926h>