

Synthesis of new *N*-confused triazole-porphyrin derivatives through the Heck procedure

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Keywords: N-confused porphyrins, triazoles, Heck reaction

INTRODUCTION

Porphyrins have already demonstrated significant applications in several areas; the one in Medicine (PDT-Photodynamic Therapy of cancer cells) is highly significant.¹ Over the last years, the efforts to develop new methodologies to convert porphyrins into new derivatives, with better structural and spectroscopic characteristics for becoming efficient PDT photosensitizers, have been outstanding.¹ One possibility consists in the coupling of porphyrins to compounds with well-established pharmacological activities.² The efficiency of triazoles as antibacterial, antifungal, antiviral and as antitumor agents is well established;³ so coupling porphyrins to triazoles may lead to new drugs. Recently the synthesis of novel porphyrin isomers, such as *N*-confused porphyrins, has attracted the attention of many chemists.⁴ Such compounds have energy absorption bands at wavelengths higher than 700 nm, a good feature for a compound to be used as a photosensitizer in PDT.¹ In this communication we will report our studies on the synthesis of novel N-confused triazole-porphyrin derivatives through the Heck procedure.

RESULTS AND DISCUSSION

Synthesis of *N*-confused tetraphenylporphyrin (NC-TPP) was achieved *via* condensation of pyrrole and benzaldehyde under methanesulfonic acid catalysis.⁵ The Heck reactions were carried out with the silver complex of the mono-brominated NC-TPP (1); such complex was prepared, in a good yield, by reaction of NC-TPP with NBS, followed by the silver complexation, (Scheme 1).



Scheme 1. Synthesis of mono-brominated *N*-confused tetraphenylporphyrin (1).

The Heck reactions of *N*-confused porphyrin **1** with vinyl-triazoles **2a-e** were performed in the presence of Pd(0) (Scheme 2). It was possible to isolate the

new *N*-confused triazole-porphyrin conjugates **3a-e**, in good yields.



Scheme 2. Heck reactions between *N*-confused porphyrin 1 and vinyl-triazoles 2a-e

CONCLUSION

In this work the Heck procedure has been used to prepare novel *N*-confused triazole-porphyrin conjugates in good yields. These derivatives might exhibit important biological activities, since they combine the photosensitizer properties of the *N*-confused porphyrin group with the biological activity of triazole moieties.

ACKNOWLEDGEMENTS

Thanks are due to Fundação para a Ciência e a Tecnologia (FCT, Portugal), European Union, QREN, FEDER and COMPETE for funding the QOPNA research unit (project PEst/QUI/UI0062/2011) and the Portuguese NMR Network. Thanks are also due to the Universities of Aveiro and Federal Fluminense and to the FCT/GRICES/CAPES collaborative programme for funding this work. ATP Gomes also thanks FCT for her post-doctoral grant (SFRH/BPD/79521/2011).

REFERENCES

¹ Kadish, K.M., Smith, K.M.; Guilard, R. *Handbook of Porphyrin Science*, World Scientific Publishing Company Co:Singapore, 2010, vol. 1-12.

² M. M. Bastos, A. T. P. C. Gomes, M. G. P. M. S. Neves, A. M. S. Silva, O. A. Santos-Filho, N. Boechat, J. A. S. Cavaleiro, *Eur. J. Org. Chem.*, **2013**, 1485-1493.

³ Genin, M.J.; Allwine, D.A.; Anderson, D.J.; Barbachyn, M.R.; Emmert, D.E.; Garmon, S.A.; Graber, D.R.; Grega, K.C.; Hester, J.B.; Hutchinson, D.K.; Morris, J.; Reischer, R.J.; Ford, C.W.; Zurenko, G.E.; Hamel, J.C.; Schaadt, R.D.; Stapert, D.; Yagi, B.H. *J. Med. Chem.* **2000**, *43*, 953-970.

⁴ (a)Maeda, H.; Furuta, H. *Pure Appl. Chem.*, 2006, 78, 29–44. (b)
Furuta, H.; Asano, T.; Ogawa, T. *J. Am. Chem. Soc.* 1994, *116*, 767-768.
⁵ Geier, G. R.; Haynes, D. M.; Lindsey, J. S. *Org. Lett.*, 1999, 1, 1455 -

⁵ Geier, G. R.; Haynes, D. M.; Lindsey, J. S. *Org. Lett.*, 1999, 1, 1455 - 1458.

15th Brazilian Meeting on Organic Synthesis – 15th BMOS – November 10-13, 2013 - Campos do Jordão, Brazil