

Synthesis of Chiral Nitrogen Heterocycles *via* the Narasaka-Heck Reaction

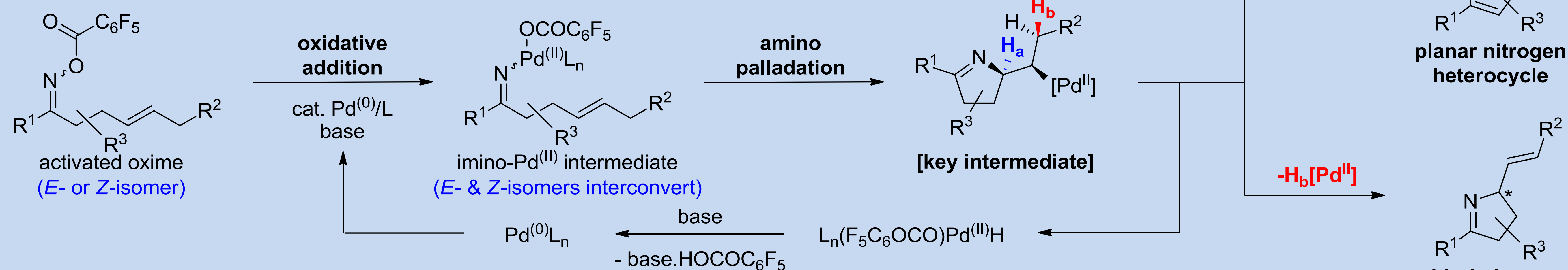


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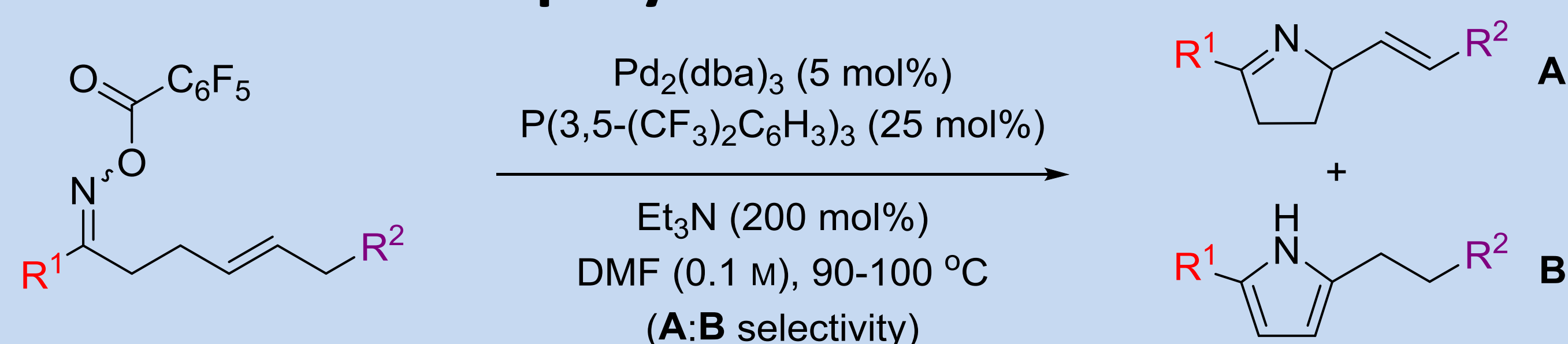
Introduction

- The Narasaka-Heck reaction has been applied to the synthesis of a range of achiral nitrogen heterocycles (e.g. pyrroles, pyridines, imidazoles)⁽¹⁾

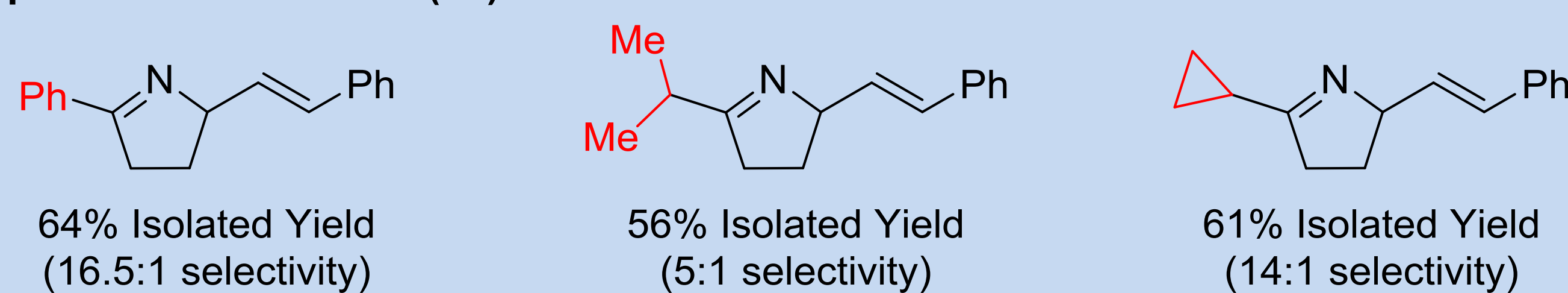


- Work in the Bower group is extending this reaction towards the formation of chiral nitrogen heterocycles⁽²⁾
- In this study presented, cyclisation onto 1,2-disubstituted alkenes affords the key intermediate shown – elimination of H_b delivers the chiral product⁽³⁾

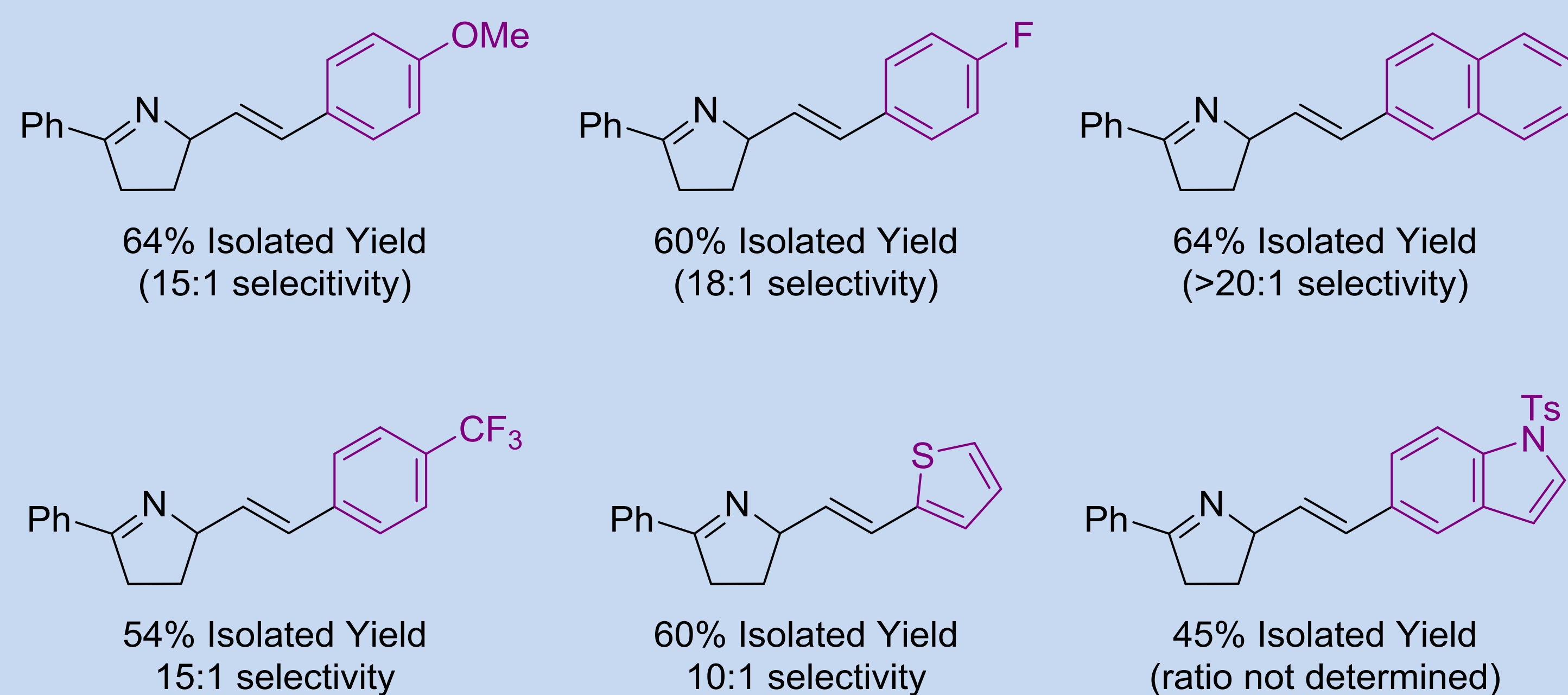
Substrate controlled β -hydride elimination



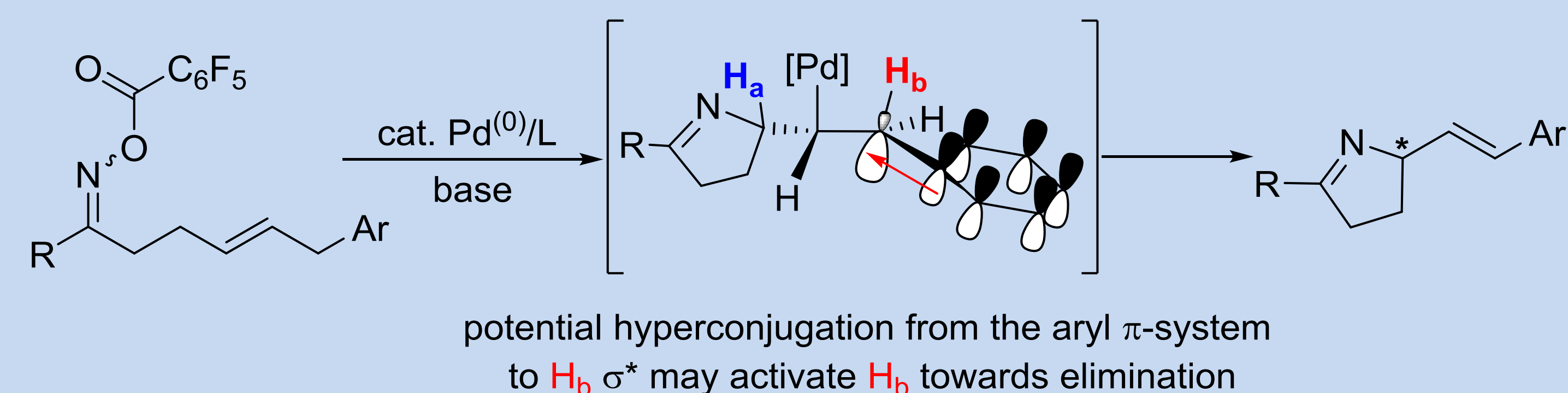
Scope of the oxime ester (R^1)



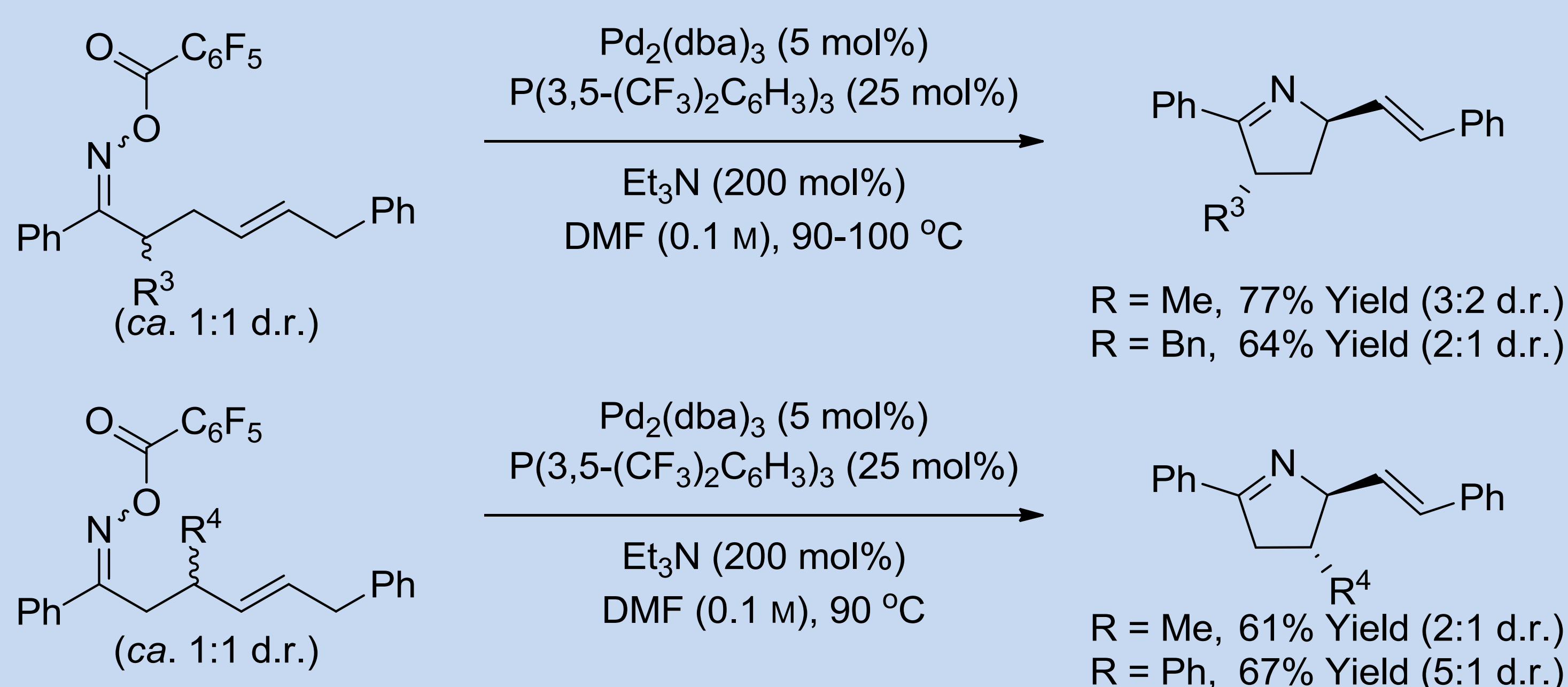
Scope of the aryl group (R^2)



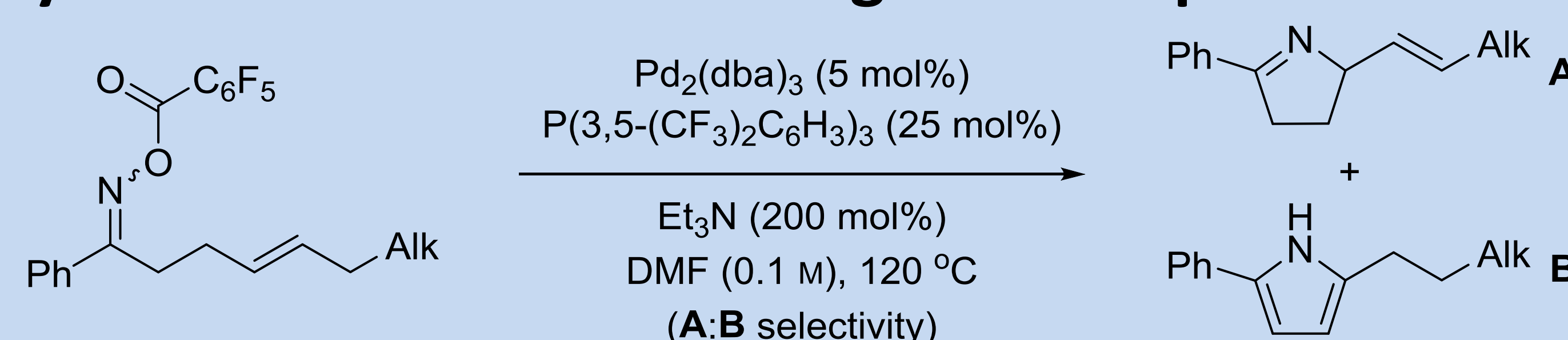
Aryl directed β -hydride elimination?⁽⁴⁾



Diastereoselectivity in the cyclisation

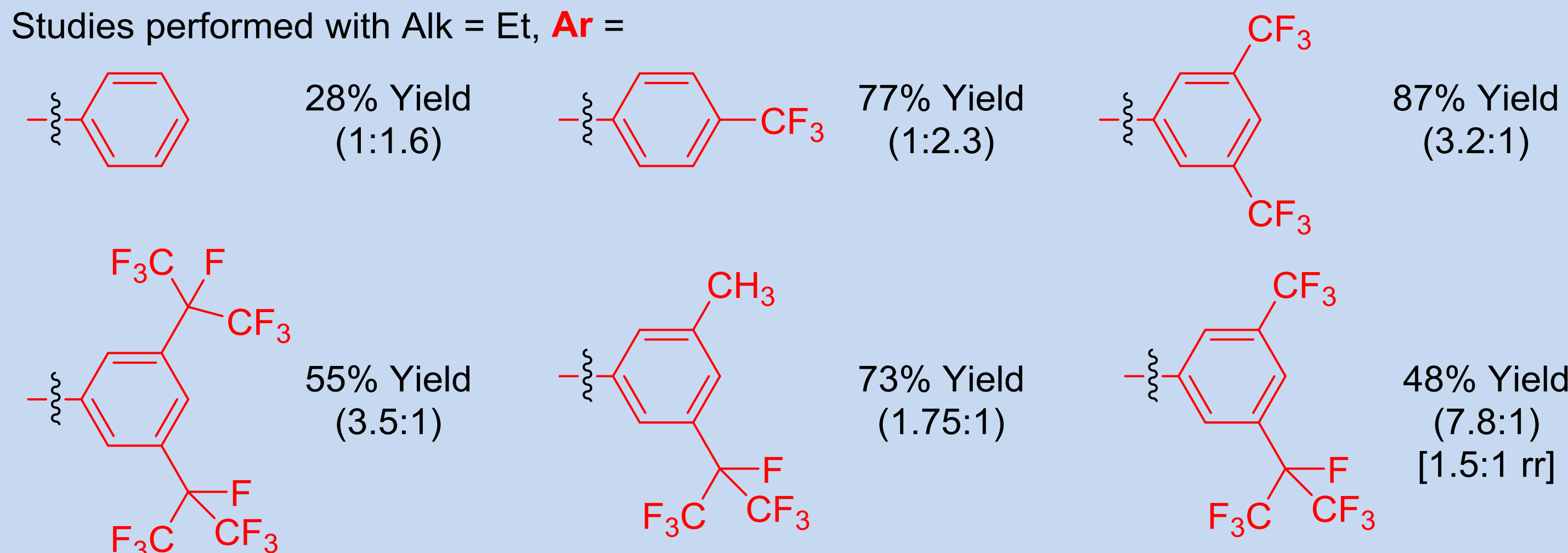


Alkyl-substituted alkenes also give chiral products

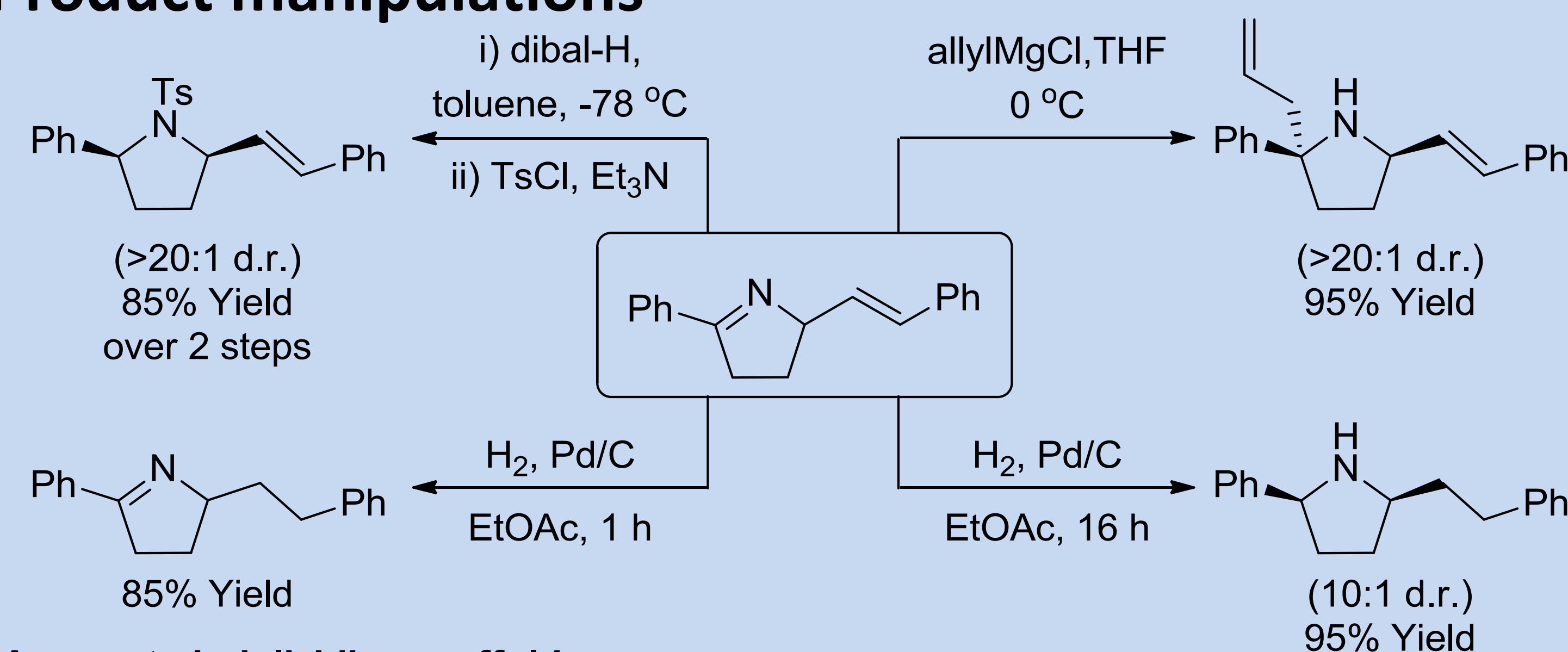


Choice of ligand (PAr_3) affects product ratio:

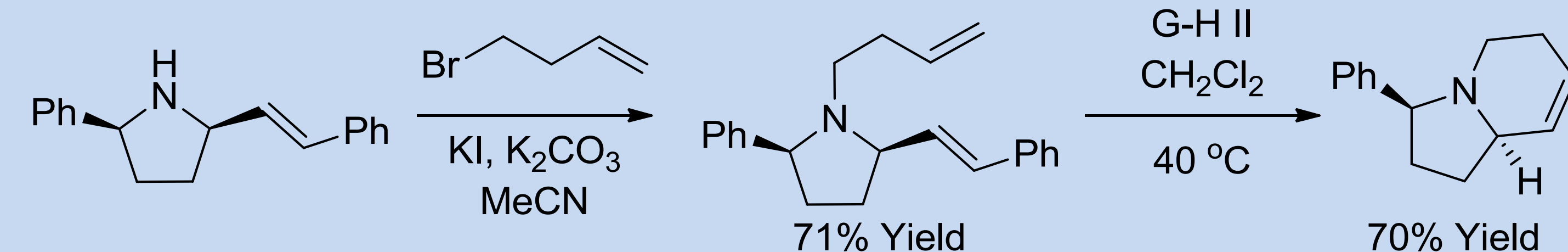
Studies performed with Alk = Et, Ar =



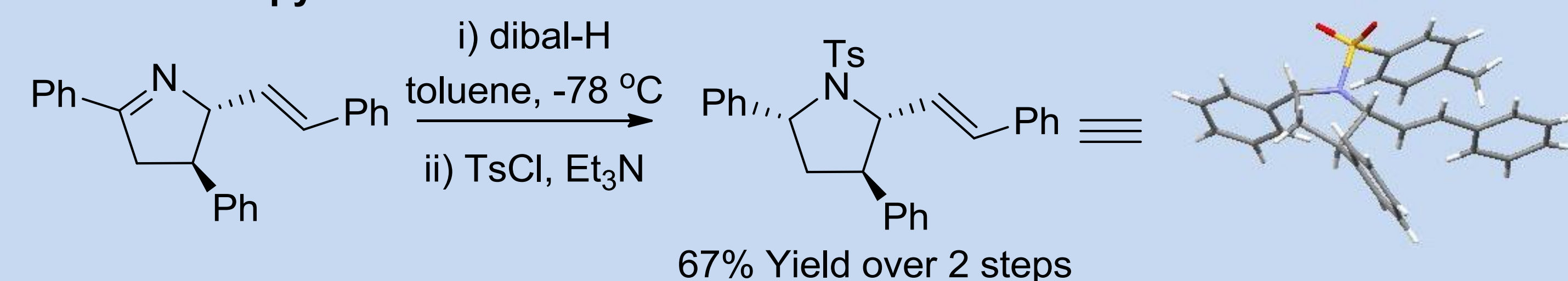
Product manipulations



Access to indolizidine scaffolds:



Trisubstituted pyrrolidine:



Acknowledgements

Bristol Chemical Synthesis Centre for Doctoral Training and the University of Bristol, for the provision of PhD Studentship. The SCl is acknowledged for the award of a Messel Scholarship (NJR).

References

- (1) Kitamura, M.; Narasaka, K. *Chem. Rec.* **2002**, 2, 268 (2) a) Faulkner, A.; Bower, J. F. *Angew. Chem. Int. Ed.* **2012**, 51, 1675 b) Faulkner, A.; Scott, J. S.; Bower J. F. *Chem. Commun.* **2013**, 49, 1521
- (3) Race, N. J.; Bower, J. F. *Org. Lett.* **2013**, 15, 4616 (4) Werner, E. W.; Sigman, M. S. *J. Am. Chem. Soc.* **2011**, 133, 9692