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Palladium-Catalyzed Ligand-Directed C–H Functionalization Reactions

Thomas W. Lyons and Melanie S. Sanford*

Department of Chemistry, University of Michigan, 930 North University Avenue, Ann Arbor, Michigan 48109

Reporter: Yi Zhang

Superior: Prof. Yong Huang

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Introduction:



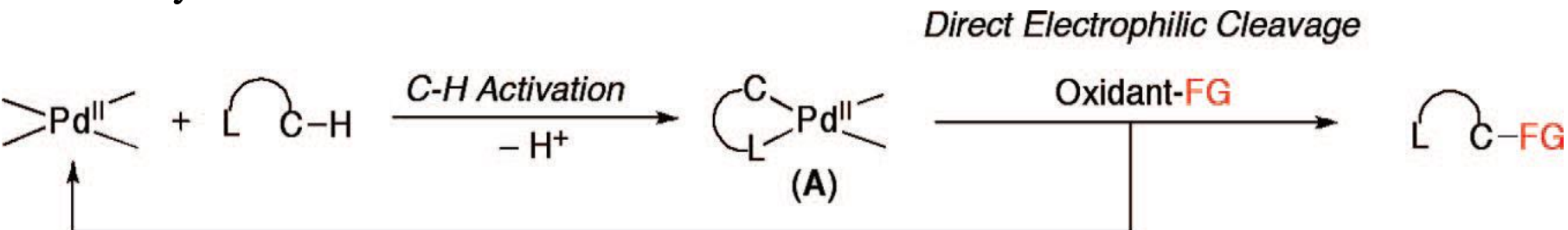
Thomas Lyons was born in Ottawa, IL, in 1983. He received his B.S. degree with honors in Chemistry from DePaul University in 2005 under the mentorship of Professor Matthew Dintzner. He is currently a Ph.D. candidate in Professor Melanie Sanford's research laboratory, where he is studying regioselective oxidative coupling reactions.



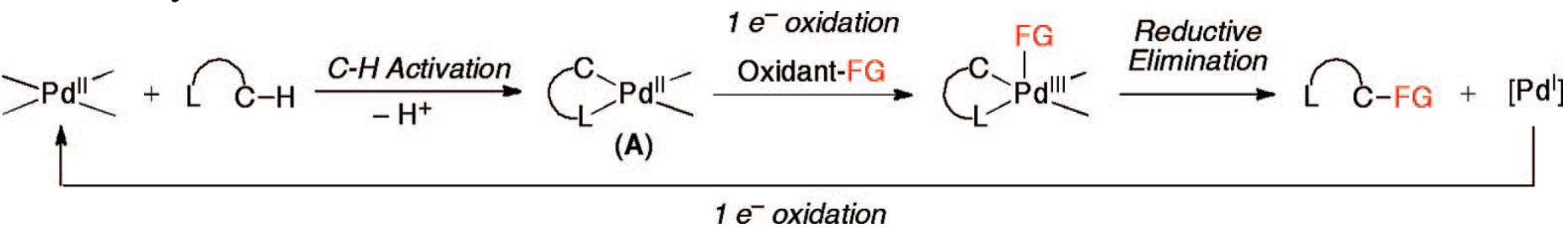
Melanie Sanford received her B.S. and M.S. degrees at Yale University, where she carried out undergraduate research in the laboratory of Professor Robert Crabtree. She pursued graduate studies at the California Institute of Technology working with Professor Robert Grubbs. Following postdoctoral work at Princeton University with Professor John Groves, she joined the faculty at the University of Michigan in the summer of 2003 as an Assistant Professor of Chemistry. In spring 2007 she was promoted to her current position of Associate Professor of Chemistry. She has been recognized with a number of awards, including a Presidential Early Career Award in Sciences and Engineering, an Arthur Cope Scholar award from the American Chemical Society, and the BASF catalysis award. Research in the Sanford group focuses broadly on the development and mechanistic study of new transition-metal-catalyzed reactions for applications in organic synthesis. The group is working to develop a diverse set of transformations for the direct conversion of unactivated carbon–hydrogen bonds into new functional groups with high levels of chemo-, regio-, and stereoselectivity.

Electrophilic Functionalization Pathway

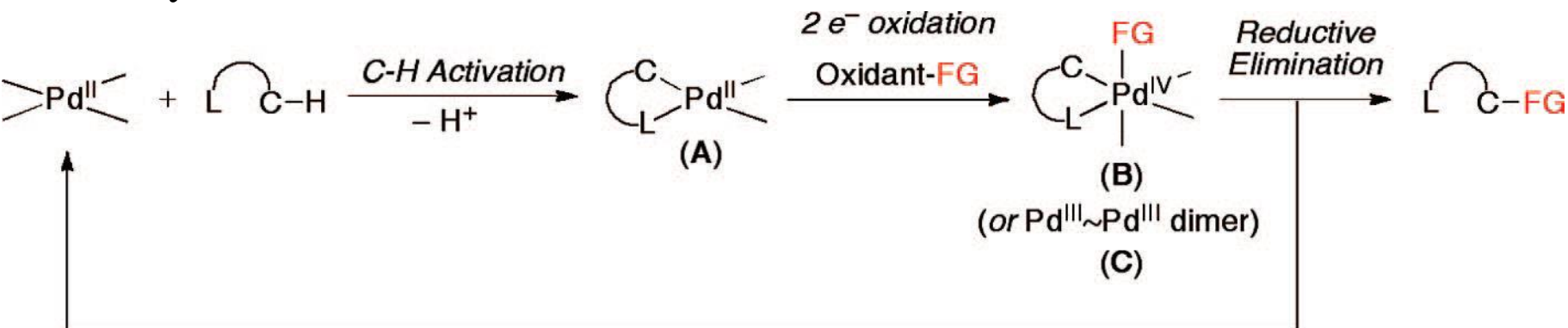
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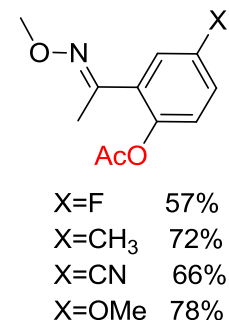
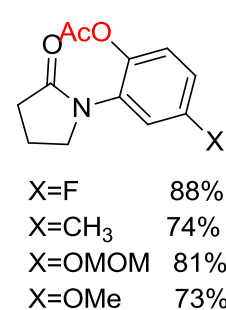
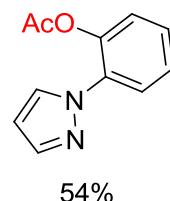
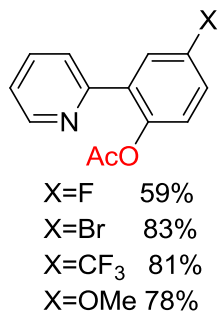
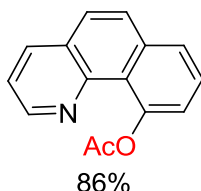
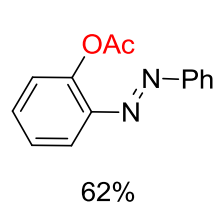
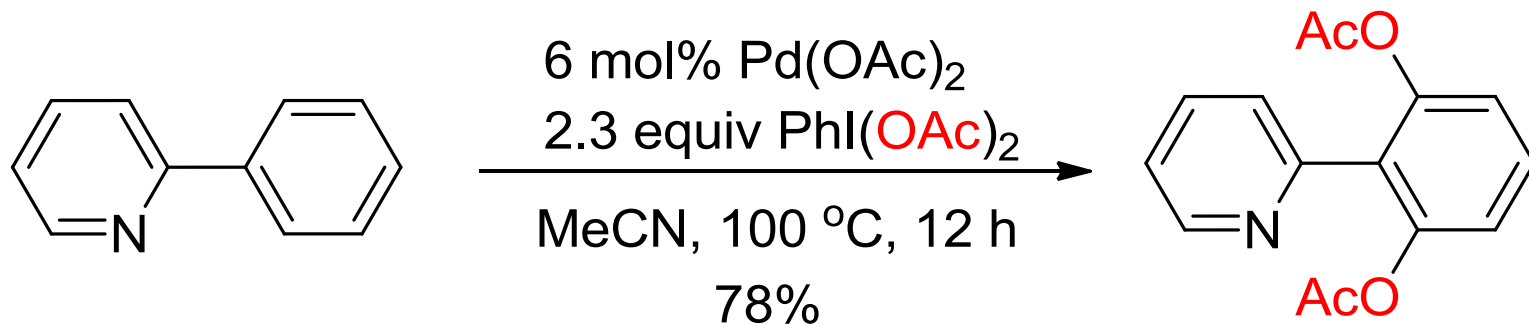
Pathway B:



Pathway C:



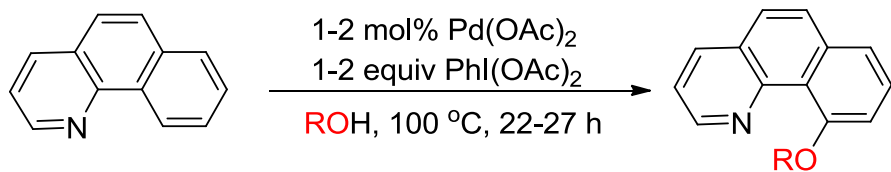
Acetoxylation with $\text{PhI}(\text{OAc})_2$



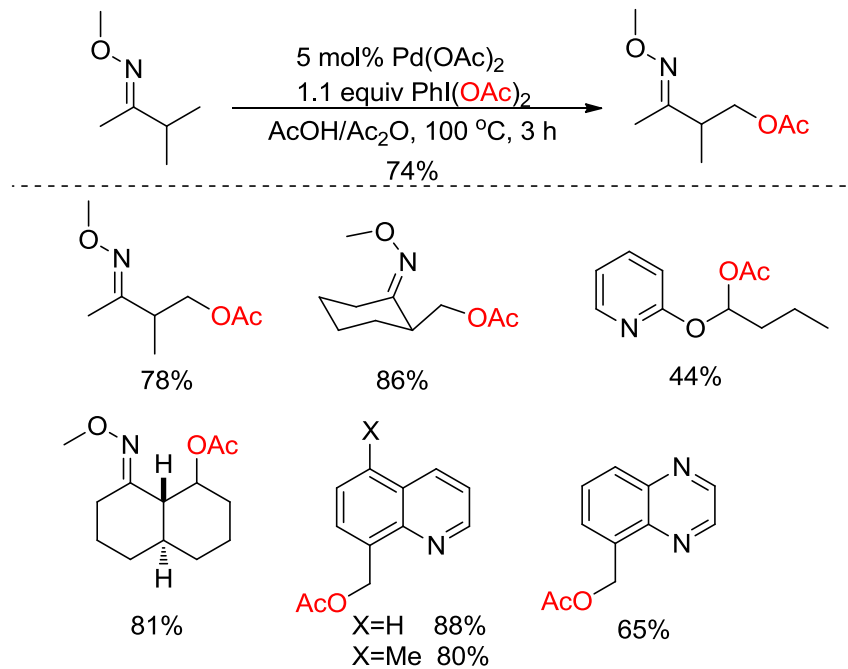
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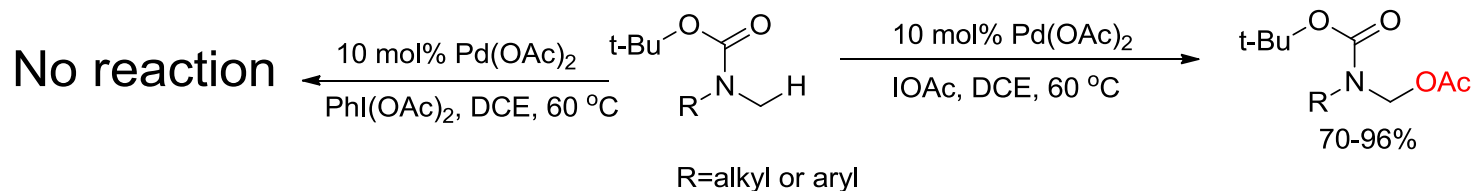


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R=Et 80%
R=i-Pr 72%
R=CF₃CH₂ 71%

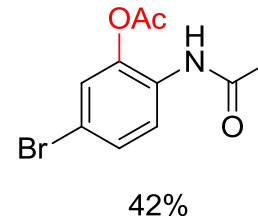
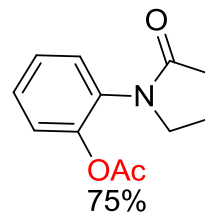
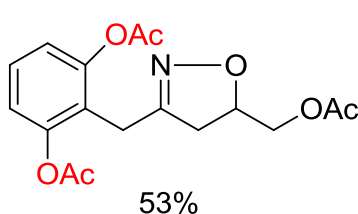
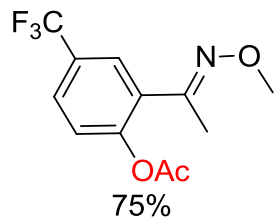
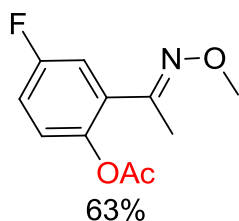
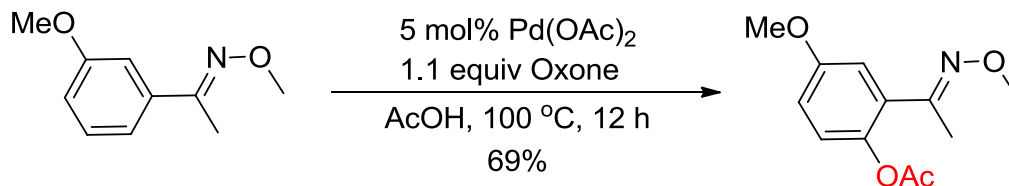


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Acetoxylation with other oxidants

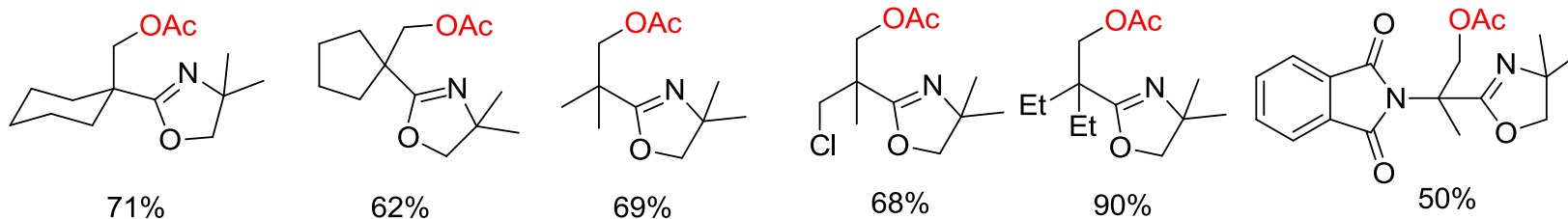
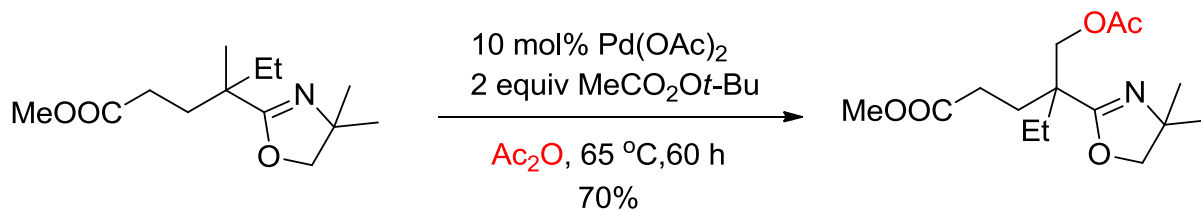


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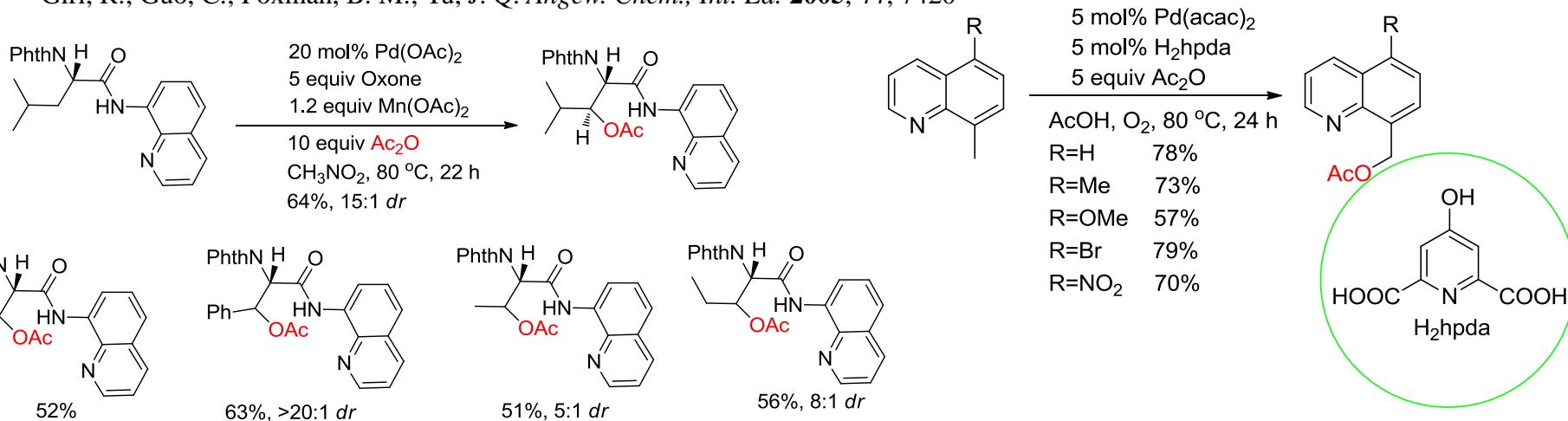


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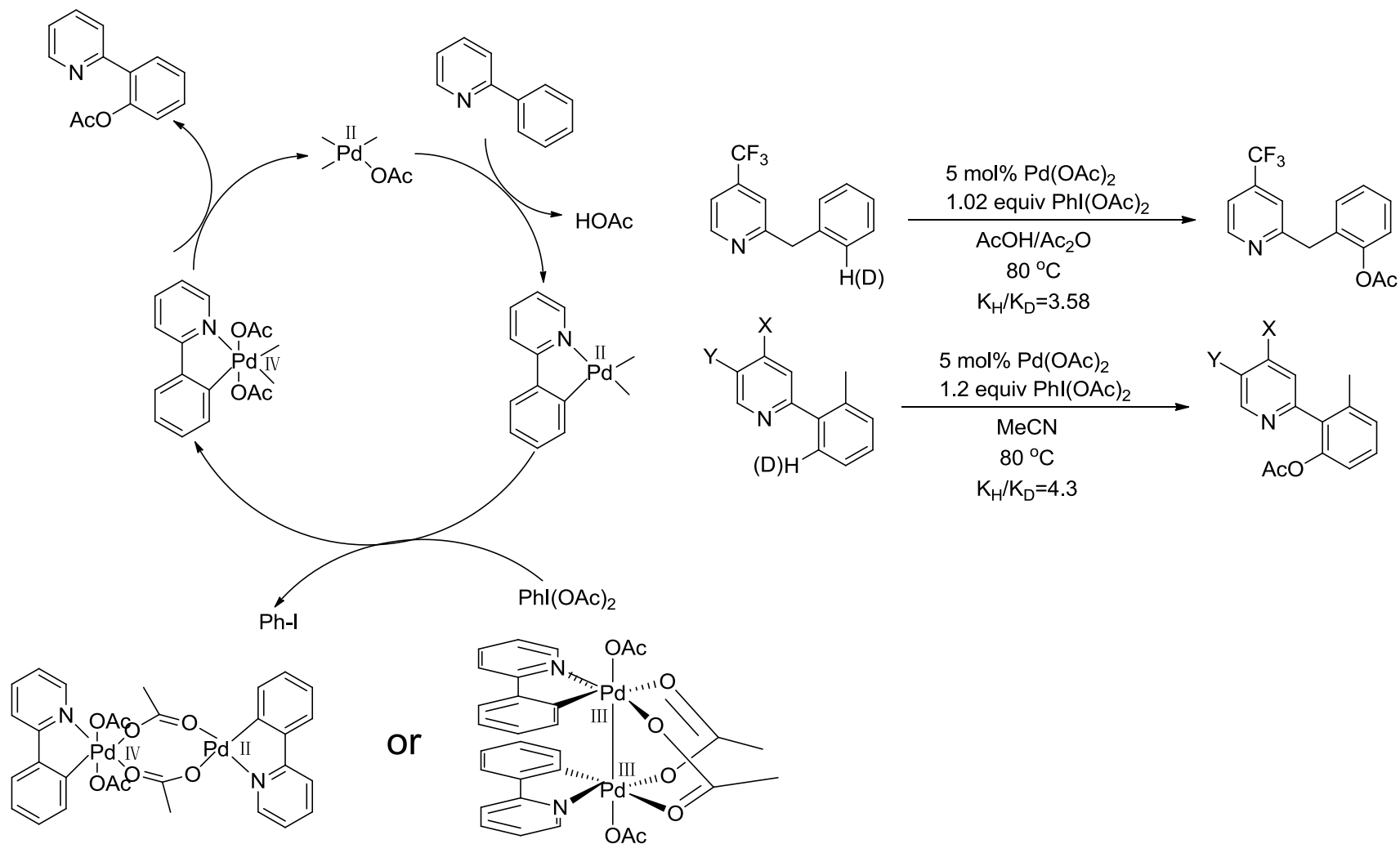
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Reddy, B. V. S.; Reddy, L. R.; Corey, E. J. *Org. Lett.* **2006**, *8*, 3391

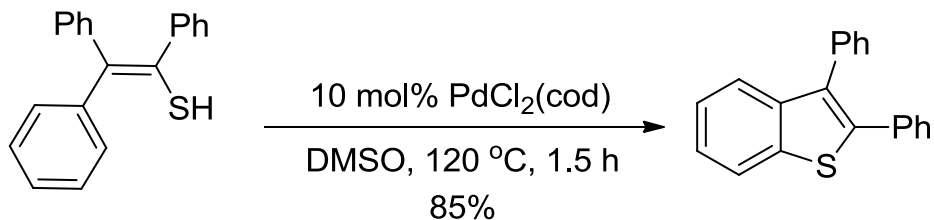
Zhang, J.; Khaskin, E.; Anderson, N. P.; Zavalij, P. Y.; Vedernikov, A. N. *Chem. Commun.* **2008**, 3625

Mechanism of Acetoxylation

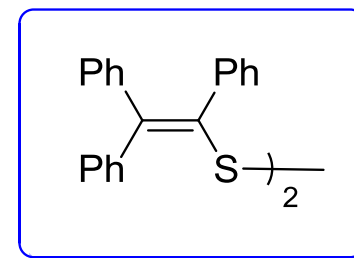
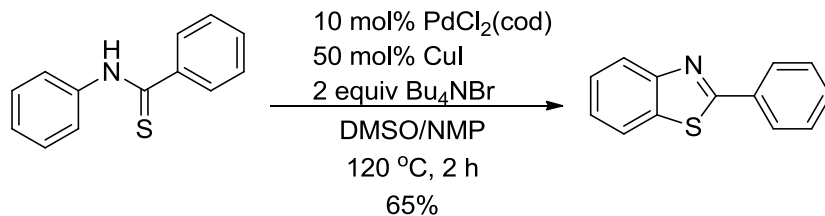


Dick, A. R.; Kampf, J. W.; Sanford, M. S. *Organometallics* **2005**, *24*, 482
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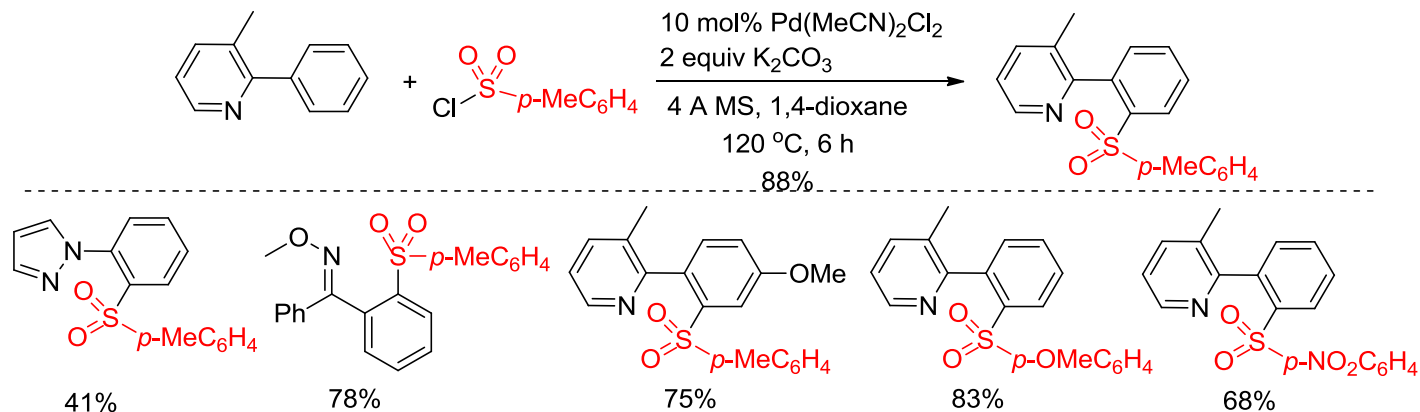
C-S Bond Formation



Inamoto, K.; Arai, Y.; Hiroya, K.; Doi, T. *Chem. Commun.* **2008**, 5529

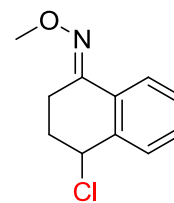
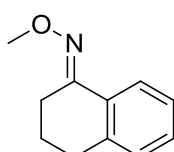
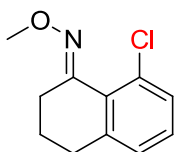
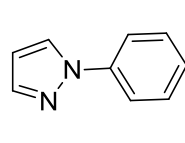
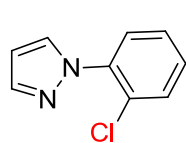
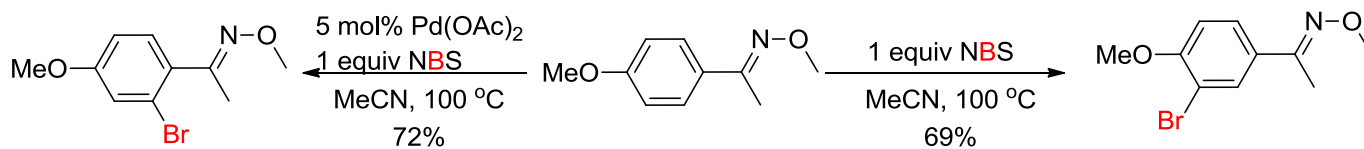
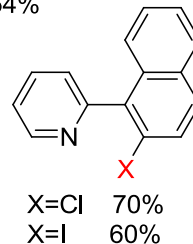
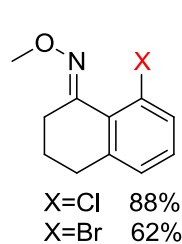
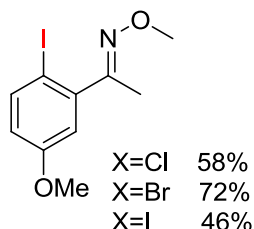
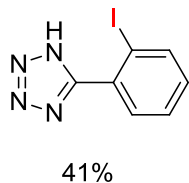
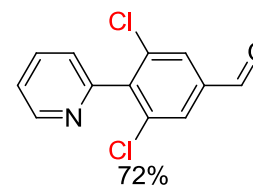
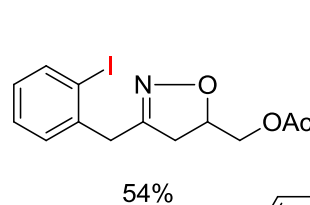
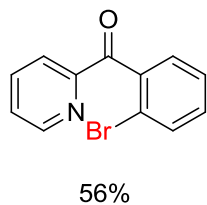
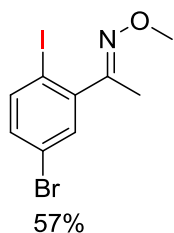
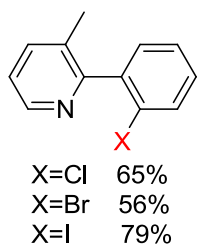
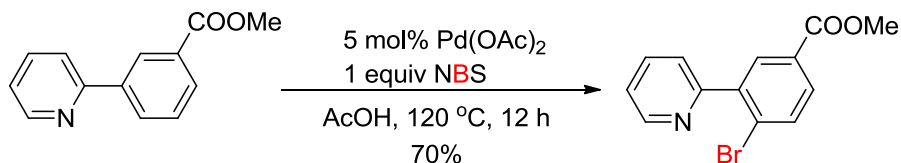


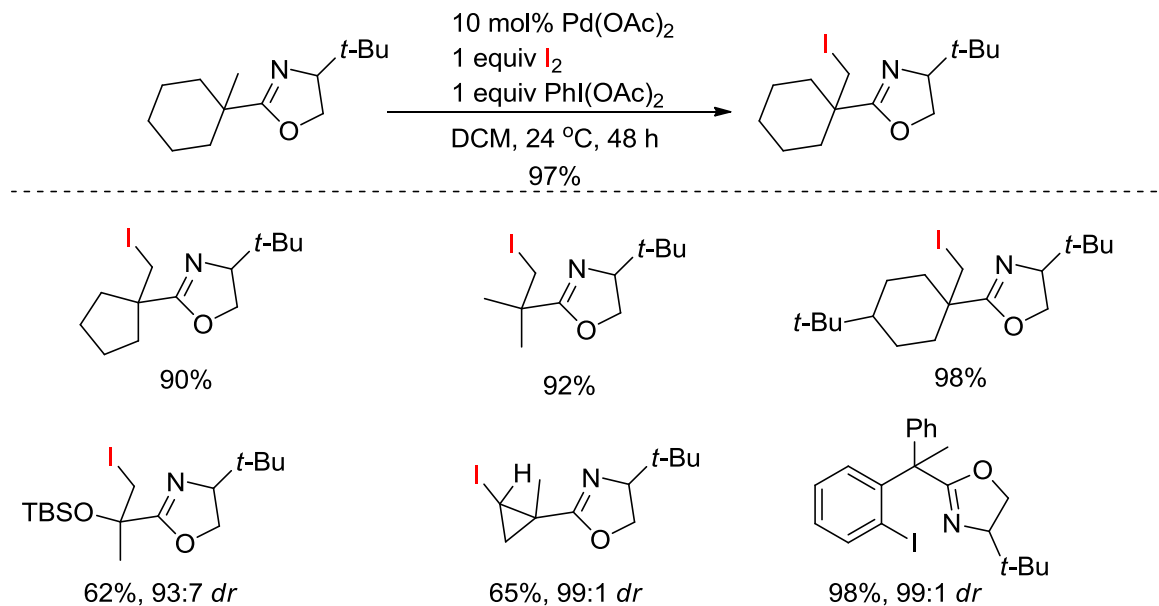
Inamoto, K.; Hasegawa, C.; Hiroya, K.; Doi, T. *Org. Lett.* **2008**, 10, 5147



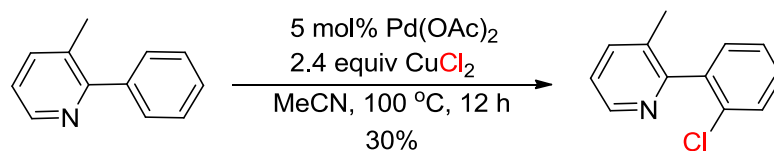
Zhao, X.; Dimitrijevic, E.; Dong, V. M. *J. Am. Chem. Soc.* **2009**, 131, 3466

C-X Bond Formation

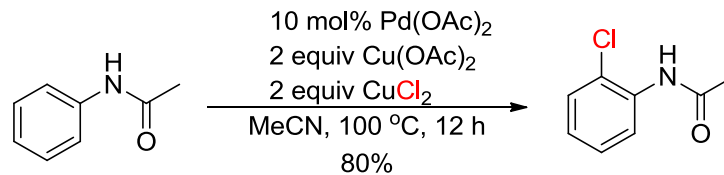




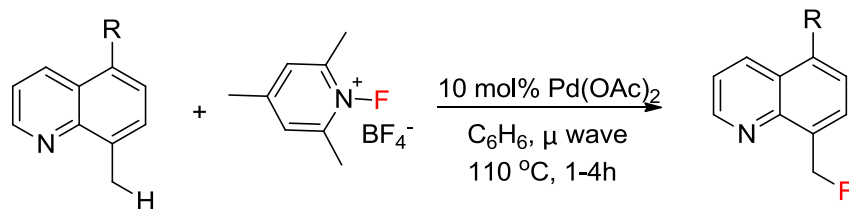
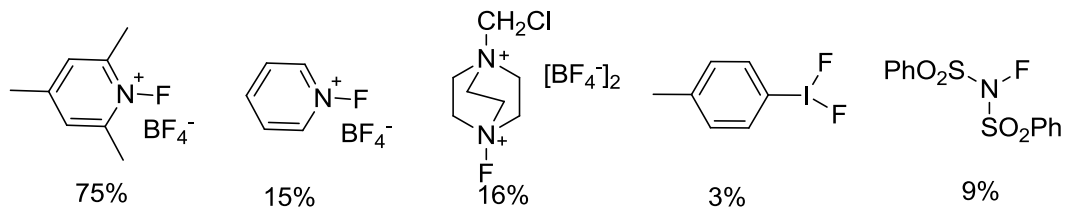
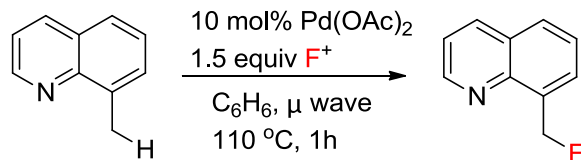
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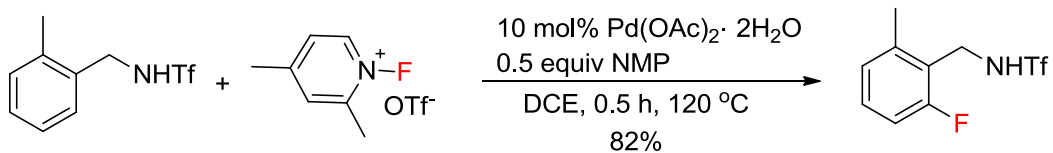
Kalyani, D.; Dick, A. R.; Anani, W. Q.; Sanford, M. S. *Org. Lett.* **2006**, *8*, 2523



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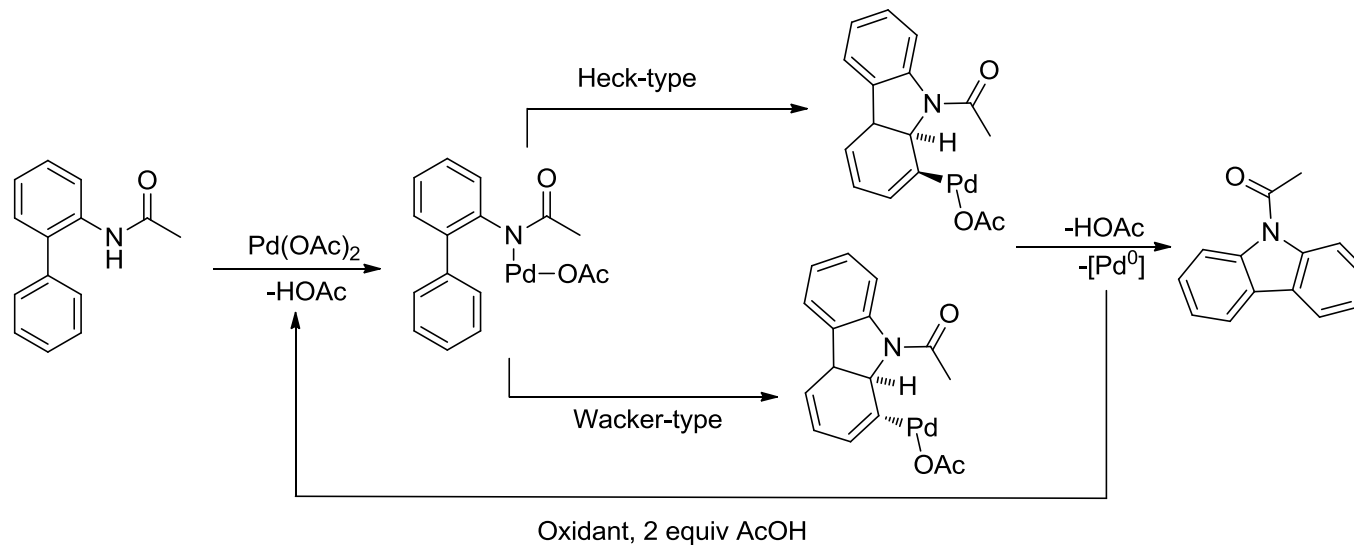
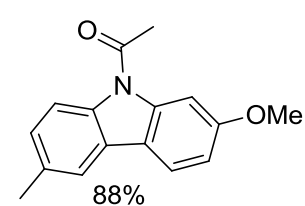
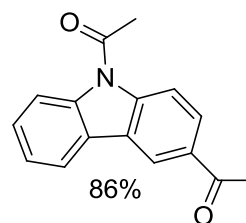
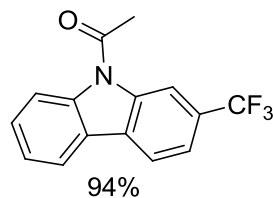
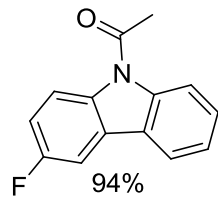
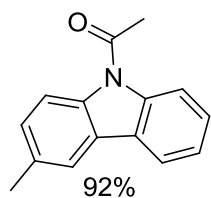
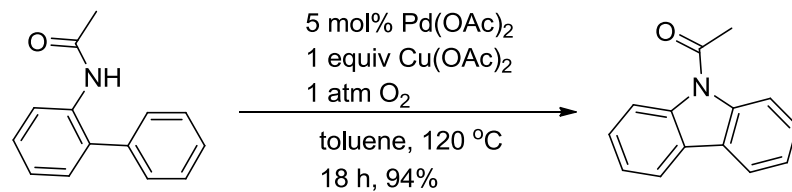
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 R=Me 57%
 R=F 49%
 R=Br 53%

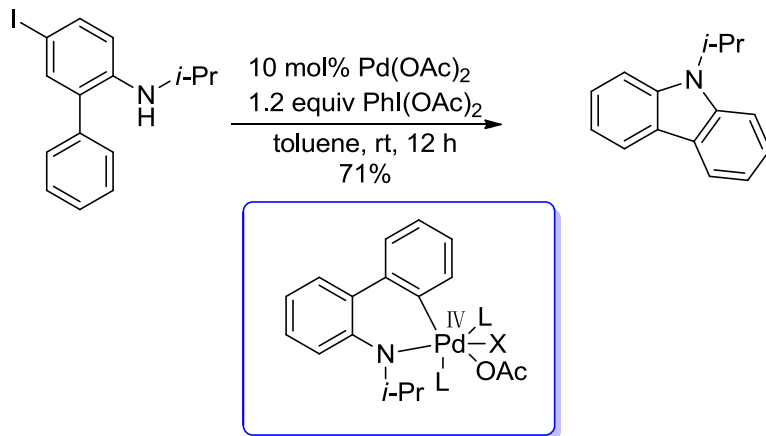


Hull, K. L.; Anani, W. Q.; Sanford, M. S. *J. Am. Chem. Soc.* **2006**, *128*, 7134

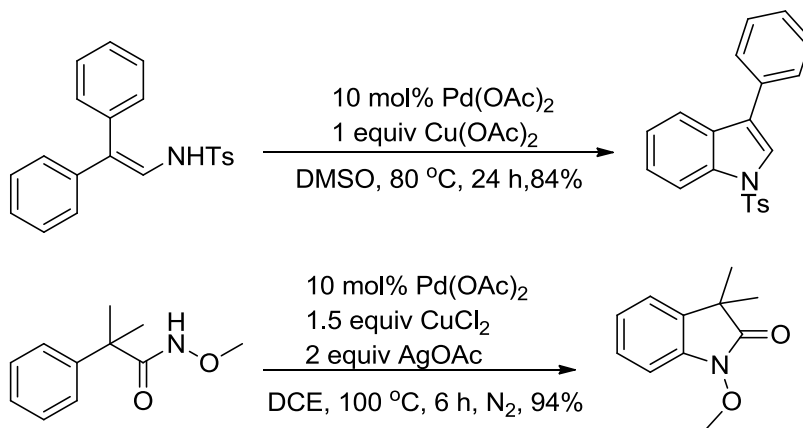
Wang, X.; Mei, T. S.; Yu, J. Q. *J. Am. Chem. Soc.* **2009**, *131*, 7520

C-N Bond Formation

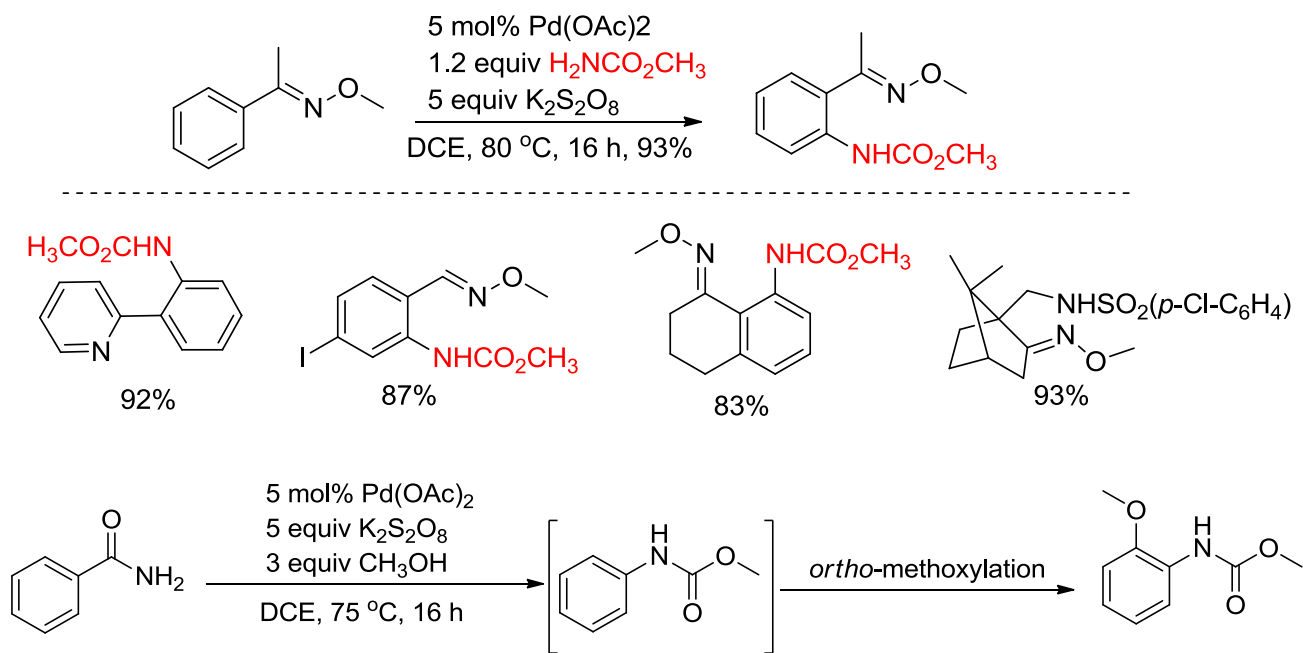




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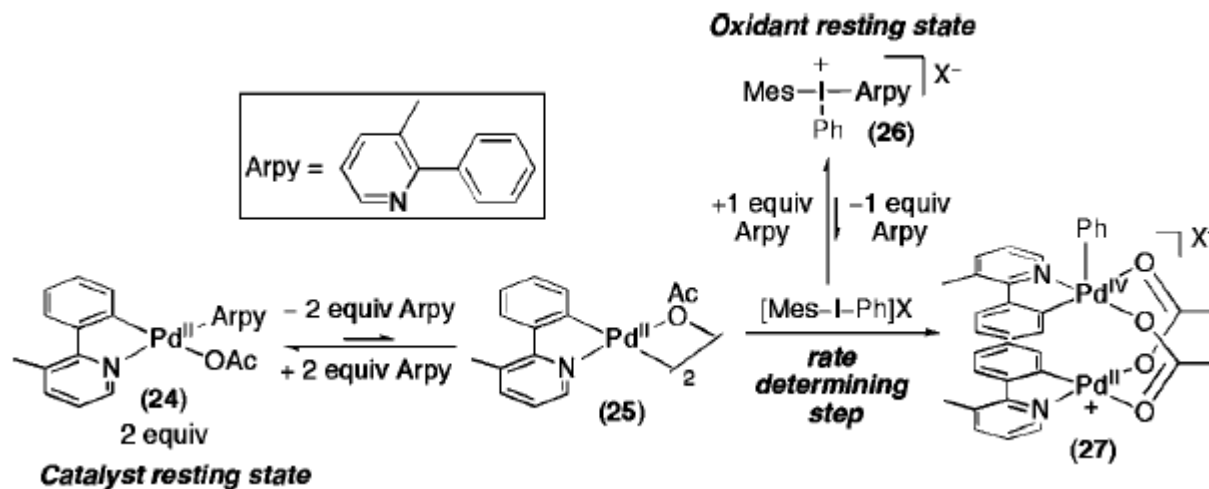
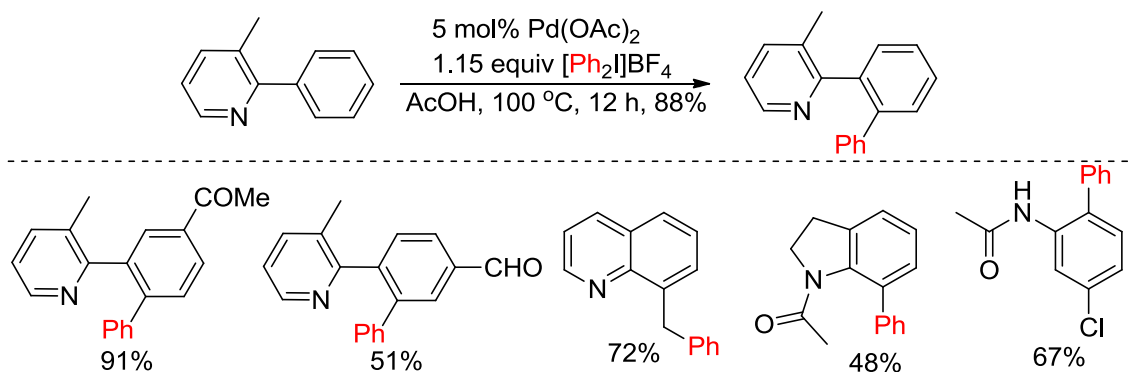


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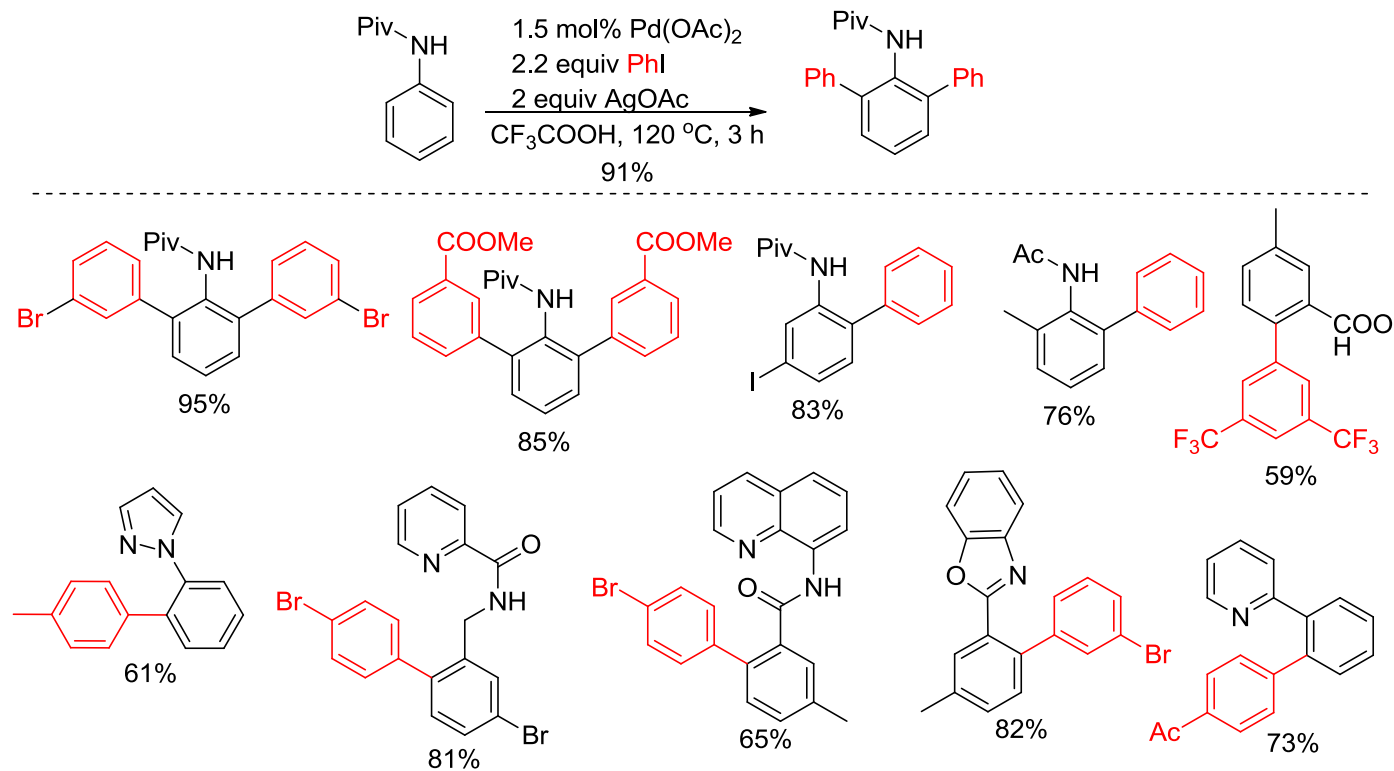
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C-C Bond Formation



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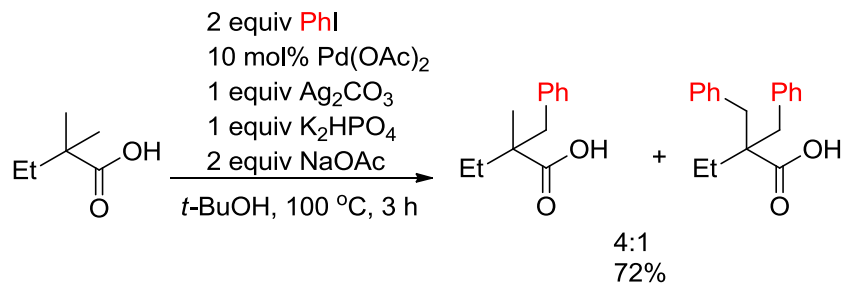
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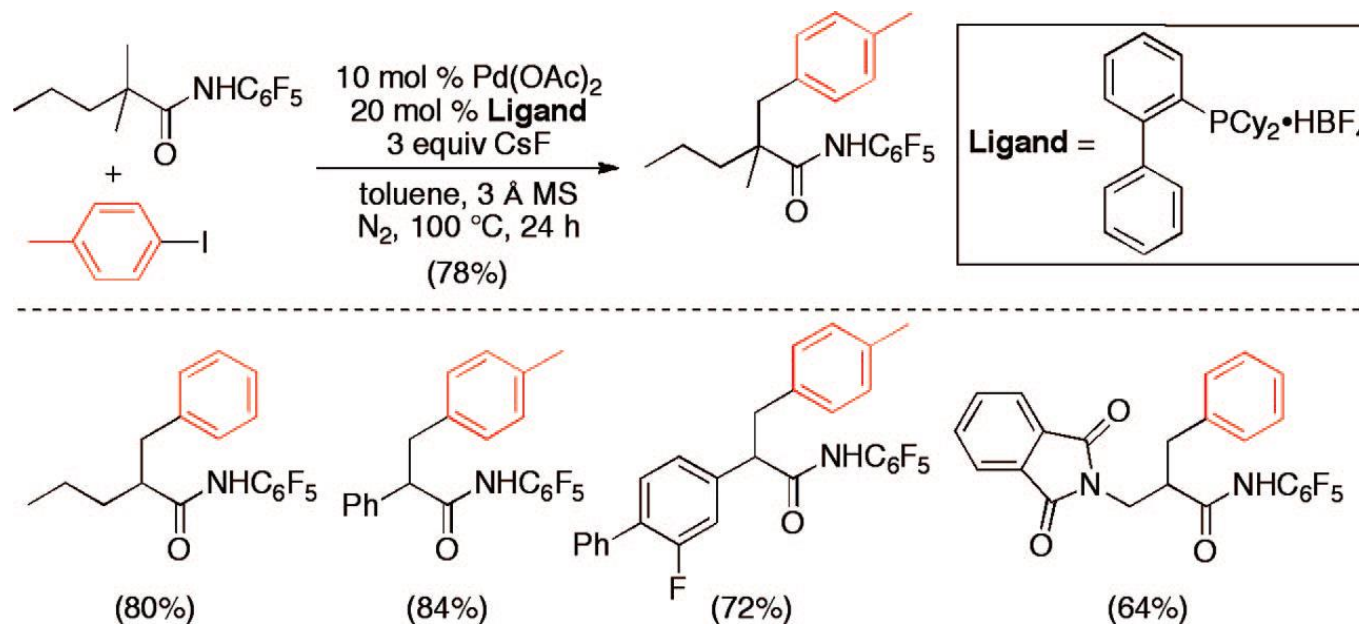
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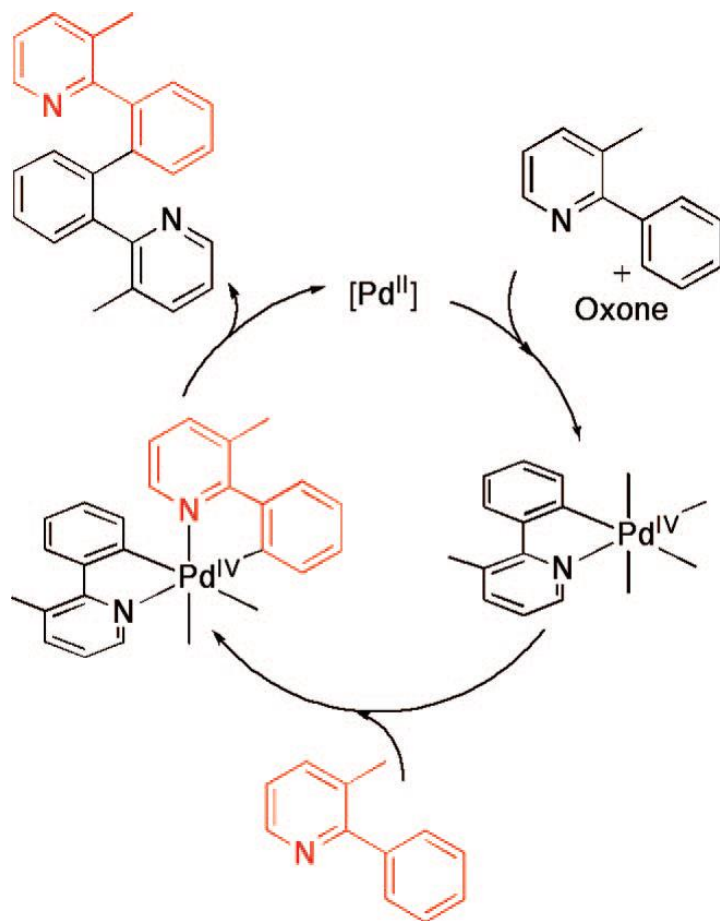
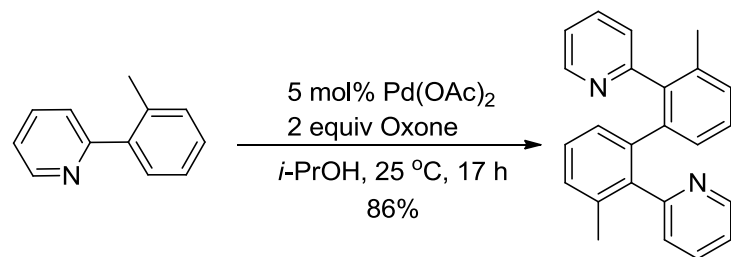
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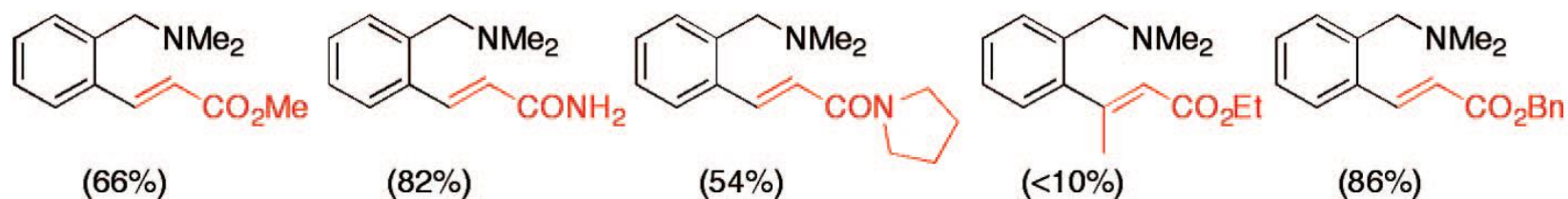
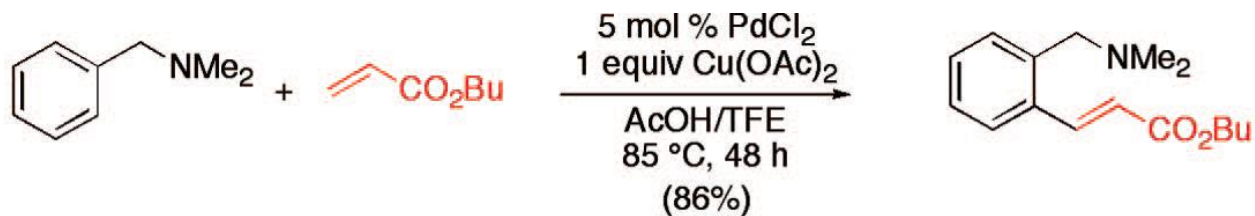


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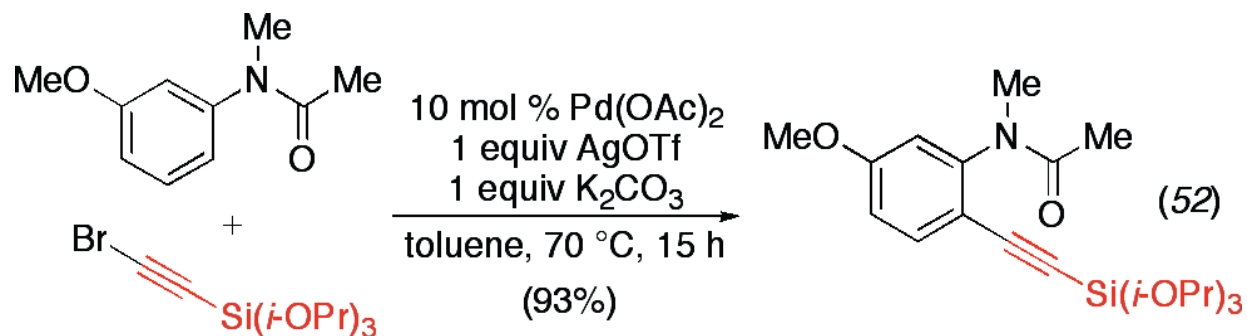


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Tobisu, M.; Ano, Y.; Chatani, N. *Org. Lett.* **2009**, *11*, 3250

Outlook

- New Directing Groups
- New Oxidants
- Asymmetric C-H Bond Activations

Acknowledgement:

- **Prof. Yong Huang**
- **All members of E201**

Thank you for your attention !