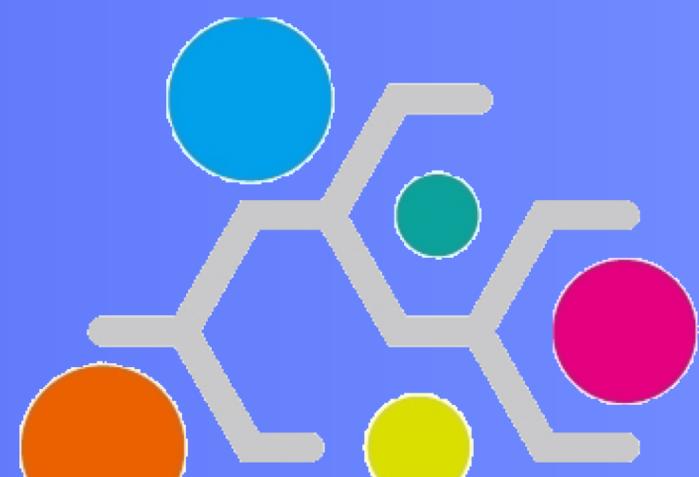


# Tf<sub>2</sub>O-mediated Reaction of Alkenyl Sulfoxides with Unprotected Anilines in Flow Microreactors



Alexandre Baralle,<sup>1</sup> Tomoaki Inukai,<sup>1</sup> Tomoyuki Yanagi,<sup>1</sup> Keisuke Nogi,<sup>1</sup> Aiichiro Nagaki,<sup>\*2</sup> Jun-ichi Yoshida,<sup>3</sup> Hideki Yorimitsu<sup>\*1</sup>

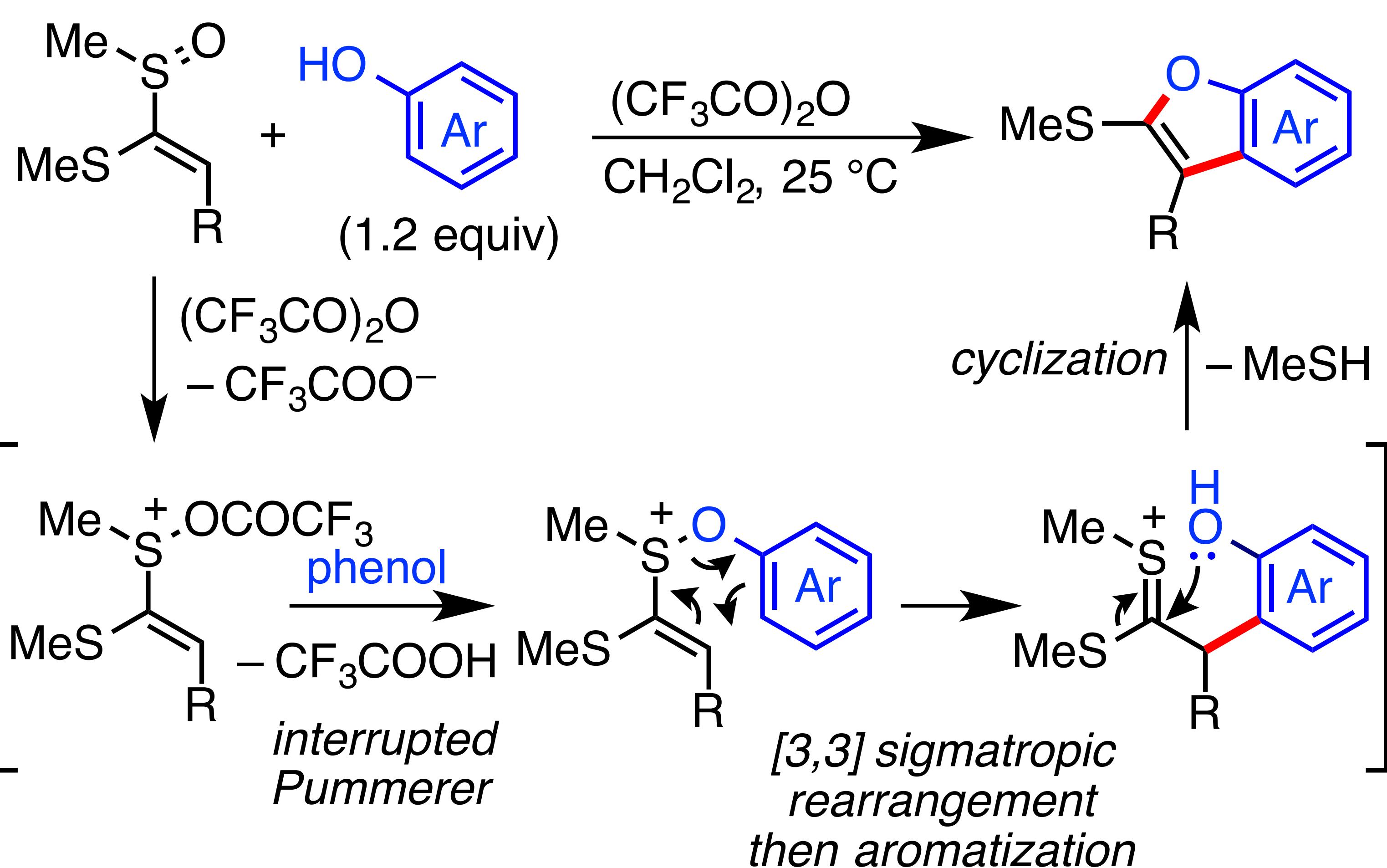
<sup>1</sup>Department of Chemistry, Graduate School of Science, Kyoto University

<sup>2</sup>Department of Synthetic Chemistry and Biological Chemistry, Graduate School of Engineering, Kyoto University

<sup>3</sup>National Institute of Technology, Suzuka College

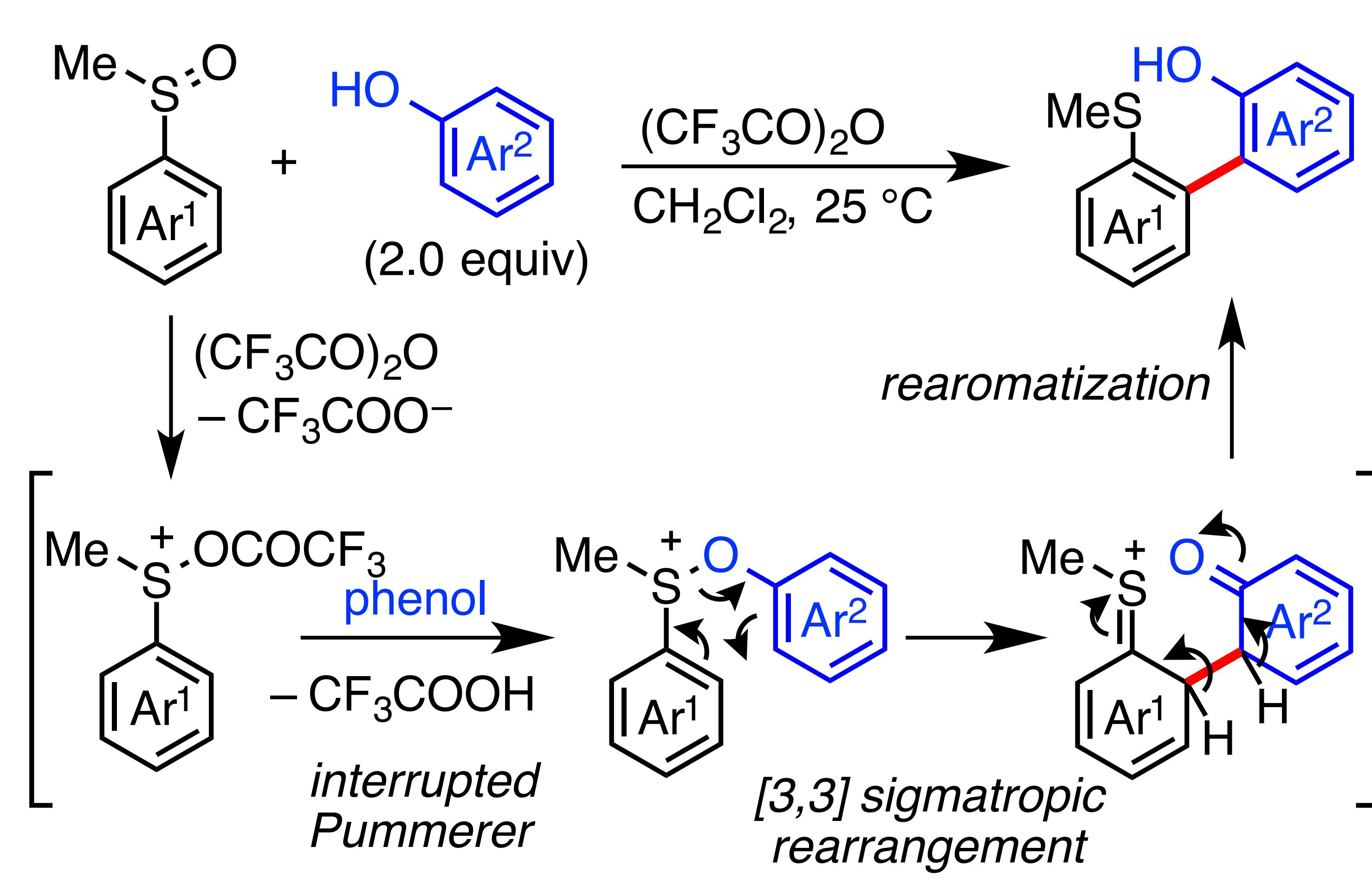
## Previous Work: Extended Pummerer Reactions with Phenols

### Benzofuran synthesis



JACS, 2010, 132, 11838; CEJ, 2012, 18, 12690; ACIE, 2014, 53, 7510.

### Dehydrogenative cross-coupling



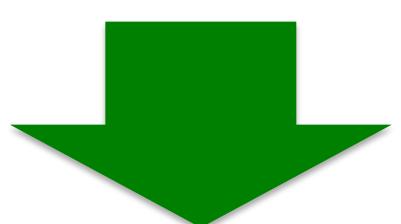
JACS, 2016, 138, 14582.

## Our Plan

### Extended Pummerer Reaction with Anilines

#### Problem

Anilines react with acid anhydrides directly in a **batch reactor**

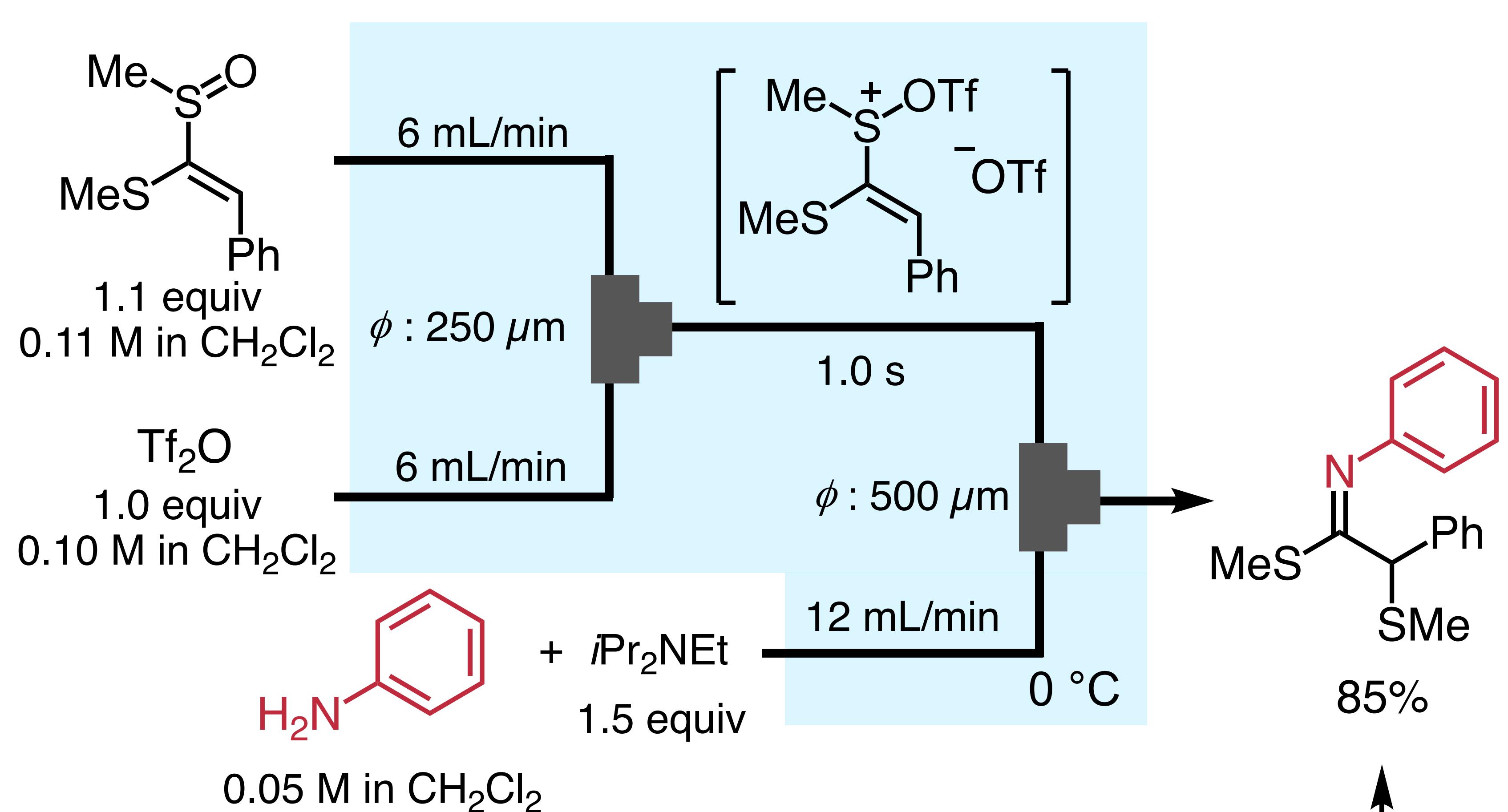


#### Solution

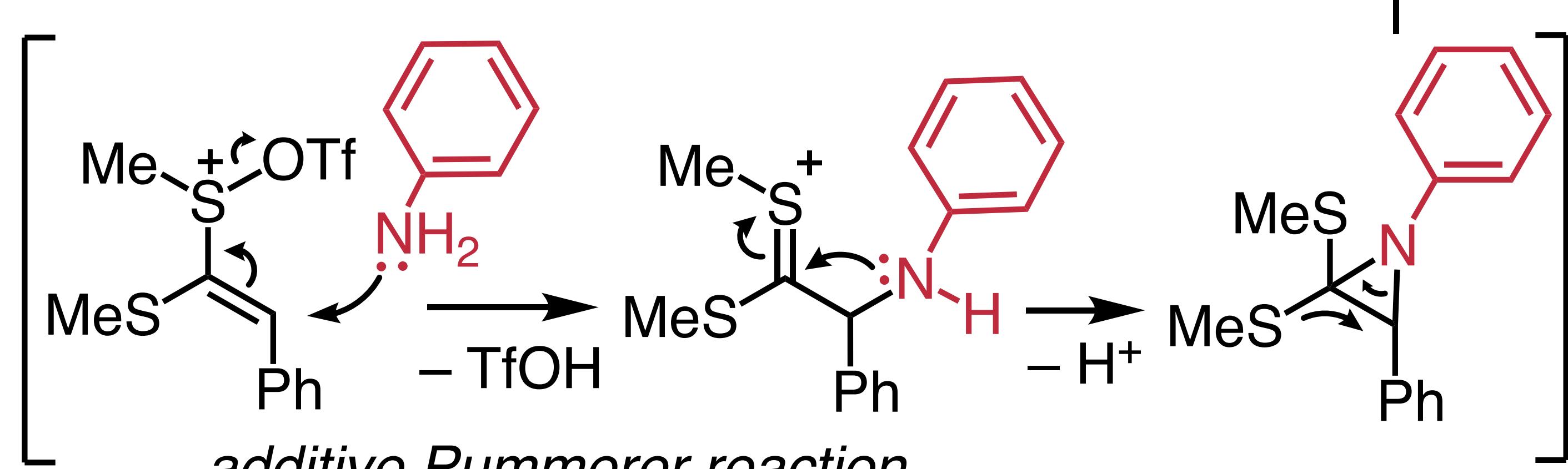
Formation of sulfonium intermediates in a **flow microreactor**

## This Work: Flow Microreactor System for Pummerer Chemistry with Anilines

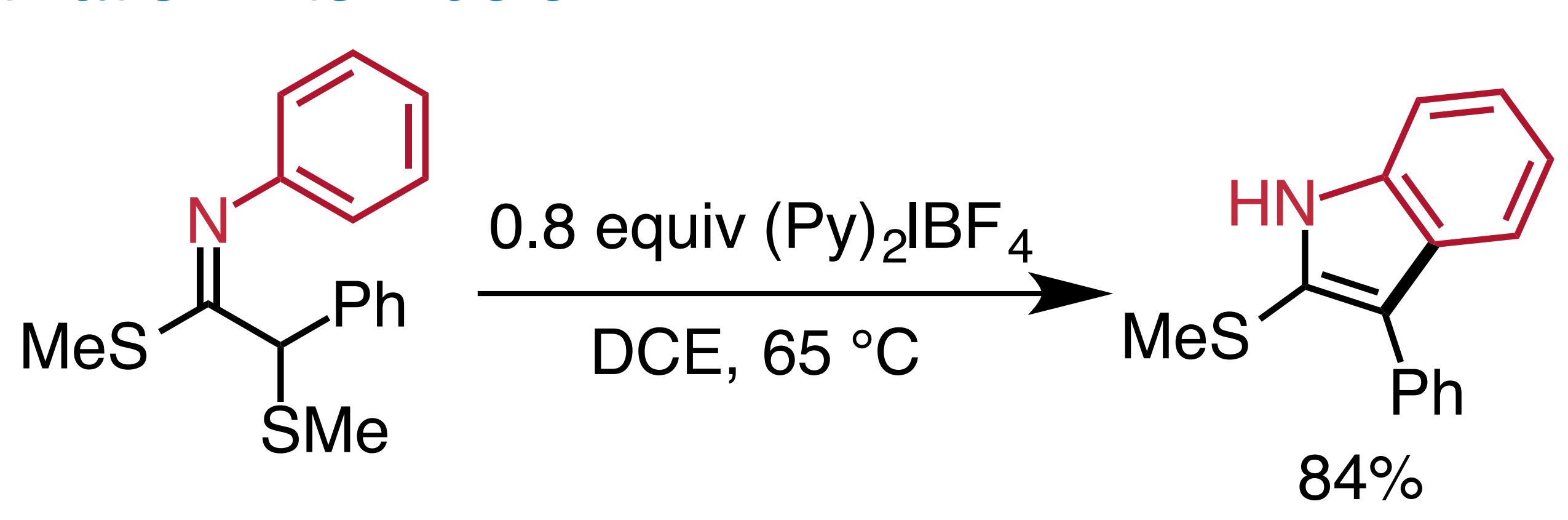
Chem. Lett. 2020, 49, 160-163.



### A possible reaction mechanism



### Derivatization into indole



### Scope of substrates

		77%
MeS-Substituted Aniline	$R^2 = H$	68%
	$= OMe$	63%
	$= CF_3$	63%
$R^1 = 4\text{-OMe}$	78%	
$= 4\text{-CF}_3$	69%	
$= 4\text{-COOEt}$	82%	
$= 4\text{-CN}$	79%	
$= 2\text{-Me}$	75%	
$= 3\text{-OMe}$	77%	
	53%	
	63%	

### Temperature-residence time map

(for Pummerer activation step)

