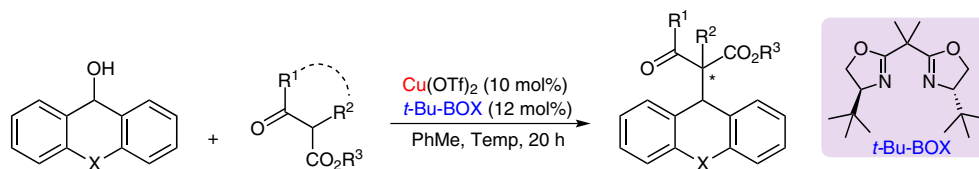
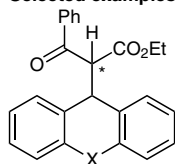


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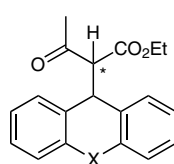
Copper-Catalyzed Asymmetric Alkylation of  $\beta$ -Keto Esters with Xanthydrols*Adv. Synth. Catal.* **2013**, 355, 2815–2821.Enantioselective Alkylation of  $\beta$ -Keto Esters

X = O, S

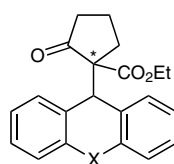
Selected examples:



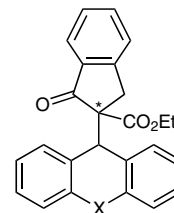
X = O,  $-50\text{ }^\circ\text{C}$   
 82% yield, 80% ee  
 X = S,  $0\text{ }^\circ\text{C}$   
 45% yield, >99% ee



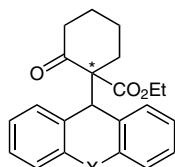
X = O,  $-50\text{ }^\circ\text{C}$   
 91% yield, 0% ee  
 X = S,  $-20\text{ }^\circ\text{C}$   
 75% yield, 69% ee



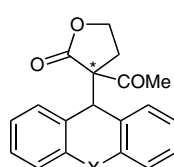
X = O,  $-50\text{ }^\circ\text{C}$   
 87% yield, 73% ee  
 X = S,  $-20\text{ }^\circ\text{C}$   
 37% yield, 90% ee



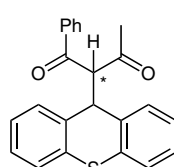
X = O,  $-50\text{ }^\circ\text{C}$   
 89% yield, 82% ee  
 X = S,  $0\text{ }^\circ\text{C}$   
 71% yield, 63% ee



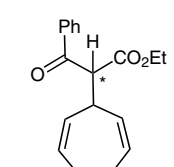
X = O,  $-50\text{ }^\circ\text{C}$   
 90% yield, 16% ee  
 X = S,  $0\text{ }^\circ\text{C}$   
 42% yield, 34% ee



X = O,  $-78\text{ }^\circ\text{C}$   
 87% yield, 56% ee  
 X = S,  $0\text{ }^\circ\text{C}$   
 38% yield, 71% ee

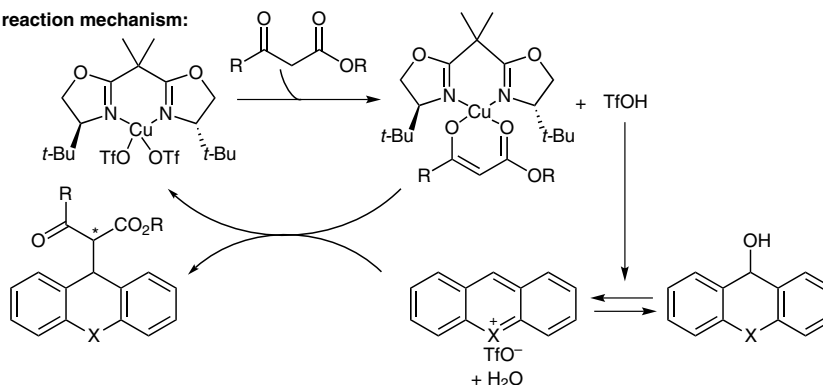


$-50\text{ }^\circ\text{C}$   
 37% yield, 85% ee



$0\text{ }^\circ\text{C}$   
 92% yield, 30% ee

Proposed reaction mechanism:



**Significance:** The  $\text{Cu(OTf)}_2$ /*tert*-butyl-bis-oxazoline catalyst system allows the asymmetric alkylation of  $\beta$ -keto esters with free benzylic alcohols, for example, xanthydrols as alkylating agents. The reaction is environmentally benign as it generates only water as by-product.

**Comment:** The reaction between the asymmetric  $\text{Cu(II)}$ - $\beta$ -keto ester derivative and the in situ generated carbocation proceeds via an  $\text{S}_{\text{N}}1$  mechanism. Albeit unknown, the water produced has a specific role to facilitate the reaction.

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