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Palladium-Catalyzed Tandem Reaction of α-Aminophenols, Bromoalkynes and Isocyanides to Give 4-Amine-benzo[b][1,4]oxazepines


Palladium-Catalyzed Synthesis of Benzo[b][1,4]oxazepines

**Significance:** Reported is the palladium-catalyzed synthesis of benzo[b][1,4]oxazepines 4 via the annulation of α-aminophenols 1 with bromoalkynes 2 and isocyanides 3. Substrate-scope investigation revealed broad tolerance to variation of all components 1–3, particularly across sterically and electronically differentiated aryl bromoalkynes 2, with a reduction in yield noted for alkyl bromoalkynes (4o). Experiments demonstrating the competency of 5 under the standard reaction conditions are offered in support of the proposed mechanism.

**Comment:** Building on their previous investigations into combining the nucleophilic addition of isocyanides 3 to bromoalkynes 2 with palladium catalysis (Chem. Commun. 2012, 48, 3545), the current report extends this methodology allowing the synthesis of benzoxazepines traditionally synthesized by multiple-step procedures. Taking advantage of the established addition of phenols to bromoalkynes (For furan synthesis, see: S. Wang et al. Org. Lett. 2011, 13, 5968) the current report, intercepting intermediate 5, appears to have exceptional scope.