Enantioselective Modular Approach to 3-Azabicyclohexanes

**Significance:** 3-Azabicyclo[3.1.0]hexanes are present in a wide range of bioactive compounds. Besides many common methods to access this scaffold, the 1,3-dipolar cycloaddition of azomethine ylides to cyclopropanes (A. S. Filatov et al. *J. Org. Chem.* 2017, 82, 959) and multicomponent reactions in water (M. Ghorbani et al. *Org. Lett.* 2016, 18, 4759) have recently been described. The present work takes advantage of the high electrophilic character of the intermediate alkylfluoro-substituted ketamine 2 to produce highly substituted 3-azabicyclo[3.1.0]hexanes 3 by addition of nucleophiles. The presence of the strained cyclopropane ring ensures the diastereoselective control of the addition.

**Comment:** Reported is the enantioselective palladium-catalyzed cyclization of imidoyl chlorides 1 to produce cyclopropane-fused dihydropyrrole 2. The scope of this transformation is broad, and 1 with various substituents gave products 2 in high yields and high enantioselectivities. When R₂ = H, the reaction proceeded with low yield, although the er was unaffected. Cyclopropane C–H functionalization was observed exclusively in the presence of an aryl substituent (R₁ = Ar), to give dihydropyrroles 2. However, switching the ligand to Ph₃P reversed the chemoselectivity to aryl C–H functionalization, producing spirocyclic dihydroisoquinolines 4. The reaction of electrophilic ketimines 2 with various nucleophiles gave pyrrolidines 3 diastereoselectively. Moreover, 3 can be accessed directly from 1 in a one-pot manner without any significant loss in enantioselectivity.