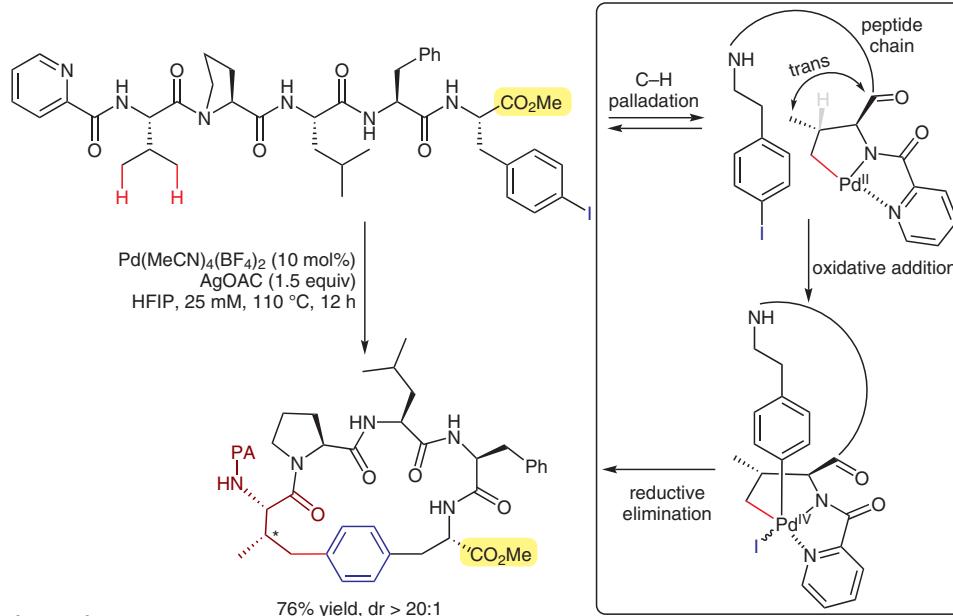
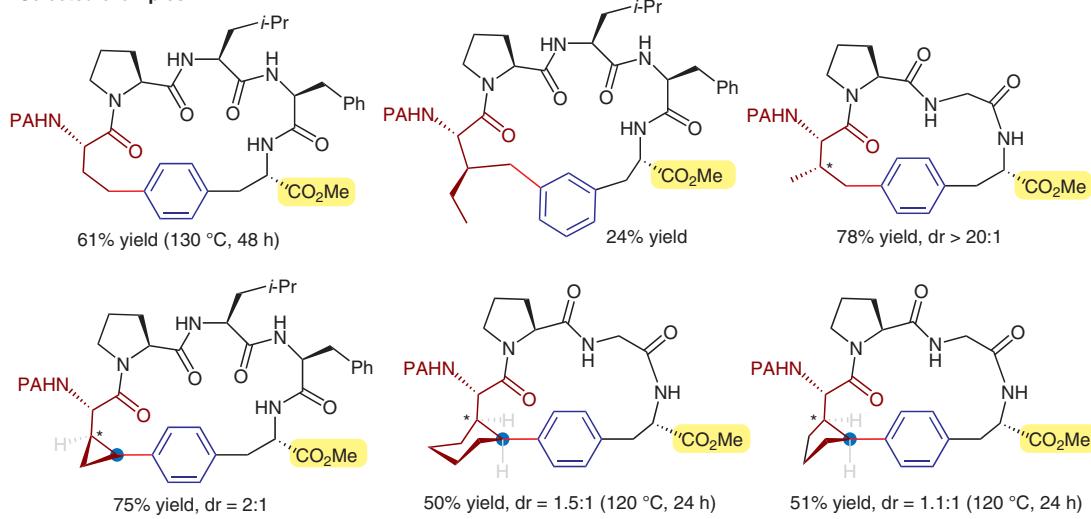


# Palladium-Catalyzed Synthesis of Cyclophane-Braced Peptide Macrocycles

Pd-catalyzed picolinamide-directed intramolecular C–H arylation of peptides:



Selected examples:



**Significance:** Cyclic peptides possess great potential for modulating challenging biological processes. The authors have developed a general method for the synthesis of cyclophane-braced peptide macrocycles through palladium-catalyzed  $\text{C}(\text{sp}^3)\text{–H}$  arylation.

**Comment:** Cyclophane motifs with unusual structural and stereochemical complexity were obtained by the picolinamide-directed intramolecular  $\gamma$ -selective  $\text{C}(\text{sp}^3)\text{–H}$  arylation of the N-termini of peptide substrates at both the methyl and methylene positions.