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Stereospecific Nickel-Catalyzed Cross-Coupling Reactions of Alkyl Grignard Reagents and Identification of Selective Anti-Breast-Cancer Agents

Angew. Chem. Int. Ed. 2014, 53, 2422-2427.

Stereospecific Kumada Cross-Coupling of Alkyl and Aryl Grignards

$$\begin{split} &\text{Ar} = \text{Naph, 6-MeONaph, benzothiophenyl, benzofuranyl} \\ &\text{R}^1 = \text{Et, } \dot{\text{-}}\text{Bu, CH}_2\text{Cy, Bn, CH}_2\text{Bn, } (\text{CH}_2)_2\text{CHCMe}_2 \\ &\text{R}^2 = \text{various Alk, Ar} \end{split}$$

Selected examples:

Significance: Herein, the authors report an efficient nickel-catalyzed cross-coupling reaction of long-chain alkyl and various aryl Grignards with benzylic ethers providing the corresponding products with high enantiospecificity. Furthermore, applying this reaction protocol, compounds that show promising inhibition of breast cancer cell proliferation could be prepared.

Comment: Decreasing the reaction concentration raised the efficiency of the transformation and less elimination byproduct was generated. Also, the use of more than two equivalents of Grignard reagent was disadvantageous for the reaction.

SYNFACTS Contributors: Paul Knochel, Diana Haas Synfacts 2014, 10(5), 0515 Published online: 17.04.2014 **DOI:** 10.1055/s-0033-1341218; **Reg-No.:** P04114SF

Category

Metal-Mediated Synthesis

Key words

Grignard reagents

Kumada coupling

nickel

