**MgBr₂·OEt₂ – A Versatile Reagent in Organic Synthesis**

Compiled by Suresh Kumar Tipparaju

Suresh Kumar Tipparaju obtained his BSc from Osmania University and MSc in chemistry from Sri Sathya Sai Institute of Higher Learning, Prashanthinilayam in 1998. He has been working on ‘stereoselective functionalizations on arene tricarbonylchromium template’ in Prof. Amitabha Sarkar’s group at National Chemical Laboratory. His research interests are in metal-mediated organic synthesis, asymmetric synthesis, and the chemistry of hypervalent silicon.

Division of Organic Chemistry (Synthesis), National Chemical Laboratory, Pune – 411 008, India
Fax +91(20)5893153; E-mail: tsuresh_kumar@yahoo.com

**Abstracts**

(A) Condensation of 2,5-dimethoxy-2,5-dihydrofuran with ethyl vinyl ether in the presence of a catalytic amount of MgBr₂·OEt₂ resulted in 2-furylacetaldehyde diethyl acetal in 50% yield. The reaction is a formal acetal and ethyl vinyl ether condensation followed by aromatization. The protocol has been employed in the synthesis of various 2-(2-furo)tetrahydrofuranic or -pyranic moieties in good yields.¹¹

(B) A mild and practical N-acylation of amides was possible by the dual activation of both amides and acid anhydrides with MgBr₂·OEt₂.¹² The method was applicable to amides that can undergo O-acylation and are susceptible to racemization or O,N-acyl migrations.

(C) We have recently reported a mild method for the formation of cyclic siloxanes by the exchange of the Li counterion of an intermediate alkoxide with Mg, using excess MgBr₂·OEt₂.¹³ The reaction may formally be considered to be a semi-Brook rearrangement.

(D) MgBr₂·OEt₂, in combination with Bu₃SnH, was effective in a chelation-controlled reductive opening of methoxybenzylidene acetals.¹⁴ The reaction offers a mild and efficient method for selective mono-MPM ether protection of diols. High conversions, regioselectivity and tolerance to functional groups make this a very useful protocol in natural product synthesis.

(E) MgBr₂·OEt₂ effected deprotection of aliphatic SEM ethers under extremely mild and high yielding conditions in the presence of other sensitive groups like acetones, TBS and TIPS ethers and O-silylated cyanohydrins.¹⁵ A variety of functionalities including alcohols, esters, benzyl groups, dithianes, and methoxy acetals are tolerated.

**Introduction**

Use of magnesium(II) species as Lewis acid catalysts for various functional group transformations is well documented.¹ Of these, magnesium halides are the most useful. Ready availability and ease of preparation prompted the frequent use of MgBr₂·OEt₂ in various organic transformations. The oxophilic and coordinating nature of MgBr₂·OEt₂ has been demonstrated through its use as a bidentate chelating Lewis acid in a number of chelation-controlled reactions such as cycloadditions,² asymmetric aldol reactions,³ rearrangements,² radical additions,⁵,⁶ hydrogen transfer reactions,⁷ stereoselective reductions,⁸ and anomerizations.⁹ MgBr₂·OEt₂ is commercially available as a grey solid (mp >300 °C, fp 35 °C). It can be readily prepared by reacting a slight excess of magnesium turnings with 1,2-dibromoethane in anhydrous diethyl ether.¹⁰ The solution can be stored at room temperature for several months and the solid can be stored in a vacuum desiccator for indefinite periods without any loss of activity.
(F) The MgBr₂·OEt₂–Me₅S system was used for a mild and chemoselective deprotection of p-methoxybenzyl (PMB) ethers in the presence of 1,3-diene, β-butyldimethylsilyl (TBDMS) ether, benzoate, benzyl ether, and acetonides. The method is especially effective for 1,3-diene systems that tend to isomerize rapidly when other protocols are employed.¹⁷

(G) A stereodivergent opening of the oxirane ring with MgBr₂·OEt₂ was recently described. While MgBr₂·OEt₂ alone resulted in the bromohydrin in high diastereomeric excess, use of MgBr₂·OEt₂/Amberlyst 15 gave the syn-product in high yields.

(H) The efficacy of MgBr₂·OEt₂ as a chelating Lewis acid in highly diastereoselective addition of nucleophiles to Cr(CO)₃-complexed benzyl ether, and acetonides. The method is especially effective for the presence of 1,3-diene, β-butyldimethylsilyl (TBDMS) ether, benzoate, benzyl ether, and acetonides. The method is especially effective for 1,3-diene systems that tend to isomerize rapidly when other protocols are employed.¹⁷

(I) MgBr₂·OEt₂ could afford 3-hydroxyazetidines by a highly region- and stereoselective cyclization of 2,3-epoxy amines.²⁰

References