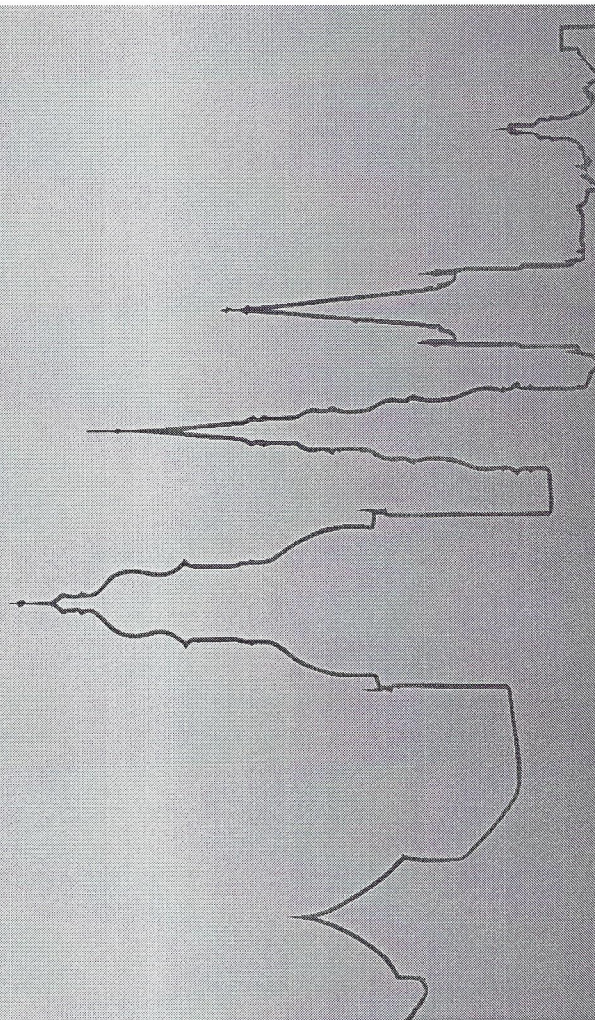


Drug Discovery Conference

August 27-29, 2015, Riga, Latvia



Abstract book

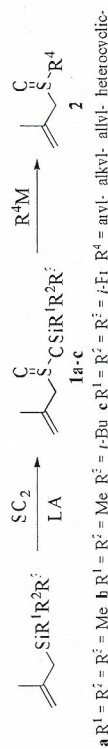
PP65. FROM METHALLYLSILANES TO METHALLYLSULFOXIDES: APPLICATION OF SILA-ENE REACTION OF SULFUR DIOXIDE

Agnese Stukute, Vilnis Peipinš, Māris Turks

Faculty of Material Science and Applied Chemistry, Riga Technical University,
Paula Valdena Str. 3/7, LV-1048, Riga, Latvia
E-mail: maris_turks@kcf.rtu.lv

Structural motif of sulfoxides is present in many naturally occurring and biologically active compounds, that are used as medication. Sulfoxides are also well-recognized synthetic intermediates. They form complexes with transition-metals and are used in catalysis. The most common approaches for synthesis of sulfoxides are oxidation of sulfides and C-S bond formation in nucleophilic substitution reactions.

Herein we report an application of silyl sulfonates **1** in the synthesis of variously substituted methallylsulfoxides **2** (see Scheme 1). In order to optimize the reaction conditions for sulfoxide **2** synthesis we investigated influence of solvent, temperature, organometallic reagent and Lewis acid additive on sulfoxide **2** yield. We have also diversified silyl moiety in sulfonate **1** structure, examining trimethylsilyl- (**1a**), *tert*-butyldimethylsilyl- (**1b**) and triisopropylsilyl sulfonate (**1c**) in order to increase the yields of sulfoxides **2**.



Scheme 1. Strategy of sulfoxide **2** synthesis

Experiments with Grignard reagents as nucleophiles and silyl sulfonates **2a,b** as sulfinyl transfer reagents showed the most promising results. The nucleophilic attack of Grignard reagents was accelerated in toluene and in the presence of LiCl or ZnCl₂ as Lewis acidic additives. The scope of the method has been demonstrated with the successful incorporation of aryl-, alkyl-, and heterocyclic Grignard reagents. Under the given experimental conditions trialkylsiloxy groups act as good leaving groups. Above described method gives opportunity to synthesize methallyl sulfoxides **2** in up to 83% yield.

Acknowledgements:

A.S. thanks the Latvian Academy of Sciences for scholarship. This work was financed by the Latvian Council of Science Grant No 12.0291.