

SYNTHESIS OF PYRROLE AND SUBSTITUTED PYRROLES (REVIEW)

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ABSTRACT

Pyrrole is widely known as a biologically active scaffold which possesses a diverse nature of activities. The combination of different pharmacophores in a pyrrole ring system has led to the formation of more active compounds. Pyrrole containing analogs are considered as a potential source of biologically active compounds that contains a significant set of advantageous properties and can be found in many natural products.

The present review highlights the synthetic methods of representatives of nitrogen heterocycles such as pyrrole, substituted pyrroles and other related compounds. The aim of this review is to indicate and summarise the different methods for the synthesis of nitrogen containing heterocycles from the group of pyrrole and pyrrole related structures.

Keywords: pyrrole, synthesis, pharmacological activity.

INTRODUCTION

Pyrrole is a five membered heterocyclic compound, corresponding to the C_4H_5NH general formula [1]. It is a colorless volatile liquid, unstable in the presence of air, where it easily darkens. Thus a preliminary distillation before use is necessary [2].

Pyrrole is included in the group of aromatic compounds, and its hydrogenated is difficult. The Diels-Alder reactions or usual olefin reactions are not characteristic for this ring. Due to the fact, that it can easily polymerize, most of the electrophilic reaction, used in benzene chemistry, are not applicable to pyrroles. On the other hand, the substituted pyrrole derivatives have been included in various transformations [3].

REACTION OF PYRROLE WITH ELECTROPHILES

In general pyrroles most commonly react with electrophiles at α -position, due to the highest stability of the obtained intermediate protonation (Scheme 1).

From the group of electrophilic addition, pyrroles are easily nitrated, halogenated and sulfonated, where normally polyhalogenated derivatives are obtained, but monohalogenation may also occur (Scheme 2)[6]. When the pyrrole nitrogen is silylated, a halogenation on 3rd position is also possible, thus this is considered as a useful procedure for functionalization of the less active 3rd position [4].

Origin of pyrrole

In 1834 F.F. Runge has detected pyrrole for the first time as a constituent of coal tar [5]. Later in 1857 it has



Scheme 1. Mechanism of the reaction of pyrrole with electrophiles.

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