

OP24. ISOQUINOLINE ALKALOIDS AS A PROMISING FRAMEWORK FOR CHEMICAL MODIFICATIONS TO ENHANCE BIOACTIVITY

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Investigation of natural products is not limited only to the discovery and isolation of novel bioactive compounds; but equally important, it would provide valuable lead molecules that can be modified chemically and serve as a template for the design of semi-synthetic potent molecules. Numerous medications have been developed or inspired by the chemical modification of their natural parent molecules, such as development of atracurium from tubocurarine and dextromethorphan from morphine.

Isoquinoline alkaloids are one of the largest groups of natural products that have been known as fruitful sources in the drug discovery. Some isoquinoline alkaloids, such as galantamine and morphine, have found their place in therapy as naturally occurring molecules, and yet there are many other well-investigated isoquinoline alkaloids with promising bioactivity profiles, providing a natural potent scaffold. Derivatization and preparation of analogues of the natural isoquinoline-bearing alkaloids can be utilized not only as a beneficial strategy for improving the potency of these complex structures but also to improve the toxicity or bioavailability profile of the molecule, while avoiding the complicated and expensive total-synthesis procedure.

In this presentation, we will describe the perspective designed study for preparing semi-synthetic derivatives of a few selected potent isoquinoline alkaloids, which are available in natural sources in sufficient amounts, to enhance their targeted biological activity. Moreover, the possible challenges of chemical modification of alkaloids will be discussed, and achieved results will be presented.

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