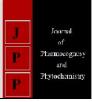


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Natural product repertoire of the marine actinomycetes *Nocardia* sp.

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Abstract

Actinomycetes are a very attractive and promising source in the production of novel bioactive compounds. They exhibit a wide range of pharmacological activities. The genus *Nocardia* consists of various species, such as *Nocardia brasiliensis*, *Nocardia fukayae*, *Nocardia interforma*, *Nocardia gloverula* and *Nocardia* sp. *Nocardia* sp is a rich source of metabolites containing diverse chemical classes, including benzoquinones and different classes of peptides as thiazolyl peptide and lipopeptides with various valuable biological activities. The review provides a survey for all the compounds isolated from *Nocardia* sp, along with their corresponding biological activities, whenever applicable.

Keywords: Nocardia sp, actinomycetes, peptides, antibiotic, anticancer

1. Introduction

Actinomycetes derived from marine source considered one of the richest source of secondary metabolites with therapeutic effect ^[1]. 13,700 bioactive compounds were identified from actinomycetes up to 2010^[2]. Sixty percent of the applied antibiotics between 1950 and 1970 were isolated from actinomycetes [3]. Nocardia sp (Kingdom: Bacteria: phylum: Actinobacteria; class: Actinobacteria; order: Actinomycetales; Suborder: Corynebacterineae; family: Nocardiaceae, Genus: Nocardia) is reported to be a very rich source of bioactive metabolites. Nocardia species, reported as an important producer of antibiotics ^[4]. Many chemical and biological examinations carried especially for Nocardia sp, found in the previous literature. These investigations confirmed that Nocardia sp is rich with different classes of natural products as peptides with various subclasses as thiazolyl peptides ^[5] and Lipopeptides ^[6], benzoquinones ^{[7],[8]} and picolinic acid ^[9]. *Nocardia* sp collected from different regions including Florida ^[6], Egypt ^[7], Japan ^[10]. Biological examination of *Nocardia* species provides that the most important activity is its antibacterial activity as it considered an antibiotic reservoir, moreover, it have antitumor ^[7, 11] and cytotoxic activities ^[12]. Our aim in this review is to present the chemical structures of the natural products isolated from Nocardia sp, their places of collection and their biological activities whenever applicable.

2. Peptides

Peptides are one of the most abundant metabolites from the marine actinomycete *Nocardia* sp. With different sub classes for example, lipopeptides as peptidolipins B–E (1–4), isolated from a marine *Nocardia* sp. (strain WMMB215), cultivated from the ascidian *Trididemnum orbiculatum*^[1]. They possess antibacterial activities as; they examined against methicillin-resistant *Staphylococcus aureus* (MRSA), methicillin-sensitive Staphylococcus aureus (MSSA) and showed that Peptidolipins B (1) and E (4) have antibacterial activity with MIC 64 µg/mL against MRSA and MSSA. While, Peptidolipins C (2), D (3) have an activity of more than 64 µg/mL with MIC against MSSA and MRSA ^[1]. Moreover, another subclass of peptides is thiazolyl peptide as nocathiacin I-III (5-7) ^[5] they were considered as an antibiotic complex . Previously reported from *Nocardia* sp. isolated from *Amycolatopsis* sp ^[5].

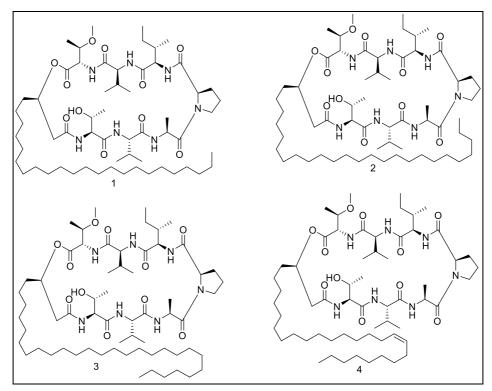


Fig 1: The chemical structure of compound (1-4)

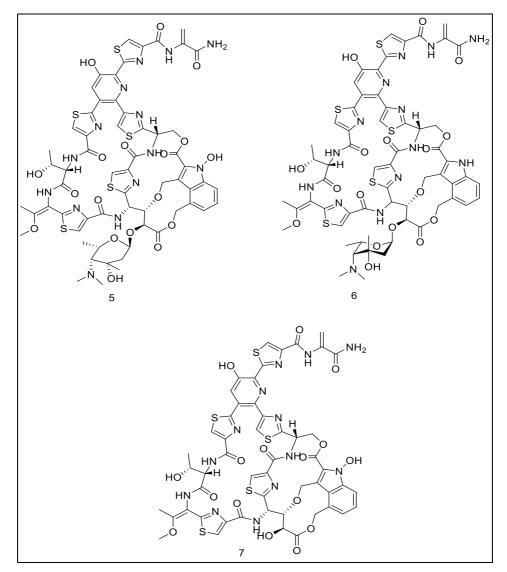


Fig 2: The chemical structures of the compounds (5-7)

3. Benzoquinones

Potent antibiotic antitumor compounds isolated from *Nocardia* sp is Macbecin I -II (8-9) considered a benzenoid ansamycins ^{[10].} Macbecin I-II have several biological activities as antiprotozoal, antibacterial and antifungal activities ^{[10].} Moreover, ubiquinone Q9 (10) ^[8] isolated from *Nocardia* sp. KMM 3749 was isolated from an unidentified marine sediment at Simushir Island, Kuril Islands ^{[8].}

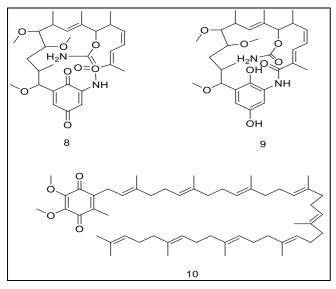


Fig 3: The chemical structures of the compounds (8-10)

4. Picolinic acid

Piclonic acid is a metabolite of tryptophan.one of its derivative is 5- Hydroxy picolinic acid (11) which is isolated from *Nocardia* sp. isolated from marine macrophytes at Japan ^[9]. Exhibited a variety of biological activities as immunological, neuroprotective anti-proliferative affects within the body ^{[13].}

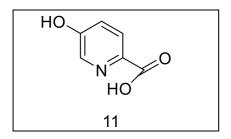


Fig 2: The chemical structures of the compounds (11)

5. Conclusions

Actinomycetes represents a huge reservoir of undiscovered natural products by having a wide range of pharmacological activities. *Nocardia* undescribed species considered an important source of secondary metabolite thus, new species discovery confirmed that there is still a chance in isolating new natural metabolites with expected strong biological activities. Peptides represents the highest distribution among the different chemical classes. Followed, by benzoquinones then acids. These chemical leads showed a huge diversity of pharmacological activities such as anticancer, antibacterial, and antiprotozoal activities, therefore, we must held further investigations for *Nocardia* sp. as it is a promising source for discovery of new bioactive natural metabolite.

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