

Cordyceps sinensis (Berk.) Sacc.

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48.1 Introduction

Dongchongxiacao, Cordyceps, is the dry complex composed of the sclerotium of the fungus *Cordyceps sinensis* (Berk.) Sacc. (Clavicipitaceae) and the larva corpses of insects of the family Hepialidae, on which the fungus is parasitic. It is officially listed in the Chinese Pharmacopoeia and is used as hemostatic, mycolytic, antiasthmatic, and expectorant in the treatment of respiratory diseases and as a tonic.

48.2 Chemical Constituents

The chemical constituents of *C. sinensis* were first studied by Chatterjee et al. in 1957 [1]. A crystalline substance was isolated and named "cordycepic acid", and was then identified by Sprecher and Sprinson as D-mannitol [2]. Further studies on the chemical constituents of Cordyceps revealed the presence of a series of known substances, but new structures or compounds with significant pharmacological efficacy were not found. The chemical constituents isolated from *C. sinensis* were amino acids, stearic acid, D-mannitol, mycose, ergosterol, uracil, adenine, adenosine [3,4], palmitic acid, cholesterolpalmitate and 5a-8a-epidioxy-5a-ergosta-6,22 dien-3 β -ol (48-1) [5].

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Since the production of *C. sinensis* has not satisfied demand, *C. hawkesii* was found as a substitute for *C. sinensis*. Comparison of the chemical constituents between *C. hawkesii* and *C. sinensis* based on thin-layer chromatography has shown that amino acid, alkaloid, sterol, and organic acid contents are similar [6,7]. Moreover, Cordyceps can be cultured submerged. Ergosterol, stearic acid, p-mannitol, mycose, uracil, uridine, adenine, adenosine. and 13 amino acids were separated in both the cultured broth and mycelium of *C. sinensis* 181.

In addition, *C. bamesii* [9, 10], *C. liangshanensis* [11], *C. militaris* [12], and *C. shanxiensis* [13] were investigated botanically and chemically. They were all used as substitutes for *C. sinensis*. The D-mannitol contents in *C. bamesii* and *C. sinensis* were 8.7% and 7.8%, respectively, and the ergosterol contents in *C. bamesii* and *C. sinensis* were 1.2% and 1.1% respectively. The contents of proteins, total amino acids, and alkaloids in both species were also similar [9].

Cultivation of *C. bamesii* was also reported [14].

From ascocarps of *C. sinensis*, a water-soluble polysaccharide was isolated and purified by ethanol fractionation and gel filtration. D-Galactose and D-mannose in a molar ratio of 1:1 were obtained by acid hydrolysis of purified polysaccharide. Chemical degradation and carbon 13 NMR spectrometric analysis showed that this polysaccharide is a highly branched galactomannan with a mannan core and galactosyl oligomer branches [15]. A water-soluble, minor galactomannan containing a small proportion of protein was further isolated from a 5% Na₂CO₃ extract of *C. sinensis*. It showed a homogeneous pattern in gel filtration and the molecular weight was estimated to be about 23,000. This minor polysaccharide was mainly composed of D-mannose and D-galactose in a molar ratio of 35 and had a highly branched structure [16].

48.3 Pharmacology

Both natural *Cordyceps* and cultured mycelia of *C. sinensis* have significant effects on the immune system of mice. They can increase the size of the spleen, decrease the size of the thymus, and prevent atrophy of spleen and liver and hypertrophy of the thymus in mice induced by cyclophosphamide [17]. The DNA, RNA, and protein contents in the enlarged spleen were significantly increased. The effect of the thymus was abolished by adrenalectomy. The extract increased the incorporation of [³H] thymidine into spleen DNA in vivo and the proliferation of splenocytes in vitro. The active principle was found to be present in the stroma rather than in the lanra [18].

The aqueous extracts of natural *Cordyceps* and the cultured mycelia of *C. sinensis* can enhance the production of macrophages and activate the functions of the phagocytic system. They not only enhance the phagocytic activity of the macrophages, but also increase the alkaline phosphatase activity of the macrophages [19]. In addition, serum hemolysin and splenocytic immunohemolytic activities were elevated in mice immune suppressed by hydrocortisone. In normal mice, however, no such regulatory effect on humoral immunity was observed [20].

The polysaccharide of *C. sinensis* also showed immunostimulating activity in mice. It activated the phagocytic function of the reticuloendothelial system and of macrophages in the abdominal cavity and increased blood serum IgG and plasma corticosterone levels and spleen weight. The polysaccharide also antagonized spleen atrophy and leukocyte decrease induced by cortisone and cyclophosphamide, and the reduction of phagocytic function of macrophages in the abdominal cavity, but did not inhibit the anti-inflammatory function of cortisone [21].

Some results of clinical studies with *C. sinensis* for treatment of tinnitus [22], chronic nephritis [23,24], arrhythmia [25], and sexual hypofunction [26] were recently reported. For instance, treatment with *C. sinensis* significantly decreased proteinuria of the patients with chronic nephritis in 24 h. Two patients with hematuria as the major clinical manifestation responded with strongly decreased erythrocyte counts in urine [23].

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