

Allicillin™



— With Stabilized and Standardized Allicin Metabolites —

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Designs for Health is pleased to present Allicillin™ with 100 mg Garlicillin™, and Allicillin Pro™ with 200 mg Garlicillin™.

Garlicillin™ is garlic oil macerate standardized to 1% ajoene (pronounced ah-hoe-ene) and also contains dithiins and allyl sulfides. These two Allicillin™ formulations are the first ever commercially available garlic oil macerate products containing a standardized level of ajoene, **the most active compound formed from garlic.**

What is Ajoene?

Ajoene is an organic trisulfur compound formed from three molecules of allicin¹. According to Dr. Eric Block, leading expert in garlic sulfur compounds, ajoene is the most active of the three heavily researched allicin metabolites that also include dithiins (pronounced di-thigh-ins) and allyl sulfides². Ajoene consists of two isomers (E and Z) and is chemically represented by 4,5,9-trithiadodeca-1,6,11-triene-9-oxide³. Since the discovery and identification of ajoene, there have been many studies that have demonstrated its related health benefits, which include:

- Anti-bacterial⁴⁻⁶
- Anti-fungal⁷⁻¹²
- Anti-parasitic¹³
- Anti-thrombotic and Anti-platelet¹⁴⁻¹⁵
- Anti-lipidemic¹⁶⁻¹⁷
- Anti-inflammatory¹⁸⁻¹⁹
- Anti-tumorigenic and Anti-mutagenic²⁰⁻²²

What is Garlic Oil Macerate?

Garlic oil macerate is a popular health food in Europe^{5, 23}. It is prepared by mixing mashed or chopped garlic in vegetable oil. Garlic oil macerate, which has been a dietary supplement for many decades, is the only garlic supplement that contains significant quantities of ajoene and dithiins^{5, 23-24}. Lawson et al. reported that garlic oil macerates contain between 15 to 115 microgram/gram of ajoene and 70 to 690 microgram/gram dithiins²⁴.

Garlicillin™ is a powdered ingredient from garlic oil macerate that is currently standardized with 1% ajoene (10,000 mcg/g), and also contains significant levels of dithiins and allyl sulfides. Allicillin Pro™ with 200mg Garlicillin™ is up to over 2000 times more potent than a typical 500 mg non-standardized garlic oil macerate product. **No other commercially available garlic preparation contains near the level of ajoene available from Allicillin Pro™.**

What about Allicin?

The chemistry of garlic is extremely complex, but research has shown that it is garlic's unique organosulfur compounds that promote its broad range of potential health benefits. Garlic has many bioactive components, the best known and studied of which is allicin. It is ironic that allicin does not exist in fresh, undamaged garlic cloves. The predominant garlic sulfur compound found in the garlic plant is alliin. Garlic also contains high levels of an enzyme called allinase. Alliin and allinase are held in different compartments of the garlic plant, by design, to react only when the plant is injured. When fresh garlic cloves are crushed or chopped, or when garlic powder (that has been carefully dried to preserve its alliin/allinase content) is added to water, allicin is produced in seconds by the action of allinase on alliin²⁵.

Many dietary supplement companies claim to provide a product that delivers allicin. Due to its instability allicin is often listed on labels as "allicin potential" or "allicin yield". Allicin potential is measured in a laboratory by using dried garlic powder that is added to water so that the alliin and allinase can quickly react to form allicin. The amount of allicin produced is the measure of allicin potential. However, the situation is very different when such garlic supplements are swallowed. The allinase enzyme is rapidly and completely destroyed by stomach acid. Allicin cannot be made from alliin in the absence of allinase enzyme. Some garlic products claim to address this issue by using an enteric coated delivery method. Unfortunately, such methods do not work well at all. Lawson and Wang reported the results of testing twenty-three enteric coated U.S. garlic supplements in 2001²⁵. Twenty of twenty-three failed to

Allicillin Pro™ 200 mg

Supplement Facts

Serving Size 1 capsule
Servings Per Container 60

Amount Per Serving	% Daily Value
Garlicillin™	200 mg *
[Garlic oil macerate standardized to contain [1% ajoene from natural allicin; [also contains dithiins]	

*Daily Value not established.

Other Ingredients: Microcrystalline cellulose, rice flour, magnesium stearate, tapioca.

release even 15 percent of their claimed allicin potential when placed in simulated intestinal fluid. Lawson and Wang concluded that allicin potential is an extremely poor measure of garlic supplement activity in the human body and should not be used for standardization of garlic supplements. Considering the questionable utility of allicin potential, technology was developed to produce the inherently stable metabolite of allicin, ajoene.

The Research on Ajoene is Broad and Impressive

In 1983, Apitz-Castro et al. isolated three garlic compounds that inhibited human platelet aggregation²⁶. Two of the compounds were identified as an allyl sulfide and a dithiin but the third, which was four times more potent than the other two, was not clearly identifiable. Subsequently that third unidentified compound was later named ajoene from "ajo" the Spanish word for garlic, by Block et al (1984)³. From published literature search it is apparent that the antimicrobial (antibacterial and antifungal) properties of ajoene have received considerable attention. Studies show that ajoene exhibits broad spectrum antimicrobial activity²⁷. Naganawa et al., found that ajoene inhibited the growth of gram positive bacteria such as *Bacillus cereus*, *B. subtilis*, *Mycobacterium smegmatis* and *Streptomyces griseus* at 5 ug ajoene per ml and *Staphylococcus aureus* and *Lactibacillus plantarum* below 20 ug per ml⁴. They also reported that growth of gram-negative bacteria such as *Escherichia coli*, *Klebsiella pneumonia* and *Xanthomonas maltophilia* were also inhibited by ajoene at higher doses, 100 to 160 ug/ml. Ajoene from garlic oil macerate likewise inhibited the growth of *Helicobacter pylori* at 10 to 25 ug/ml⁶.

Yoshida et al showed that ajoene is more effective than allicin against Aspergillus niger (16.6 ug/ml vs. 30.9 ug/ml) and Candida albicans (7.6 ug/ml vs. 17.3 ug/ml)¹². In clinical studies ajoene was shown to be as effective or better than terbinafine in the treatment of tinea pedis⁹, tinea corporis, and tinea cruris¹⁰. Ajoene was also effective against *Paracoccidioides brasiliensis*, a common fungus that causes systemic mycoses in Latin America^{7, 11}. When tested against *Scedosporium prolificans*, a fungus that is very difficult to treat, ajoene had a minimum inhibition concentration (MIC) of 2.0 to 8.0 mg/l compared to 2.0 to >16 mg/l for amphotericin B and >16 mg/l for itraconazole⁸. Incredibly, ajoene has even successfully treated malaria in an in vivo animal model¹³.

Ajoene and other garlic extracts including allicin were tested in vitro against several viruses including herpes simplex virus type 1, herpes simplex virus type 2, parainfluenza virus type 3 and human rhinovirus type 2²⁸. **Ajoene was found to have the greatest virucidal activity compared to allicin and other garlic extracts tested.** Ajoene and dithiins have chemopreventive properties with their ability to inhibit aflatoxin B1-induced mutagenesis²⁹.

Who should take Allicillin™?

Patients with recurring yeast infections, bacterial or viral infections, lipid abnormalities, platelet aggregation, inflammation, immune deficiency and/or history of cancer or heart disease. Consider Allicillin™ supplementation during antibiotic usage to prevent yeast overgrowth, a common side effect of antibiotic therapy. Allicillin Pro™ may help improve symptoms of Lyme disease. This product may be used in high doses for acute conditions and can be taken as directed daily for prevention.

Allicillin Pro™ has blood thinning capabilities and reduces platelet aggregation. Be cautious when recommending to patients taking Coumadin (warfarin) or other anti-coagulant medications.

References

1. Managoli, N. Local Drug Delivery System in Coronary Stents. International Patent Application No. PCT/IN2002/000173. Publication No. WO/2003/018082.
2. Passwater R. The chemistry of garlic health benefits. Whole Foods 15(6) 22-26 (June 1992), www.drpasswater.com.
3. Block E, Ahmad S, Jain MK, Creceley RW, Apitz-Castro R, Cruz MR. (E,Z)-Ajoene: A potent antithrombotic agent from garlic. J Am Chem Soc. 1984; 106:8295-6.
4. Naganawa R, Iwata N, Ishikawa K, et al. Inhibition of microbial growth by ajoene, a sulfur-containing compound derived from garlic. Appl Environ Microbiol. 1996 Nov;62(11):4238-42.
5. Yoshida H, Katsuzaki H, Ohta R, et al. An organosulfur compound isolated from oil-macerated garlic extract, and its antimicrobial effect. Biosci Biotechnol Biochem. 1999 Mar;63(3):588-90.
6. Ohta R, Yamada N, Kaneko H, et al. In vitro inhibition of the growth of *Helicobacter pylori* by oil-macerated garlic constituents. Antimicrob Agents Chemother. 1999 Jul;43(7):1811-2.
7. Thomaz L, Apitz-Castro R, Marques AF, et al. Experimental paracoccidioidomycosis: alternative therapy with ajoene, compound from *Allium sativum*, associated with sulfamethoxazole/trimethoprim. Med Mycol. 2008 Mar;46(2):113-8.
8. Davis SR, Perrie R, Apitz-Castro R. The in vitro susceptibility of *Scedosporium prolificans* to ajoene, allitridium and a raw extract of garlic (*Allium sativum*). J Antimicrob Chemother. 2003 Mar;51(3):593-7.
9. Ledezma E, Marciano K, Jorquera A, et al. Efficacy of ajoene in the treatment of tinea pedis: a double-blind and comparative study with terbinafine. J Am Acad Dermatol. 2000 Nov;43(5 Pt 1):829-32.
10. Ledezma E, López JC, Marin P, et al. Ajoene in the topical short-term treatment of tinea cruris and tinea corporis in humans. Randomized comparative study with terbinafine. Arzneimittelforschung. 1999 Jun;49(6):544-7.
11. San-Blas G, San-Blas F, Gil F, Mariño L, Apitz-Castro R. Inhibition of growth of the dimorphic fungus *Paracoccidioides brasiliensis* by ajoene. Antimicrob Agents Chemother. 1989 Sep;33(9):1641-4.
12. Yoshida S, Kasuga S, Hayashi N, et al. Antifungal activity of ajoene derived from garlic. Appl Environ Microbiol. 1987 Mar;53(3):615-7.
13. Perez HA, De la Rosa M, Apitz R. In vivo activity of ajoene against rodent malaria. Antimicrob Agents Chemother. 1994 Feb;38(2):337-9.
14. Apitz-Castro R, Badimon JJ, Badimon L. Effect of ajoene, the major antiplatelet compound from garlic, on platelet thrombus formation. Thromb Res. 1992 Oct 15;68(2):145-55.
15. Apitz-Castro R, Ledezma E, Escalante J, Jain MK. The molecular basis of the antiplatelet action of ajoene: direct interaction with the fibrinogen receptor. Biochem Biophys Res Commun. 1986 Nov 26;141(1):145-50.
16. Ferri N, Yokoyama K, Sadilek M, et al. Ajoene, a garlic compound, inhibits protein prenylation and arterial smooth muscle cell proliferation. Br J Pharmacol. 2003 Mar;138(5):811-8.
17. Gebhardt R, Beck H, Wagner KG. Inhibition of cholesterol biosynthesis by allicin and ajoene in rat hepatocytes and HepG2 cells. Biochim Biophys Acta. 1994 Jun 23;1213(1):57-62.
18. Dirsch VM, Vollmar AM. Ajoene, a natural product with non-steroidal anti-inflammatory drug (NSAID)-like properties? Biochem Pharmacol. 2001 Mar 1;61(5):587-93.
19. Dirsch VM, Kierner AK, Wagner H, Vollmar AM. Effect of allicin and ajoene, two compounds of garlic, on inducible nitric oxide synthase. Atherosclerosis. 1998 Aug;139(2):333-9.
20. Taylor P, Noriega R, Farah C, et al. Ajoene inhibits both primary tumor growth and metastasis of B16/BL6 melanoma cells in C57BL/6 mice. Cancer Lett. 2006 Aug 8;239(2):298-304. Epub 2005 Oct 10.
21. Nishikawa T, Yamada N, Hattori A, et al. Inhibition by ajoene of skin-tumor promotion in mice. Biosci Biotechnol Biochem. 2002 Oct;66(10):2221-3.
22. Ishikawa K, Naganawa R, Yoshida H, et al. Antimutagenic effects of ajoene, an organosulfur compound derived from garlic. Biosci Biotechnol Biochem. 1996 Dec;60(12):2086-8.
23. Hattori A, Yamada N, Nishikawa T, et al. Protective effect of ajoene on acetaminophen-induced hepatic injury in mice. Biosci Biotechnol Biochem. 2001 Nov;65(11):2555-7.
24. Lawson, LD, Wang ZJ, Hughes B. Identification and HPLC quantitation of the sulfides and dialk(en)yl thiosulfonates in commercial garlic products. Planta Med. 1991. 57:363-370.
25. Lawson, L. & Wang, Z. "Low allicin release from garlic supplements: a major problem due to the sensitivities of allinase activity" J Agric Food Chem 49(2001): 2592-99.
26. Apitz-Castro R, Cabrera S, Cruz MR, et al. Effects of garlic extract and of three pure components isolated from it on human platelet aggregation, arachidonate metabolism, release reaction and platelet ultrastructure. Thromb Res. 1983 Oct 15;32(2):155-69.
27. Rose P, Whiteman M, Moore PK, Zhu YZ. Bioactive S-alk(en)yl cysteine sulfonolipids in the genus *Allium*: the chemistry of potential therapeutic agents. Nat Prod Rep. 2005 Jun;22(3):351-68. Epub 2005 May 10.
28. Weber ND, Andersen DO, North JA, et al. In vitro virucidal effects of *Allium sativum* (garlic) extract and compounds. Planta Med. 1992 Oct;58(5):417-23.
29. Tadi PP, Teel RW, Lau BH. Organosulfur compounds of garlic modulate mutagenesis, metabolism, and DNA binding of aflatoxin B1. Nutr Cancer. 1991;15(2):87-95.