

CYTOCHALASIN

Product Number C6637, D1641, C6762, C0138, C8273, C1139, C0889, C2149
Storage Temperature -20°C

CAS #: See table

Product Description

See Table 1

The Cytochalasins (Greek *cytos*, cell; *chalasis*, relaxation) are a group of related fungal metabolites. They were discovered in 1964 during the screening of mold filtrates for possible biological activity on cells. These fungal toxins are related by chemical structure. All are characterized by a highly substituted hydrogenated isoindole ring to which is fused a macrocyclic ring. The macrocyclic ring may vary from 11 to 14 atoms and may be either a carbocycle or lactone. These fungal toxins also share a number of unusual, interesting, and characteristic effects on the animal cell.

Cytochalasin B (Product No.C6762) is a metabolite of the fungus *Drechslera* (previously *Heiminthosporium*) *dematioideum*. It was originally isolated from cultures of a *Phoma* species and, therefore, was sometimes referred to as phomin. Cytochalasin B is cell membrane permeable. It inhibits cell division by blocking formation of contractile microfilaments. It inhibits cell movement 1.2 and induces nuclear extrusion. 1.2,3,4 It shortens actin filaments by blocking monomer addition at the fast growing end of the polymer. It impairs maintenance of long term potentiation (LTP) of action filaments. It inhibits glucose transport 6,7,8,33 and platelet aggregation. It inhibits glucose transport 1,2,3,4 and platelet aggregation. It blocks adenosine-induced apoptotic body formation without affecting activation of endogenous ADP-ribosylation in leukemia LH-60 cells.

Dihydrocytochalasin B (dihydro-CB) (Product No. D1641), the saturated derivative of Cytochalasin B, induces changes in morphology and motility, but has little effect on sugar transport. 14,15,16 Dihydrocytochalasin B and its γ -lactone are useful probes for studying cytochalasin binding sites. 17,18

Dihydrocytochalasin B γ -lactone does not appear to have the same effects on cell motility and morphology as Cytochalasin B or Dihydrochalasin B. Like Dihydrochalasin B, the gamma-lactone does not appear to inhibit glucose transport.

ProductInformation

Cytochalasin A (Product No. C6637) is a metabolite of the fungus *Drechslera* (previously *Heiminthosporium*) *dematioideum*.¹⁹ Cytochalasin A is sulfhydryl-reactive. It was shown to inhibit growth and sugar uptake in a Saccharomyces strain.²⁰ Unlike Cytochalasin B, Cytochalasin C (Product No. C0138) and Cytochalasin D (Product No. C8273) are isomeric metabolites of *Metarrhizium anisopliae*.²¹ The cytochalasin D possesses antibiotic²² and antitumor²³ activity. It also impairs maintenance of long term potentiation (LTP) of actin filaments.³² It is implicated in promoting conditions favorable for depolymerizing actin.³⁴

Cytochalasin E (Product No. C2149) is a metabolite of *Rosellinia necatrix*. ^{24,25} Cytochalasin E is unique in producing a "halo" around the nucleus more often than nuclear extrusion. ⁴ Cytochalasin E is a inhibitor of angiogenesis and tumor growth. ³⁰

Cytochalasin H (Product No. C0889) and Cytochalasin J (Product No. C1139) are metabolites of *Phomopsis paspali* found on *Paspalum scrobiculatum* Linn. (a millet consumed in India). Cytochalasins H and J have shown Central Nervous System activity. All Cytochalasin J blocks or slows chromosome motion and affects spindle architecture.

Disclaimer/Precautions

Cytochalasins are regarded as highly toxic and possible teratogens. Handle in a manner to avoid/minimize direct body contact and inhalation.

Preparation Instructions

Solubility

492 mg/ml in dimethylformamide at room temperature 371 mg/ml in dimethyl sulfoxide (DMSO) at room temperature

35 mg/ml in ethanol at room temperature 10 mg/ml in acetone at room temperature Essentially insoluble in water

It is advisable to make a 1000X stock solution in DMSO (the final concentration of DMSO in the aqueous medium should not exceed 0.1% because greater DMSO concentrations can adversely affect many cultured cells). Dilute the stock in the appropriate aqueous medium to provide a physiologically acceptable final concentration (must be within the low

solubility limit of cytochalasins in the chosen aqueous medium). The physiologically desired working concentrations vary for different applications. Examples: 10 μM Cytochalasin B can completely block adenosine-induced apoptotic body formation in cultured HL-60 cells. 13 According to Theodoropoulos 5 , 30 μM Cytochalasin B can shorten actin filaments by blocking monomer addition at the fast growing end of the polymer. 5

Storage/Stability

Cytochalasin B is a solid believed to be photostable in the solid form and reasonably stable in solution. Solutions of Cytochalasin B in dimethyl sulfoxide have shown no decrease in potency when stored at 4°C for more than three years.²⁹

Cytochalasin A, C, D and E should be stored in the dark since the conjugated double bond undergoes slow isomerization from *trans* to *cis* in the presence of light.

References

- 1. Carter, S.B., Nature, 213, 261 (1967).
- 2. Krishan, A. J., Cell Biol., 54, 657 (1972).
- 3. Prescott, D.M. *et al.*, J. Exp. Cell Res., **71**, 480 (1972).
- 4. Carter, S.B., Endeavor, 31, 77 (1972).
- 5. Theodoropoulos, P.A. *et al.*, Biochem. Pharmacol., **47**(10), 1875 (1994).
- Kletzien, R.F. et al., J. Biol. Chem., 247, 2964 (1972).
- 7. Mizel, S.B. and Wilson, L., J. Biol. Chem., **247**, 4102 (1972).
- 8. Estensen, R.D. and Plagemann, P.G., Proc. Nat. Acad. Sci. U.S.A., **69**, 1430 (1972).
- 9. Shepro, D. et al., J. Cell Biol., 47, 544 (1970).
- White, J.G., Roussel Conference on Platelet Aggregation, Masson, Paris, 4th March, 1971.
- 11. Haslam, R.J., Biochem. J., 127, 34P, (1972).

- 12. Majno, G. *et al.*, Thromb. Diath. Haemorrh. **28**, 49 (1972).
- 13. Tanaka, Y. et al., Exp. Cell Res., 213, 242 (1994).
- 14. Atlas, S.J. and Lin, S., J. Cell Biol., 76, 360 (1978).
- 15. Lin, S. *et al.*, Proc. Nat. Acad. Sci. U.S.A., **75**, 329 (1978).
- 16. Lin, S. and Spudich, J.A., J. Biol. Chem., **249**, 5778 (1974).
- 17. Lin, D.C. and Lin, S., J. Biol. Chem., **253**, 1415, (1978).
- 18. Rampal, A.L. et al., Biochemistry, 19, 679, (1980).
- 19. Aldridge, D.C. et al., J. Chem. Soc. (C):1667, 1967.
- 20. Kuo, S. –C. and Lampen, J.O., Ann. N.Y. Acad. Sci., **235**, 137, (1974).
- 21. Aldridge, D.C. and Turner, W.B., J. Chem. Soc. (C) 923, (1969).
- Betina, V. and Micekova, D.Z. Allg. Mikrobiol., 12, 355, (1972) and Chem. Abstr. 77, 160508q, (1972).
- 23. Katagiri, K. and Matsuura, S., J. Antibiot., **24**, 722, (1971).
- 24. Aldridge, D.C. *et al.*, Chem. Commun. p. 148, (1972).
- 25. Aldridge, D.C. *et al.*, Chem. Commun. p. 551, (1973).
- 26. Pendse, G.S., Experientia, 30, 107, (1974).
- 27. Padwardhan, S.A. *et al.*, Phytochemistry, **13**, 1985 (1974).
- 28. Deshmukh, P.G *et al.*, Acta. Microbiol. Acad. Sci. Hung., **22**, 253 (1975).
- 29. Sigma Data.
- 30. Udagaura, T., *et al.*, J. Pharmacol. Exp. Ther., **294**, 421-427 (2000).
- 31. Snyder, J.A., and Cohen, L., Cell Motil. Cytoskeleton, **32**, 245-257 (1995).
- 32. Krucker, T. *et al..*, Proc. Natl. Acad. Sci. USA, **97**, 6856 (2000).
- 33. Lachaael, M. *et al.*, J. Biol. Chem., **271**, 5225 (1996).
- Dubinsky, W.P. *et al.*, Proc. Natl. Acad. Sci. USA, 96, 9421, (1999).

CMK 03/31/99

Table 1

Product Number	CAS#	Appearance	Molecular Formula	Molecular Weight in Daltons	Melting Point	Specific Rotation
C6637	14110-64-6	White powder	$C_{29}H_{37}NO_5$	477.61	193-195°C	
C0138	22144-76-9	White powder	$C_{30}H_{37}NO_6$	507.63	260°C	
C6762	14930-96-2	White powder	C ₂₉ H ₃₇ NO ₅	479.62	218-221°C	+86.7 (039% solution w/v in MeOH @ 21°C)
D1641	39156-67-7	White powder	C ₂₉ H ₃₇ NO ₅	481.64	198-203°C	
C1139	56144-22-0	Faint yellow powder	C ₂₈ H ₃₇ NO ₄	451.61	137-139°C	+42.2 (0.97 g/100 ml CH ₃ OH)
D6016	14110-71-5	White powder	C ₂₉ H ₃₇ NO ₅	481.64	192-193°C	
C2149	36011-19-5	White powder	C ₂₈ H ₃₃ NO ₇	495.58	206°C	-25.6 (1g/100 ml MeOH @ 25°C)
C8273	22144-76-9	White powder or white powder with yellow cast	C ₃₀ H ₃₇ NO ₆	507.63	268-271°C	-7.5 (55% solution w/v in dioxane @ 25°C)
C0889	53760-19-3	White powder	$C_{30}H_{39}NO_{6}$	493.63	268-271°C	

$$CH_{2} - CH_{2}$$
 $CH_{2} - CH_{2}$
 $CH_{2} - CH$
 $CH_{3} - CH$
 $CH_{4} - CH$
 $CH_{5} - CH$
 $CH_{5} - CH$
 $CH_{5} - CH$
 $CH_{6637} - CH$
 $CH_{2} - CH_{2}$
 $CH_{2} - CH_{2}$
 $CH_{3} - CH$
 $CH_{2} - CH$
 $CH_{2} - CH$
 $CH_{3} - CH$
 $CH_{2} - CH$
 $CH_{4} - CH$
 $CH_{5} - CH$
 $CH_{5} - CH$
 $CH_{6637} - CH$
 CH_{6637

$$O=C$$
 CH_3
 $O=C$
 CH
 CH_3
 CH_3
 CH_4
 CH_5
 CH_5
 CH_5
 CH_6
 CH_7
 CH_7
 CH_7
 CH_7
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8
 CH_8