

Table S1. Alkaloids from endophytic fungi and their biological activities, metabolite class, fungus, host plant(s), reference.

Metabolite Class	Fungus	Host Plant(s)	Compounds Isolated	Biological Target	Biological Activity	Reference
Cytochalasans	<i>Xylaria</i> sp.	<i>Casearia sylvestris</i>	Cytochalasins B (1)	AChE inhibitory activity	Potent AChE inhibition at 60	13
			Cytochalasin C (2)		µg	
			Cytochalasin D (3)		Inactive	
	<i>Aspergillus terreus</i>	<i>Artemisia annua</i>	Cytochalasin E (4)	AChE inhibitory activity	IC ₅₀ , 146.1±6.5, 176.0±11.5, and	14
			5,6-Dehydro-7-hydroxy cytochalasin E (5)		110.9±13.7 µM, respectively	
			Δ6,12-Isomer of 5 (6)			
			Rosellichalasin (7)		IC ₅₀ > 200 µM	
	<i>Phomopsis</i> sp.	<i>Senna spectabilis</i> (Fabaceae)	Cytochalasins J (8)	AChE inhibitory activity	Inactive	15
	<i>Phomopsis</i> sp., <i>Phomopsis</i> IFB-E060	<i>Senna spectabilis</i> (Fabaceae), <i>Vatica mangachapoi</i>	Cytochalasins H (9)	AChE inhibitory activity	Minimum quantity of 25.0 µg	15, 18
	<i>Aspergillus flavipes</i>	<i>Hevea brasiliensis</i>	Bisaspochalasin D (10)	Neurite growth activity of PC12 cells	Inducing a differentiation rate of 12.52% at 10 µM	16
			Bisaspochalasin E (11)			
			Aspochalasin D (12)		Inactive	
			Aspochalasin B (13)			
	<i>Chaetomium globosum</i>	<i>Panax notoginseng</i>	Chaetoglobosin A (14)	AChE inhibitory activity	Inhibition rate < 10% at 50 µM	17
Chaetoglobosin B (15)						
Chaetoglobosin E (16)			Inactive			
Chaetoglobosin F (17)						

			Chaetoglobosin Fex (18)			
			Penochalasin F (19)			
			Penochalasin G (20)			
Chaetomium globosum WQ	Imperata cylindrical		Cytoglobosin A (21)		EC ₅₀ , 0.879 mM, 472.87 mM, and 0.023 mM, respectively	18
			Penocha-lasin C (22)	Scavenging DPPH and ABTS activity,	E _{max} , 7.1%, 5.9% and 40.3%, respectively	
			Isochaetoglobosin D (23)	H ₂ O ₂ -induced PC12 cell damage inhibitory activity	EC ₅₀ , 0.002 mM, 0.002 mM, and 0.009 mM, respectively	
Phomopsis IFB-E060	sp.	Vatica mangachapoi	18-Methoxycytochalasin J (24)		EC ₅₀ : 481.121 mM, 1.052 mM, and 0.004 mM, respectively	
Westerdykella nigra	Avicennia marina (Forssk.) Vierh		Westalsan (25)		IC ₅₀ , 0.056±0.003, 0.088±0.005 and 0.140±0.007 μM, respectively	19
			Phomacin B (26)	AChE inhibitory activity		
			19-Hydroxy-19,20-dihydro phomacin C (27)			
Diketopiperazine Derivatives	Acrostalagmus luteoalbus TK-43	Codium fragile	Acrozine A (28, 29)		IC ₅₀ , 9.5, 60.7 and 130.5 μM, respectively	20
			Acrozine B (30, 31)			
			Acrozine C (32, 33)			
			Acrozine D (34)	AChE inhibitory activity	Weak activity at 32 μM	21
			Acrozine E (35)			
			Acrozine F (36)		IC ₅₀ , 8.4 μM	
			Acrozine G (37)		Weak activity at 32 μM	
			Pseudellone D (38)			
			Lasiodipine E (39)	–	–	
Chetoseminudin C (40)						

			Chetoseminudin B (41)			
			T988 C (42)			
			T988 B (43)			
<i>Aspergillus tetteus</i>	<i>Artemisia annua</i>		Acetylpoaranotin (44)	AChE inhibitory activity	IC ₅₀ , 127.4±17.3µM	14
<i>Neosartorya fischeri</i>	<i>Glehnia littoralis</i>		Cyclotryprostatin B (45)	Neuroprotective activity		22
<i>JS0553</i>			Fumitremorgin B (46)	against	Obvious neuroprotection at 20	
			Fumitremorgin A (47)	glutamate-mediated HT22 cell death.	µM	
			Fumitremorgin C (48)			
			Brevianamide F (49)			
			Spirotryprostatin A (50)			
<i>Talaromyces</i>	sp.		6-Methoxyspirotryprostatin B (51)	AChE inhibitory activity	Inactive	
<i>LGT-2</i>	<i>Tripterygium wilfordi</i>		3-Dehydroxymethylbisdet			23
			hio-3,10a-bis(methylthio)gliotoxin (52)			
			Bisdethiobis(methylthio)gliotoxin (53)		Weak activity	
			Didehydrobisdethiobis(methylthio)gliotoxin (54)	–	–	
<i>Nigrospora</i>			Nigrosporaamide A (55)			
<i>camelliae-sinensis</i>	<i>Lumnitzera littorea</i>		Nigrosporaamide B (56)	H ₂ O ₂ -mediated cytotoxicity	Inactive at 100 µM	24
<i>S30</i>			Cyclo-(L-Pro-L-Phe) (57)	for HT22 cell		
			Cyclo[L-(4-hydroxyprolinyl			

l)-L-Leu] (58)						
<i>Nigrospora camelliae-sinensis</i> S30, <i>Penicillium</i> sp.1	<i>Lumnitzera littorea</i> , <i>Alibertia macrophylla</i> (Rubiaceae)	Cyclo-(L-Val-L-Pro) (59)	AChE inhibitory activity	A detection limit of 10.0 µg	24, 25	
<i>Nigrospora camelliae-sinensis</i> S30	<i>Alibertia macrophylla</i> (Rubiaceae)	Cyclo-(L-Leu-L-Pro) (60)	H ₂ O ₂ -mediated cytotoxicity for HT22 cell	Inactive at 100 µM	24	
		Cyclo-(R-Leu-R-Pro) (61)				
		Cyclo-(L-Ile-L-Pro) (62)				
		Cyclo-(4-methyl-R-Pro-S-N va) (63)				
<i>Colletotrichum gloeosporioides</i>	<i>Michelia champaca</i>	Cyclo-(S-Pro-S-Tyr) (64)	AChE inhibitory activity	Moderate activity at 200 µg	26	
		Cyclo-(S-Pro-S-Val) (65)				
<i>Colletotrichum crassipes</i>	<i>Casearia sylvestris</i>	Cyclo(D)-Pro-(L)-Val (66)	AChE inhibitory activity	Inactive at 100 µg	13	
		Cyclo(D)-Pro-(D)-Tyr (67)				
		Cyclo(D)-Val-(D)-Tyr (68)				
		Cyclo(D)-Hyp-(L)-Ile (69)				
		Cyclo(D)-Pro-(D)-Leu (70)				
		Cyclo(D)-Pro-(L)-Ile (71)				
		Cyclo(D)-Pro-(L)-Phe (72)				
		Cyclo(D)-Pro-(D)-Phe (73)				
Indole		16α-Hydroxy-5N-acetylardeemin (74)	AChE inhibitory activity	IC ₅₀ , 58.3±3.3, 149.4±11.3, and 116.9±3.7 µM, respectively	14	
Alkaloids	<i>Aspergillus terreus</i>	<i>Artemisia annua</i>				5N-Acetylardeemin (75)
						15b-β-Hydroxy-5N-acetylardeemin (76)
<i>Colletotrichum</i>	<i>Michelia champaca</i>	2-Phenylethyl	AChE inhibitory activity	Moderate activity at 200 µg	26	

<i>gloeosporioides</i>			1H-indol-3-yl-acetate (77)			
<i>Penicillium chrysogenum</i> No. 005	<i>Cistanche deserticola</i>		Chrysogenamide A (78)	Neuroprotection for H ₂ O ₂ -mediated SH-SY5Y cell death	Enhancing cell viability of 59.6% at 1×10 ⁻⁴ μM	27
			9-Deacetylfumigaclavine C (79)			
<i>Aspergillus fumigatus</i>	<i>Cynodon dactylon</i>		9-Deacetoxymumigaclavine C (80)	Neurite outgrowth of PC12	Inactive	28
			Fumigaclavine C (81)			
<i>Neosartorya fischeri</i> JS0553	<i>Glehnia littoralis</i>		Aszonalenin (82)	Neuroprotection against glutamate-mediated HT22 cell death	Inactive	22
			Acetylazonalenin (83)			
<i>Alternaria alternate</i>	<i>Psidium littorale</i> Raddi		Alternatine A (84)	PC12 cell injury induced by glutamate	Inactive	29
<i>Alternaria. Alternate/Epicoccum nigrum</i>	<i>Psidium littorale</i> Raddi/ <i>Houttuynia cordata</i>		1H-indole-3-carboxylic acid (85)	Scavenging DPPH and ABTS activity	IC ₅₀ , 88.97 μg/mL for DPPH, EC ₅₀ , 21.48 ± 0.88 μM for ABTS	29,30,31
<i>Alternaria alternate</i>	<i>Psidium littorale</i> Raddi		Indole-3-methylethanoate (86)	Neuroprotection on glutamate-mediated PC12 cell death	Cell viabilities improved with 75.6 ± 4.2% and 84.8 ± 6.5 at 40 and 80 μM	29
			Colletotryptin A (87)			
			Colletotryptin B (88)			
			Colletotryptin C (89)			
<i>Colletotrichum</i> sp. SC1355	<i>Cleistocalyx operculatus</i>		Colletotryptin D (90)	AChE inhibitory activity	Inactive at 50 μM	32
			Colletotryptin E (91/92)			
			Colletotryptin F (93)			

Other of Alkaloids	Types	<i>Colletotrichum gloeosporioides</i>	<i>Michelia champaca</i>	Uracil (94)	AChE inhibitory activity	Moderate activity at 200 µg	26
				4-Hydroxy-benzamide (95)			
		<i>Botryosphaeria dothidea</i> KJ-1	<i>Melia azedarach</i> L	3-Hydroxy-2-methoxy-5-methylpyridin-2(1H)-one (96)			
		<i>Nigrospora camelliae-sinensis</i> S30	<i>Lumnitzera littorea</i>	5-(40-Hydroxybenzyl)hydantoin (97)	H ₂ O ₂ -mediated cytotoxicity for HT22 cell	Inactive	24
				(±)-Oxypenicinoline (98/99)	A		
				(±)-Oxypenicinoline (100/101)	B		
				(±)-Oxypenicinoline (102/103)	C		
				(±)-Oxypenicinoline (104/105)	D	Inactive	
		<i>Penicillium steckii</i> SCSIO 41025	<i>Avicennia. Marina</i> (Forsk.) Vierh (Trichomaceae)	Penicnoline F (106)	AChE inhibitory activity		34
				Penicnoline G (107)			
				1,2,3,11b-Tetrahydroquinolactacide (108/109)			
				Quinolactacide (110)			
				Penicnoline (111)		IC ₅₀ , 87.3 µM	
				Methyl-penicnoline (112)		Inactive	
				Penicnoline E (113)		IC ₅₀ , 68.5 µM	
				Quinolonimide (114)		Inactive	
				4-Oxo-1,4-dihydroquinolin			

			e-3-carboxamide (115)			
			Huptremule A (116)			
			Huptremule B (117)			
<i>Ceriporia lacerate</i> HS-ZJUT-C13A	<i>Huperzia Serrata</i>		Huptremule C (118)	AChE inhibitory activity	IC ₅₀ , 0.99±0.03, 2.17±0.05, 0.11±0.01, 0.06±0.00, and 12.11±0.34 μM, respectively	35
			Huptremule D (119)			
			8α,15α-Epoxyhuperzine A (120)			
<i>Aspergillus terreus</i> (No. GX7-3B)	<i>Bruguiera gymnoihiza</i>		8-O-methylbostrycoidin (121)	AChE inhibitory activity	IC ₅₀ , 6.71 μM	36
<i>Fusarium</i> sp. HP-2	Chinese "Qi-Nan"	agarwood	Lumichrome (122)	AChE inhibitory activity	Inactive at 50 μM	37
<i>Phomopsis</i> sp. xy21	Thai	<i>Xylocarpus granatum</i>	Phomopsol A (123)	Neuroprotection corticosterone-mediated PC12 cell injury	Cell viabilities of 76% at 40.0 μM	38
<i>Aspergillus fumigatus</i>	<i>Cynodon dactylon</i>		14-Norpseurotin (124)	Inducing neurite outgrowth	Promoting neurite outgrowth of PC12 at 10.0 μM	28
			Pseurotin A (125)			
<i>Neosartorya fischeri</i> JS0553	<i>Glehnia littoralis</i>		Fischerin (126)	Neuroprotection against glutamate-mediated HT22 cell death	100% cell viability at 5 μM	22
			Pyripyropene A (127)		Inactive	
<i>Penicillium</i> sp. JVF17	<i>Vitex rotundifolia</i>		Penazaphilone E (128)	Neuroprotection against glutamate-mediated HT22 cell death	Neuroprotective activities with almost 100% at 25 μM	39
			Isochromophilone VI (129)			
			Peniazaphilone D (130)			
<i>Cochliobolus lunatus</i>			Sinulariapeptide A (131)	AChE inhibitory activity	IC ₅₀ , 1.8 ± 0.12 and 1.3 ± 0.11	40

SCSIO41401			Sinulariapeptide B (132)		μM, respectively		
<i>Rhizopycnis</i>	<i>vagum</i>	<i>Nicotiana tabacum</i>	Rhizovagine A (133)	AChE inhibitory activity	IC ₅₀ , 43.1 μM	41	
Nitaf22							
<i>Talaromyces</i>	sp.	<i>Tripterygium wilfordi</i>	Pseurotin A1 (134)	AChE inhibitory activity	Weaker	23	
LGT-2			Pseurotin A2 (135)				

“_” not test. IC₅₀, half maximal effective concentration; EC₅₀, half effective concentration; E_{max}, maximum effect; AChE, acetylcholinesterase; DPPH, 2,2-diphenyl-1-picrylhydrazyl; ABTS, 2,2-azino-bis(3-ethylbenzothiazoline-6-sulfonic acid) diammonium salt.