

Supporting Information for

Secondary Metabolites of the Genus *Didemnum*: A Comprehensive Review of Chemical Diversity and Pharmacological Properties

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NOTE: In all below tables, yellow highlight indicates that the method used bioassay-guided fractionation.

Table S1. Compounds with antitumor/anticancer and antiproliferative activities.

Compound Name	Chemical Class	Source organism/Location	Solvent of Extraction	Purification of the compounds	Structural Determination	Ref.
Lamellarin- ζ (69)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, and 1D NMR (^1H , ^{13}C) 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin- η (70)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV and IR, MS, 1D NMR (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin- ϕ (71)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV and IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin- χ (72)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV and IR spectra, MS, 1D NMR (^1H and ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin K (74)	Alkaloid	<i>Didemnum sp.</i> / Wasp Island, New South Wales	Aqueous EtOH	-LLPC and HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
		<i>Didemnum sp.</i> / Wasp Island, New South Wales	Aqueous EtOH	-LLPC, HPLC-UV-DAD	-UV, HRMS, 1D (^1H and ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
Lamellarin J (76)	Alkaloid	Dark purple encrusting colonies of <i>Didemnum sp.</i> / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel) and RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT and NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	³⁵
		<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
		<i>Didemnum sp.</i> / Australia	EtOH	-LLP, GPC-Sephadex LH-20, C18-Sep Pak, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2NMR (HETCOR, COLOC, COSY)	³⁶
		<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel) (GFC-Sephadex LH-20), C18 HPLC	UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
		<i>Didemnum sp.</i> / West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel) and RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT and NOE) and 2D NMR (XHCORRD, HMQC and COLOC)	³⁵

Lamellarin K triacetate (77)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin L triacetate (78)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin F (73)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin T diacetate (79)	Alkaloid	<i>Didemnum obscurum</i> / Tiruchandur coast in the Gulf of Mannar, Tamilnadu, India	MeOH	-TLC, CC (silica gel), GFC-Sephadex LH-20, C18 HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³³
Lamellarin S (87)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales	Aqueous EtOH	-LLPC, and HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
		<i>Didemnum</i> sp. / Australia	EtOH	LLPC, GPC-Sephadex LH-20, C18-Sep Pak, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2NMR (HETCOR, COLOC, COSY)	³⁶
Lamellarins A1-A3 (80-82)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales	Aqueous EtOH	-LLPC, and HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC and NOESY)	³⁴
Lamellarin E (59)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales	Aqueous EtOH	-LLPC, and HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
		<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
Lamellarin M (86)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales	Aqueous EtOH	-LLPC, and HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
		Dark purple colonies of <i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel) and RP HPLC	HREIMS, 1D (^1H , ^{13}C , INEPT, NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	³⁵
Lamellarin A6 (85)	Alkaloid	<i>Didemnum</i> sp. / Northern Rottnest Shelf, Western Australia	Aqueous EtOH	-LLPC, HPLC-UV-DAD	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
		<i>Didemnum</i> sp. / Northern Rottnest Shelf, Western Australia	EtOH	-LLP, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, ROESY, HMBC).	⁸²
Lamellarin L (61)	Alkaloid	Dark purple colonies of <i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel), RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT and NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	³⁵
		<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH20, RP HPLC, NP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR, (COSY, HSQC, HMBC, ROESY)	³²
		Purple encrusting <i>Didemnum</i> sp. / Indian Ocean	70% MeOH/CHCl ₃	-LLPC, GFC (Sephadex LH-20, Spectra Gel HW-40), RP HPLC	-IR, UV, LRMS and HRMS of DEI and FAB 1D (^1H , ^{13}C) and 2D NMR (DQFCOSY, HMQC, HMBC, NOESY)	³⁷

Lamellarin β (90)	Alkaloid	Purple encrusting <i>Didemnum</i> sp. / Indian Ocean	70% MeOH/CHCl ₃	-LLPC, GFC (Sephadex LH20, Spectra Gel HW-40), RP HPLC	-IR, UV, LRMS, HRMS of DEI, FABMS, 1D (¹ H, ¹³ C) and 2D NMR (DQFCOSY, HMQC, HMBC, NOESY)	³⁷
Lamellarin K-20-sulfate (91)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Lamellarin E-20-sulfate (92)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Lamellarin A3-20-sulfate (93)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Lamellarin B1-20-sulfate (94)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Lamellarin D-8-sulfate (95)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Lamellarin B2-20-sulfate (96)	Alkaloid	<i>Didemnum ternerratum</i> / Kingdom of Tonga	MeOH	-CC (RP, HP20 and HP20ss) and C18 HPLC	- UV, HRESIMS, 1D (1H and 13C) and 2D NMR (COSY, HMBC and ROESY).	³⁸
Granulatimide (101)	Alkaloid	<i>Didemnum granulatum</i> / Brazil	MeOH	-CC Sephadex LH20, RP HPLC	- UV, IR, HRFABMS, 1D (¹ H, ¹³ C, NOE) and 2D NMR (COSY, HMQC, HMBC)	^{40, 41}
Isogranulatimide (102)	Alkaloid	<i>Didemnum granulatum</i> / Brazil	MeOH	-CC Sephadex LH20, RP HPLC	-UV, IR, HRFABMS, 1D (¹ H, ¹³ C, NOE) and 2D NMR (COSY, HMQC, HMBC)	⁴⁰
6-Bromogranulatimide (104)	Alkaloid	<i>Didemnum granulatum</i> / Brazil	MeOH followed by DMF	-FLC (Silica gel), NP HPLC	-HRFABMS, UV, IR, 1D (¹ H, ¹³ C) and 2D NMR (HMQC, HMBC)	⁴¹
Didemnolines A-C (105-107)	Alkaloid	<i>Didemnum</i> sp. / Island of Rota, Northern Mariana Islands	MeOH	-FLC (Silica gel), NP HPLC	-IR, UV, HREIMS, 1D (¹ H, ¹³ C, NOE) and 2D NMR (HMBC)	⁴²
Bengacarboline (113)	Alkaloid	<i>Didemnum</i> sp. / Fiji	MeOH	-LLP, CC (Sephadex LH-20, C18 RP)	-IR, UV, FABHRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, TOCSY, HMQC, HMBC)	⁴⁴
Fascaplysin (114)	Alkaloid	<i>Didemnum</i> sp. / Micronesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (Silica gel), RP HPLC	-FABHRMS, 1D (¹ H and ¹³ C), 2D NMR (COSY, HMQC, HMBC, NOESY)	⁴⁵
		<i>Didemnum</i> sp. / Fiji	MeOH	-LLP, CC (Sephadex LH-20, C-18)	-IR, UV, FABHRMS, 1D (¹ H and ¹³ C) and 2D NMR (COSY, TOCSY, HMQC HMBC)	^{44, 78}
3-Bromofascaplysin (115)	Alkaloid	<i>Didemnum</i> sp. / Micronesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (Silica gel), RP HPLC	-FABHRMS, 1D (¹ H and ¹³ C), 2D NMR (COSY, HMQC, HMBC, NOESY)	⁴⁵
Mollamide (122)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Net Reef, in the central section of the Great Barrier Reef, Australia	MeOH/CH ₂ Cl ₂	-CC (silica gel), RP HPLC	-IR, HRESIMS, 1D (¹ H, ¹³ C, and DEPT) and 2D NMR (COSY, HMBC NOESY), X-ray crystallographic study	⁴⁶
Mollamide C (124)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Indonesia	H ₂ O	-VLC (Silica gel), NP and RP HPLC	-UV, IR, HRESIMS, 1D (¹ H and ¹³ C) 2 D NMR (COSY, HMQC, ROESY, HMBC)	⁴⁷

Keenamide A (125)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Indonesia	H ₂ O	-VLC (Silica gel), NP and RP HPLC	-UV, IR, HRESIMS, 1D (¹ H and ¹³ C) and 2 D NMR (COSY, HMQC, ROESY, HMBC)	⁴⁷
Shishijimicins A-C (126-128)	Polyketides / enediynes	<i>Didemnum proliferum</i> / Southern Japan	MeOH and EtOH	-LLPC, and RP HPLC	-HRFABMS, 1D (¹ H and ¹³ C) and 2D NMR (COSY, HOHAHA, HMBC, HMQC and NOESY)	⁴⁸
Namenamicin (129)	Polyketide / enediynes	<i>Didemnum proliferum</i> / Southern Japan	MeOH and EtOH	-LLPC, RP HPLC	-HRFABMS, 1D (¹ H and ¹³ C) and 2D NMR (COSY, HOHAHA, HMBC, HMQC, NOESY)	⁴⁸
Plakinidine D (130)	Alkaloid / pyrroloacridine	<i>Didemnum rubeum</i> / Island of Rota, Northern Mariana Islands	EtOH MeOH	-TLC, LLP, FLC (Silica gel)	-UV, IR, HREIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, INADEQUATE, HMBC)	^{49, 50}
Ascididemin (131)	Alkaloid / Pyridoacridine	<i>Didemnum</i> sp. / Okinawa, Japan	MeOH	-LLP, CC (silica gel)	-UV, IR, HRFABM, EIMS, 1D (¹ H, ¹³ C, NOE) and 2D NMR (COLOC)	⁵¹
		<i>Didemnum rubeum</i> / The Republic of Palau	MeOH	-CC (silica gel, Sephadex LH-20)	-UV, IR, HREIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC)	⁵⁰
		Red <i>Didemnum</i> sp. / Indonesia	MeOH	-CC (silica gel and LH-20)	-UV, IR, HREIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC)	⁵⁰
Methyl myristate (139)	Fatty acid methyl ester	<i>Didemnum psammatoches</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel and LH-20), RP HPLC (C18) and GC	-IR-FT, 1D (¹ H, ¹³ C) and 2D NMR	⁵³
Methyl palmitate (140)	Fatty acid methyl ester	<i>Didemnum psammatoches</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, LH-20), C18 HPLC, GC	-IR-FT, 1D NMR (¹ H, ¹³ C) and 2D NMR	⁵³
Methyl stearate (141)	Fatty acid methyl ester	<i>Didemnum psammatoches</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, LH-20), RP HPLC (C18) and GC	-IR-FT, 1D NMR (¹ H, ¹³ C) and 2D NMR	⁵³
Eusynstyelamide B (142)	Bis-indole alkaloid	<i>Didemnum candidum</i> / Great Barrier Reef, Australia	CH ₂ Cl ₂ /MeOH	-Semi-preparative C18 RP HPLC	-CD, MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	⁵⁴
Vitilevuamide (143)	Bicyclic depsipeptide	<i>Didemnum cuculliferum</i> / Namenalala Island, Fiji	MeOH	-FLC (silica gel), RP FLC, and RP HPLC	-UV, IR, HRMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (DQF COSY, COSY, TOCSY, HMQC, HMBC)	⁵⁵
Didemnenones C and D (144 and 145)	Fatty acid or polyketide / cyclopentenones	<i>Didemnum voeltzkowi</i> / Fringing reef, Suva Harbor, Fiji	MeOH	-LLP and CC (Sephadex LH-20)	-UV, IR, HRMS, 1D NMR (¹ H, ¹³ C)	⁵⁶
Didemnilactones A and B (146 and 147)	Fatty acid / 10-membered ring lactone	<i>Didemnum moseleyi</i> / Ago Bay, Mie Prefecture, Japan	MeOH	-CC (silica gel and alumina), RP HPLC	-UV, IR, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC)	⁵⁷
Neodidemnilactone (148)	Fatty acid	<i>Didemnum moseleyi</i> / Ago Bay, Mie Prefecture, Japan	MeOH	-CC (silica gel and alumina), RP HPLC	-UV, IR, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC)	⁵⁷
Cyclodidemnamide (149)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Philippines	30% CH ₂ Cl ₂ in MeOH	-LLP, CC (silica gel, LH-20), RP HPLC	-UV, IR, GC-MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, HMQC, ROZY)	⁵⁸

Comoramides A and B (152 and 153)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Mayotte	CHC1 ₃ /MeOH (1:2)	-VLC (silica gel)	-FT-IR, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, TCOSY, HMBC, NOESY)	⁵⁹
Mayotamides A and B (150 and 151)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Mayotte	CHC1 ₃ /MeOH (1:2)	-VLC (silica gel)	-FT-IR, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, TCOSY, HMBC, NOESY)	⁵⁹
Caledonin (154)	Peptide	<i>Didemnum rodriquesi</i> / Baie des Citrons, New Caledonia	MeOH	-LLP, CC (Silica gel, Sephadex LH-20), RP HPLC	-UV, IR, HRMS, FABMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (DQF-COSY, TOCSY, HMQC, ROZY	⁶⁰
Hexamollamide (161)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Okinawa, Japan	MeOH	-LLP, RP CC (ODS), RF HPLC and Chiral HPLC	-ESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	⁶²

Table S2. Compounds with antimicrobial activities.

Compound Name	Chemical Class	Source organism / Location	Solvent of Extraction	Purification of the Compounds	Structural Determination	Ref.
Rodriguesines A and B (162 and 163)	Peptide / diketopiperazine	<i>Didemnum</i> sp. / Salvador, Brazil	MeOH	-CC (C18, Sephadex LH-20), RP HPLC	-UV, IR, HRESIMS, 1D NMR (¹ H, ¹³ C) and 2D NMR (COSY, HMQC, HMBC)	⁶³
Didemnaketals A and B (179 and 180)	Polyketide / spiroketal	<i>Didemnum</i> sp. / Auluptagel Island, Palau	CH ₂ Cl ₂ /MeOH (1:1)	-Sephadex LH-20, RP HPLC	-IR, HRFABMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC, HMBC)	⁶⁹
Didemnaketals D and E (166 and 167)	Polyketide / spiroketal	<i>Didemnum</i> sp. / Sharm El-Sheikh, Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-LLP, VLC (Silica gel), Sephadex LH20, RP HPLC	-UV, IR, HRESIMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (COSY, ROESY, NOESY, TOCSY, HSQC, HMBC)	⁶⁴
Didemnaketals F and G (173 and 174)	Polyketide / spiroketal	<i>Didemnum</i> sp. / Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-VLC, Sephadex LH-20, C18 HPLC	-IR, UV, HRESIMS, 1D (¹ H, ¹³ C, DEPT) and 2D (COSY, ROESY, NOESY, TOCSY, HSQC, HMBC)	⁶⁶
Mollamides E and F (186-187)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / New Britain, Papua New Guinea	MeOH	-CC (HP20SS resin), FLC (C18), RP HPLC	-UV, IR, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (gCOSY, HMBC, ROESY)	⁷²
Molleurea A (5)	Alkaloid / phenylethylamine	<i>Didemnum molle</i> / New Britain, Papua New Guinea	MeOH	-CC (HP20SS resin), FLC (C18) and RP HPLC	-IR, UV, HRESIMS, 1D NMR (¹ H, ¹³ C) 2DNMR (gCOSY, HMBC, ROESY)	⁷²
		<i>Didemnum molle</i> / Mayotte	MeOH/CH ₂ Cl ₂ (1:1)	-Kupchan solvent partition procedure, CC (Sephadex LH-20), RP HPLC	-UV, IR HRMS, LCMS, 1D (¹ H, ¹³ C, DEPT) and 2D (COSY, TOCSY, HSQC, HMQC, HMBC, ROESY)	¹⁷
3,5-Diiodo-4-methoxyphenethylamine (27)	Alkaloid / phenylethylamine	<i>Didemnum</i> sp. / Northwest end of Cocos Lagoon, Guam, United States	Organic solvents	-Crystallization from a CHC ₁₃ extract	-HRMS, 1D NMR (¹ H, ¹³ C)	⁵²
		<i>Didemnum rubeum</i> / Republic of Palau	EtOH MeOH	-LLP, and FLC (Silica gel)	-UV, IR, HREIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, INADEQUATE, HMBC)	^{49, 50}
Kakelokelose homopolysaccharide (188)	Carbohydrate / sulfated mannose	<i>Didemnum molle</i> / Pohnpei, Micronesia	95% EtOH	-Ultrafiltration membranes are expressed as Molecular Weight Cutoff (MWCO)	-FT-IR, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (COSY, HMQC)	⁷³
Cyclodidemniserinol trisulfate (182)	Fatty acid / serinolipid	<i>Didemnum guttatum</i> / Ngerchaol Island, Palau	MeOH	-Purified by repeated reversed phase chromatography	-UV, IR, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC-TOCSY, HMBC)	⁷¹
(R)-(E)-1-Aminotridec-5-en-2-ol (191)	Alkaloid	<i>Didemnum</i> sp. / Great Barrier Reef, Australia	MeOH	-Kupchan solvent partition procedure, CC (Sephadex LH-20), FLC (C18) RP HPLC	-UV, IR, MS, and 1D NMR (¹ H, ¹³ C, DEPT) 2D NMR (COSY, HMQC)	⁷⁵
Divamides A and B (189 and 190)	Cyclic peptide / lanthipeptide	<i>Didemnum molle</i> / Eastern Fields, Papua New Guinea	MeOH / metagenomics method	-HP20, HPLC (C18)	-UV, MS, 1D NMR (1H, 13C) 2D NMR (COSY, TOCSY, HMQC, HMBC, ROESY, NOESY), metagenome sequencing	⁷⁴

Table S3. Compounds with antiprotozoal activities.

Compound Name	Chemical Class	Source organism / Location	Solvent of Extraction	Purification of the compounds	Structural Determination	Ref.
Lepadins D-F (194-196)	Alkaloid / decahydroquinoline	<i>Didemnum</i> sp. / Stanley Reef, the Great Barrier Reef, Australia	Acetone	-TLC, NP VLC, NP HPLC	-HREIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, ROESY, NOE)	⁷⁶
Albopunctatone (197)	Polyketide / anthrone-anthraquinone	<i>Didemnum albopunctatum</i> / Swain Reefs in the Southern Great Barrier Reef, Australia	MeOH	-NP HPLC, RP HPLC	-UV, IR, MS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	⁷⁷
Homofascaplysin A (199)	Alkaloid	<i>Didemnum</i> sp. / Fiji	MeOH	-LLP, CC (Sephadex LH-20, C-18 RP)	-UV, IR, FABHRMS, ECD, 1D (^1H and ^{13}C) and 2D NMR (COSY, TOCSY, HMQC, HMBC)	⁷⁸
Mollamide B (123)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Indonesia	H ₂ O	-VLC (Silica gel), NP and RP HPLC	-UV, IR, HRESIMS, 1D (^1H and ^{13}C) 2 D NMR (COSY, HMQC, HMBC, ROESY)	⁴⁷
Didemnidine B (202)	Alkaloid / indole spermidine	<i>Didemnum</i> sp. / New Zealand	MeOH	-FLC (C8 and C18 RP), Sephadex LH-20, RP HPLC	-UV, IR, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	⁷⁹

Table S4. Compounds with antidiabetic activities.

Compound Name	Chemical Class	Source organism / Location	Solvent of Extraction	Purification of the compounds	Structural Determination	Ref.
<i>N,N'</i> -Diphenethyl-urea (11)	Alkaloid / phenylethlyamine	<i>Didemnum mole</i> / Okinawa, Japan	MeOH	-LLP, fractionation using ODS RP	-UV, IR, MS, 1D (^1H , ^{13}C)	⁸⁰
		<i>Didemnum ternatanum</i> / Urukthapal, Palau	MeOH	-CC (Silica gel, Sephadex LH-20)	-IR, HRMS, EIMS, 1D NMR (^1H , ^{13}C)	¹⁶
		<i>Didemnum molle</i> / Mayotte	MeOH/CH ₂ Cl ₂ (1:1)	-Kupchan solvent partition procedure, CC (Sephadex LH-20), RP HPLC	-UV, IR, HRMS, LCMS, 1D (^1H , ^{13}C , DEPT) and 2D NMR (COSY, TOCSY, HSQC, HMQC, HMBC, ROESY)	¹⁷

Table S5. Compounds that affect the central nervous system.

Compound Name	Chemical Class	Source organism / Location	Solvent of Extraction	Purification of the compounds	Structural Determination	Ref.
Ningalin B-D (204-2067)	Alkaloid	<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	EtOH	-LLP, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, ROESY, HMBC)	⁸²
		<i>Didemnum sp.</i> / near Ningaloo Reef, Western Australia	70% MeOH in CH_2Cl_2	-LLP, GFC (Sephadex LH-20), C18 HPLC, GPC (Spectra Gel HW-40)	-UV, IR, HRFABMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	⁸¹
Ningalin E-G (207-209)	Alkaloid	<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	EtOH	-LLP, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, ROESY, HMBC)	⁸²
Lamellarin G (60)	Alkaloid	<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	EtOH	-LLP, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, ROESY, HMBC)	⁸²
		<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	Aqueous EtOH	-LLPC, HPLC-UV-DAD	-HRMS, UV, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMBC, ROESY).	³⁴
		<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-HRESMS, UV, IR, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
		Purple encrusting <i>Didemnum</i> sp. / the Indian Ocean	70% MeOH/CHCl ₃	-LLPC, GFC (Sephadex LH-20 and Spectra Gel HW-40), RP HPLC	-IR, UV, LRMS, HRMS of DEI, FABMS, 1D (^1H , ^{13}C) and 2D NMR. (DQFCOSY, HMQC, HMBC, NOESY)	³⁷
Lamellarin Z (62)	Alkaloid	<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	EtOH	-LLP, RP HPLC	-UV, HRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, ROESY, HMBC)	⁸²
		<i>Didemnum sp.</i> / Northern Rottnest Shelf, Western Australia	Aqueous EtOH	-LLPC, and HPLC-UV-DAD	-HRMS, UV, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMBC, ROESY)	³⁴
		<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C-18 SPE cartridge	-HRESMS, UV, IR, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
Lepadin I (210)	Alkaloid	<i>Didemnum sp.</i> / Bahamas	70% MeOH in CH_2Cl_2	-LLP, CC (Silica), RP HPLC, C-18 SPE cartridge	-HRMS, ESIMS IR, UV, ECCD, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMBC, HSQC, NOESY)	⁸³

Table S6. Compounds without any reported biological activities.

Compound Name	Chemical Class	Source organism / Location	Solvent of Extraction	Purification of the compounds	Structural Determination	Ref.
Lepadin J and K (211 and 212)	Alkaloid	<i>Didemnum</i> sp. / Bahamas	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (Silica), RP HPLC, C18 SPE cartridges	-UV, ESIMS IR, HRMS, ECCD, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	⁸³
1,8-Dihydroxy-9,10 anthraquinone (199)	Polyketide / anthraquinone	<i>Didemnum albopunctatum</i> / Australia	MeOH	-NP HPLC, RP HPLC	-UV, IR, MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	⁷⁷
Salvadenosine (43)	Nucleoside	<i>Didemnum</i> sp. / Bahamas	MeOH	-LLP, RP HPLC	-HRMS and 1D (¹ H, ¹³ C), 2D NMR (COSY, HSQC, HMBC)	²⁶
Lamellarin A4 and A5 (83 and 84)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales, Australia	MeOH	-LLPC, HPLC-UV-DAD	-UV, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (HMQC, HMBC, NOESY)	³⁴
Lamellarin C (58)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales, Australia	Aqueous EtOH	-LLPC, HPLC-UV DAD	-UV, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, ROESY)	³⁴
		<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP and RP HPLC, C18-SPE cartridge	-HRESMS, UV, IR, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
		<i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast, Australia	MeOH/CH ₂ Cl ₂	-VLC (silica gel), RP HPLC	-HREIMS, 1D (¹ H, ¹³ C, INEPT and NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	³⁵
Lamellarin T (88)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales, Australia	Aqueous EtOH	-LLPC, HPLC-UV DAD	-UV, HRMS, 1D NMR (¹ H, ¹³ C) and 2D NMR, (COSY, HMBC, ROESY)	³⁴
Lamellarin X (89)	Alkaloid	<i>Didemnum</i> sp. / Wasp Island, New South Wales, Australia	Aqueous EtOH	-LLPC, HPLC-UV-DAD	-HRMS, UV, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, ROESY)	³⁴
Lamellarin B 20-sulfate (63)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D NMR (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
Lamellarin C 20-sulfate (64)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²
Lamellarin L 20-sulfate (65)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	³²

Lamellarin A (56)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	32
		<i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel), RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT, NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	35
Lamellarin B (57)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP and RP HPLC, C-18 SPE cartridge	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	32
		Dark purple encrusting colonies of <i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel), RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT, NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	35
Lamellarin D triacetate (67)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC, C18 SPE cartridge	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY).	32
		Dark purple encrusting colonies of <i>Didemnum</i> sp. / South West Cay, Lihou Reef, off the North Queensland coast	MeOH/CH ₂ Cl ₂	-VLC (silica gel), RP HPLC	-HREIMS, 1D (^1H , ^{13}C , INEPT, NOE) and 2D NMR (XHCORRD, HMQC, COLOC)	35
Lamellarin N triacetate (68)	Alkaloid	<i>Didemnum chartaceum</i> / Australia	MeOH	-LLPC, Sephadex LH-20, NP HPLC, RP HPLC and SPE cartridge packed with C18	-UV, IR, HRESMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC, ROESY)	32
Ningalin A (203)	Alkaloid	<i>Didemnum</i> sp. / near Ningaloo Reef, Western Australia	70% MeOH in CH ₂ Cl ₂	-LLP, GFC (Sephadex LH-20), C18 HPLC, GPC (Spectra Gel HW-40)	-UV, IR, HRFABMS 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	81
Didemnimate A (97)	Alkaloid	<i>Didemnum conchyliatum</i> / Sweetings Cay, Bahamas	MeOH/CH ₂ Cl ₂ (1:1)	-VLC, NP CC, C18 RP CC, RP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC), X-ray analysis	39
		<i>Didemnum granulatum</i> / Brazil	MeOH	-CC, Sephadex LH20, RP HPLC	-UV, IR, HRFABMS, 1D (^1H , ^{13}C , NOE) and 2D NMR (COSY, HMQC, HMBC)	40
Didemnimate B (98)	Alkaloid	<i>Didemnum conchyliatum</i> / Sweetings Cay, Bahamas	MeOH/CH ₂ Cl ₂ (1:1)	-VLC, NP CC, C18 RP, CC, RP HPLC	-HREIMS, UV, IR, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC), X-ray analysis	39
Didemnimate C (99)	Alkaloid	<i>Didemnum conchyliatum</i> / Sweetings Cay, Bahamas	MeOH/CH ₂ Cl ₂ (1:1)	-VLC, NP CC, C18 RP, RP HPLC	-HREIMS, UV, IR, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC), X-ray analysis	39
Didemnimate D (100)	Alkaloid	<i>Didemnum conchyliatum</i> / Sweetings Cay, Bahamas	MeOH/CH ₂ Cl ₂ (1:1)	-VLC, NP CC, C18 RP, RP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C) and 2D NMR (HMQC, HMBC), X-ray analysis	39

		<i>Didemnum conchyliatum</i> / Sweetings Cay, Bahamas	MeOH	-CC Sephadex LH20, RP HPLC	-UV, IR, HRFABMS, 1D (^1H , ^{13}C , NOE) and 2D NMR (COSY, HMQC, HMBC)	40
Didemnimide E (103)	Alkaloid	<i>Didemnum granulatum</i> / Brazil	MeOH	-CC Sephadex LH20, RP HPLC	-UV, IR, HRFABMS, 1D (^1H , ^{13}C NOE) and 2D NMR (COSY, HMQC, HMBC)	40
N-Acetyl-rodriguesines A and B (164 and 165)	Peptide / diketopiperazine	<i>Didemnum</i> sp. / Salvador, Brazil	MeOH	-LLP, CC (Sephadex LH-20, C18), RP HPLC	-IR, UV, HRESIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC)	63
Didemmiserinolipids A-C (183 and 185)	Fatty acid / serinolipid	<i>Didemnum</i> sp. / Sulawesi Island, Indonesia	MeOH	-LLPC, VLC (silica gel), RP HPLC	-UV, IR, HRESIMS, 1D (^1H , ^{13}C and DEPT) and 2D NMR (1H-1H COSY, TOCSY, HMQC and ROESY)	70
Didemnoline D (108)	Alkaloid	<i>Didemnum</i> sp. / Near Rota, Northern Mariana Islands	MeOH	-FLC (Silica gel), NP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C and NOE) and 2D NMR (HMBC)	42
Eudistomin O (109)	Alkaloid	<i>Didemnum</i> sp. / Near Rota, Northern Mariana Islands	MeOH	-FLC (Silica gel), NP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C , NOE) and 2D NMR (HMBC)	42
β-Carboline (110)	Alkaloid	<i>Didemnum</i> sp. / Near Rota, Northern Mariana Islands	MeOH	-FLC (Silica gel), NP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C , NOE) and 2D NMR (HMBC)	42
2-(2',4'-Dibromophenoxy)-3,5-dibromophenol (111)	Phenolic ether	<i>Didemnum</i> sp. / Near Rota, Northern Mariana Islands	MeOH	-FLC (Silica gel), NP HPLC	-UV, IR, HREIMS, 1D (^1H , ^{13}C , NOE) and 2D NMR (HMBC)	42
β-Carboline dimers (112)	Alkaloid	<i>Didemnum</i> sp. / Sykes Reef in the Capricorn Bunker Group of the southern Great Barrier Reef, Australia	Hexane followed by EtOAc	-CC (Silica gel)	-UV, IR, MS, 1D NMR (^1H , ^{13}C)	43
3-Bromohomo-fascaplysin A (200)	Alkaloid	<i>Didemnum</i> sp. / Fiji	MeOH	-LLP, CC (Sephadex LH-20, C18 RP)	-UV, IR, FABHRMS, ECD, 1D (^1H , ^{13}C) and 2D NMR (COSY, TOCSY, HMQC, HMBC)	78
Homofascaplysin B-1 (116)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C), 2D NMR (COSY, HMQC, HMBC, NOESY)	45
3-Bromohomo-fascaplysin B (117)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	45
Homofascaplysin C (118)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	45
3-Bromohomo-fascaplysin B-1 (119)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	45
3-Bromohomo-fascaplysin C (120)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC and NOESY)	45
Homofascaplylate A (121)	Alkaloid	<i>Didemnum</i> sp. / Indonesia	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-FABHRMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMQC, HMBC, NOESY)	45

Didemnaketal C (181)	Polyketide / spiroketals	<i>Didemnum</i> sp. / Auluptagel Island, Palau	MeOH/CH ₂ Cl ₂ (1:1)	-Sephadex LH-20, RP HPLC	-UV, IR, HRFABMS, 1D NMR (¹ H, ¹³ C)	68
Mollecarbamates A-D (1-4)	Phenylethylamine	<i>Didemnum molle</i> / Mayotte	MeOH/CH ₂ Cl ₂ (1:1)	-Kupchan solvent partition procedure, CC (Sephadex LH-20), RP HPLC	-UV, IR, HRMS, LCMS, 1D (¹ H, ¹³ C, DEPT) 2D (COSY, TOCSY, NOESY, HSQC, HMBC)	17
Molleurea B-E (5-8)	Phenylethylamine	<i>Didemnum molle</i> / Mayotte	MeOH/CH ₂ Cl ₂ (1:1)	-Kupchan solvent partition procedure, CC (Sephadex LH-20), RP HPLC	-UV, IR, HRMS, LCMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (COSY, NOESY, HMBC)	17
Molledihydroisoquinolone (10)	Phenylethylamine	<i>Didemnum molle</i> / Baie des Assassins, Madagascar	MeOH/CH ₂ Cl ₂ (1:1)	-Kupchan solvent partition procedure, CC (Sephadex LH-20), RP HPLC	-UV, IR, HRMS, LCMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (COSY, TOCSY, HSQC, HMQC, HMBC, ROESY)	17
Molleurea E (9)	Phenylethylamine	<i>Didemnum molle</i> / New Britain, Papua New Guinea	MeOH	-CC (HP20SS resin), C18 FLC, RP HPLC	-UV, IR, HRESIMS, 1D NMR (¹ H, ¹³ C) and 2D NMR (gCOSY, HMBC and ROESY)	72
Antatollamides A and B (14 and 15)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Pohnpei	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, FLC (Silica gel), RP HPLC	-UV, FTIR, HRMS, 1D (¹ H, ¹³ C, NOE) and 2D NMR (COSY, TOCSY, HSQC, HMBC, NOESY)	19
Didemnidine A (201)	Alkaloid	<i>Didemnum</i> sp. / New Zealand	MeOH	-FLC (C8, C18), Sephadex LH-20, RP HPLC	-UV, IR, HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	79
Didemnaceride A and B (168 and 169)	Fatty acid / glyceride	<i>Didemnum</i> sp. / Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-LLP, VLC, CC (Silica gel), CC (RP-C18)	-UV, IR, HR-ESI-MS, GC-MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	65
24-Ethyl-25-hydroxycholesterol (170)	Steroid	<i>Didemnum</i> sp. / Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-LLP, VLC, CC (Silica gel), CC (RP-C18)	-UV, IR, HR-ESI-MS, GC-MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	65
Cholest-6-en-3,5,8-triol (171)	Steroid	<i>Didemnum</i> sp. / Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-LLP, VLC, CC (Silica gel), CC (RP-C18)	-UV, IR, HR-ESI-MS, GC-MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	65
Cholestane-3β,5α,6β-26-tetrol (172)	Steroid	<i>Didemnum</i> sp. / Red Sea, Egypt	CH ₂ Cl ₂ /MeOH (1:1)	-LLP, VLC, CC (Silica gel), CC (RP-C18)	-UV, IR, HR-ESI-MS, GC-MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	65
Minalemines A-F (155-160)	Peptide / guanidine	<i>Didemnum rodriguesi</i> / Baie des Citrons, Noumea, New Caledonia	MeOH	-LLP, CC (Sephadex LH-20), RP HPLC	-UV, IR, HRFABMS, 1D (¹ H, ¹³ C, DEPT) and 2D NMR (COSY, TOCSY, HMQC, HMBC)	61
Cholesterol (132)	Steroid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (¹ H and ¹³ C) and 2D NMR	53
Campesterol (133)	Steroid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (¹ H and ¹³ C) and 2D NMR	53

Stigmasterol (18)	Steroid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H and ^{13}C) and 2D NMR	53
β-Sitosterol (134)	Steroid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H and ^{13}C) 2D NMR	53
Palmitic acid (135)	Fatty acid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H and ^{13}C) and 2D NMR	53
Stearic acid (136)	Fatty acid	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H , ^{13}C) and 2D NMR	53
1,2-Propanediol, 3-(heptadecyloxy) (137)	Fatty acid / glyceryl ether	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H , ^{13}C) and 2D NMR	53
Batyl alcohol (19)	Fatty acid / glyceryl ether	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H , ^{13}C) and 2D NMR	53
1,2-Propanediol,3[(met hyloctadecyl)oxy] (138)	Fatty acid / glyceryl ether	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H and ^{13}C) and 2D NMR	53
Thymidine (21)	Nucleoside	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H and ^{13}C) and 2D NMR	53
		<i>Didemnum psammatodes</i> / Shores of Ceará State, Brazil	MeOH	-CC (silica gel, Sephadex LH-20)	-UV, MS, GC-MS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	20

2'-Deoxyguanosine (23)	Nucleoside	<i>Didemnum psammatodes</i> / Intertidal zone of the beach rocks at Flexeiras Beach, Brazil	MeOH	-CC (silica gel, Sephadex LH-20), C18 HPLC, GC	-IR-FT, 1D (^1H , ^{13}C) and 2D NMR	53
		<i>Didemnum psammatodes</i> / Shores of Cear á State.	MeOH	-CC (silica gel and Sephadex LH-20)	-UV, MS, GC-MS, 1D NMR (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	20
2'-Deoxyuridine (20)	Nucleoside	<i>Didemnum psammatodes</i> / Shores of Cear á State, Brazil	MeOH	-CC (silica gel, Sephadex LH-20)	-UV, MS, GC-MS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	20
2'-Deoxyinosine (22)	Nucleoside	<i>Didemnum psammatodes</i> / Shores of Cear á State, Brazil	MeOH	-CC (silica gel and Sephadex LH-20)	-UV, MS, GC-MS, 1D (^1H , ^{13}C) and 2D (COSY, HSQC, HMBC)	20
Cholestanol (16)	Steroid	<i>Didemnum psammatodes</i> / Shores of Cear á State, Brazil	MeOH	-CC (silica gel, Sephadex LH-20)	-UV, MS, GC-MS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	20
Cholestanone (17)	Steroid	<i>Didemnum psammatodes</i> / Shores of Cear á State, Brazil	MeOH	-CC (silica gel, Sephadex LH-20)	-UV, MS, GC-MS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	20
16-Epi-18-acetyl herdmanine D (24)	Alkaloid	<i>Didemnum</i> sp. / Near Haegeumgang, Geoje in the South Sea of Korea	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (Sephadex LH-20), RP FLC, RP HPLC	-HRESIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HMBC)	21
N-(6-Bromo-1H- indole-3-carbonyl)- L-arginine (25)	Alkaloid	<i>Didemnum</i> sp. / Near Haegeumgang, Geoje, Korea	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (LH-20), RP FLC, RP HPLC	-HRESIMS, 1D (^1H , ^{13}C) and 2D NMR, (COSY, HMBC)	21
(6-Bromo-1H- indol-3- yl)oxoacetamide (26)	Alkaloid	<i>Didemnum</i> sp. / Near Haegeumgang, Geoje, Korea	MeOH/CH ₂ Cl ₂ (1:1)	-LLP, CC (LH-20), RP FLC, RP HPLC	-HRESIMS, 1D (^1H , ^{13}C) and 2D NMR, (COSY, HMBC)	21
2-(3,5-Diiodo-4- methoxyphenyl)eth anamine (27)	Phenylethylamin e	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC).	22
2-(3,5-Diiodo-4- methoxyphenyl)eth anaminium (28)	Phenylethylamin e	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC).	22
2-(3,5-Diiodo-4- methoxyphenyl)eth anaminium benzoate (29)	Phenylethylamin e	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (^1H , ^{13}C) and 2D NMR (COSY, HSQC, HMBC)	22

2-(3,5-Diiodo-4-methoxyphenyl)acetamide (30)	Phenylethylamine	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	22
N-[2-(3,5-Diiodo-4-methoxyphenyl)ethyl]formamide (31)	Phenylethylamine	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	22
N-[2-(3,5-Diiodo-4-methoxyphenyl)ethyl]benzamide (32)	Phenylethylamine	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	22
N,N'-Bis[2-(3,5-diiodo-4-methoxyphenyl)ethyl] ethanediamide (33)	Phenylethylamine	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	22
4-(2-Aminoethyl)-2-iodophenol (34)	Phenylethylamine	<i>Didemnum rubeum</i> / Chuuk Atoll	Aqueous crude extract	-CC (Sephadex LH20), RP HPLC	-LRMS, HRESIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HSQC, HMBC)	22
Minimide (35)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Solomon Islands	MeOH / metagenomics	-LCMS	-LCMS, metagenomics, synthetic biology	23
5α,8α-Epidioxycholest-6-en-3β-ol (36)	Steroid	<i>Didemnum salary</i> / Andavadoaka, Madagaskar	CHCl ₃ /MeOH (1:1)	-CC (Silica gel, Sephadex LH20)	-1D NMR (¹ H NMR)	24
5α,8α-Epidioxy-24(S)-methylcholest-6-en-3β-ol (37)	Steroid	<i>Didemnum salary</i> / Andavadoaka, Madagaskar	CHCl ₃ /MeOH (1:1)	-CC (Silica gel, Sephadex LH20)	-1D NMR (¹ H NMR)	24
5α,8α-Epidioxy-24(R)-methylcholest-6-en-3β-ol (38)	Steroid	<i>Didemnum salary</i> / Andavadoaka, Madagaskar	CHCl ₃ /MeOH (1:1)	-CC (Silica gel, Sephadex LH20)	-1D NMR (¹ H NMR)	24
5α,8α-Epidioxy-24(S)-ethylcholest-6-en-3β-ol (39)	Steroids	<i>Didemnum salary</i> / Andavadoaka, Madagaskar	CHCl ₃ /MeOH (1:1)	-CC (Silica gel, Sephadex LH20)	-1D NMR (¹ H NMR)	24
5α,8α-Epidioxy-24(R)-ethylcholest-6-en-3β-ol (40)	Steroids	<i>Didemnum salary</i> / Andavadoaka, Madagaskar	CHCl ₃ /MeOH (1:1)	-CC (Silica gel, Sephadex LH20)	-1D NMR (¹ H NMR)	24
3,5-Dibromotetra-methyltyrosine (41)	Phenylethylamine	<i>Didemnum</i> sp. / White Sands Reef, Algoa Bay, South Africa	MeOH/CH ₂ Cl ₂	-LC-ICP-MS/ESI-MS	-1D NMR (¹ H NMR)	25
3-Iodotetramethyl-tyrosine (42)	Phenylethylamine	<i>Didemnum</i> sp. / White Sands Reef, Algoa Bay, South Africa	MeOH/CH ₂ Cl ₂	-LC-ICP-MS/ESI-MS	-1D NMR (¹ H NMR)	25

6-Bromotryptamine (44)	Alkaloid / brominated indole derivative	<i>Didemnum candidum</i> / Southern Gulf of California, Mexico	MeOH	-CC (Sephadex LH-20), FLC (silica gel)	-UV, IR, MS, 1D (¹ H, ¹³ C) and 2D NMR (COSY)	27
2,2-Bis(6'-bromo-3'-indolyl)ethylamine (45)	Alkaloid / brominated indole derivative	<i>Didemnum candidum</i> / Southern Gulf of California, Mexico	MeOH	-CC (Sephadex LH-20), FLC (silica gel)	-UV, IR, MS, 1D (¹ H, ¹³ C) 2D NMR (COSY)	27
2,5-Bis(6'-bromo-3'-indolyl)piperazine (46)	Alkaloid / brominated indole derivative	<i>Didemnum candidum</i> / Southern Gulf of California, Mexico	MeOH	-CC (Sephadex LH-20) and FLC (silica gel)	-UV, IR, MS, 1D (¹ H, ¹³ C) 2D NMR (COSY)	27
Didmolamides A and B (12 and 13)	Cyclic peptide / cyanobactin	<i>Didemnum molle</i> / Madagascar	CHCl ₃ /MeOH (2:1)	-LLP, CC (Sephadex LH-20), HPLC	-FTIR, HR-EIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC, HMBC)	18
Ascidatrienolides A-C (47-49)	Eicosanoids	<i>Didemnum candidum</i> / Philippines	Acetone	-VLC (silica gel), NP HPLC	-UV, IR, HRBIMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY)	28
5'-Deoxy-3-iodotubercidin (50)	Nucleoside	<i>Didemnum voeltzkowi</i> / Philippines	MeOH	-LLP, C18 VLC, RP HPLC	-FABMS, 1D NMR (¹ H, ¹³ C NMR)	29
5'-Deoxy-3-bromotubercidin (51)	Nucleoside	<i>Didemnum voeltzkowi</i> / Philippines	MeOH	-LLP, C18 VLC, RP HPLC	-FABMS and 1D NMR (¹ H, ¹³ C NMR)	29
5'-Deoxytubercidin (52)	Nucleoside	<i>Didemnum voeltzkowi</i> / Philippines	MeOH	-LLP, C18 VLC, RP HPLC	-FABMS and 1D NMR (¹ H, ¹³ C NMR)	29
Hydroxy phenyldienoic acid (53)	Polyketide / phenylpolyene	<i>Didemnum granulatum</i> / Red Sea	EtOAc	-CC (Sephadex LH-20)	-UV, MS, 1D (¹ H, ¹³ C) and 2D NMR(COSY)	30
Enterocin (175)	Polyketide / type II PKS	<i>Didemnum</i> sp. / Australia	70% MeOH in CH ₂ Cl ₂	-LLPC and VLC, RP HPLC	-HRMS, IR, UV, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, NOSY)	67
5-Deoxyenterocin (176)	Polyketide / type II PKS	<i>Didemnum</i> sp. / Australia	70% MeOH in CH ₂ Cl ₂	-LLPC, VLC, RP HPLC	-HRMS, IR, UV, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, NOSY)	67
Enterocin-5-behenate (177)	Polyketide / type II PKS	<i>Didemnum</i> sp. / Australia	70% MeOH in CH ₂ Cl ₂	-LLPC, VLC, RP HPLC	-HRMS, IR, UV, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMBC, NOSY)	67
Asterubin (54)	Guanidine derivative	<i>Didemnum ligulum</i> / São Sebastião, Brazil	MeOH	-CC (Sephadex LH-20), RP HPLC	-HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC, HMBC)	31
N,N-Dimethyl-O-methylethanolamine (55)	Ethylamine derivative	<i>Didemnum ligulum</i> / São Sebastião, Brazil	MeOH	-CC (Sephadex LH-20), RP HPLC	-HRMS, 1D (¹ H, ¹³ C) and 2D NMR (COSY, HMQC, HMBC)	31