

Supplementary Materials

Marine Natural Products from Indonesian Waters

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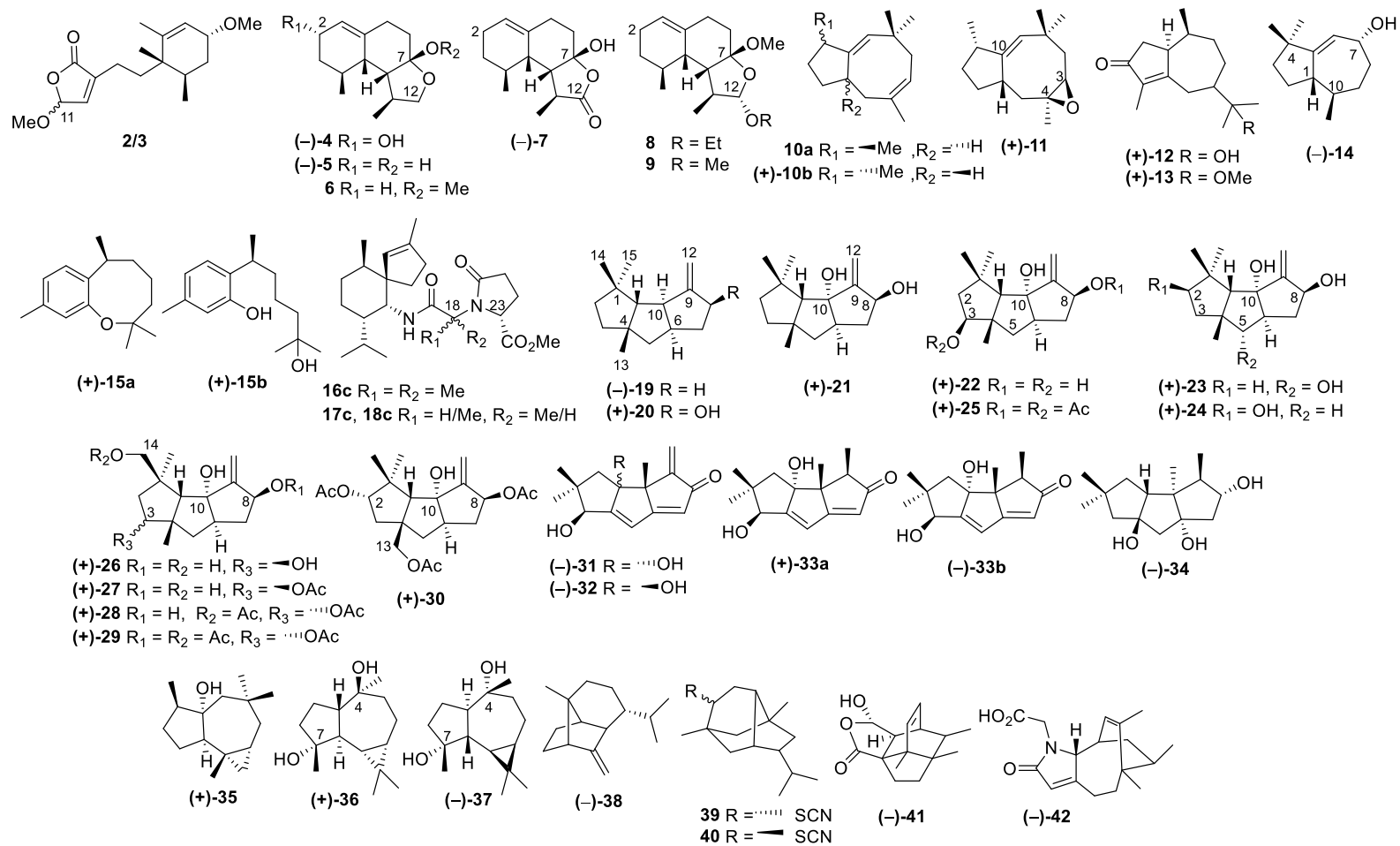


Figure S1: Structures of marine sesquiterpenoids from Indonesian waters found in 1970–2017.

Table S1: Marine sesquiterpenoids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
<i>O,O</i> -Dimethyl lingshuiolide A 2/11- <i>epi</i> 3 ^{α,β} [C ₁₇ H ₂₆ O ₄]	UV, IR, MS, NMR ECD, [α] _D	Lingshuiane*	Cytotoxic, Anti-microbial, Antiathero-sclerotic, Antiosteoclastic, Anticancer	Several cell lines, microorganisms, enzymes	NA	<i>L. herbacea</i>	NSW	[95]
(-)-Lemnacarnol 4 ^β [C ₁₅ H ₂₄ O ₃]	IR, MS, NMR, [α] _D , CT, X-ray	Nardosinane	Toxic	<i>P. micans</i> , <i>A. carterae</i>	Quan. Undetm.	<i>L. carnosa</i>	MLU	[96–99]
(-)-2-Desoxylemnacarnol 5 ^β [C ₁₅ H ₂₄ O ₂]	IR, MS, NMR, [α] _D , CT	Nardosinane	Undetm.	Undetm.	Undetm.	<i>L. africana</i> , <i>L. laevis</i> , <i>P. thyrsoidea</i>	MLU	[100]
2-Deoxy-7- <i>O</i> -methyllemnacarnol 6 ^β [C ₁₆ H ₂₆ O ₂]	UV, IR, MS, NMR	Nardosinane	Undetm.	Undetm.	Undetm.	<i>Nephtea</i> sp.	NSW	[101]
(-)-2-Desoxy-12-oxolemnacarnol 7 ^β [C ₁₅ H ₂₂ O ₃]	IR, MS, NMR, [α] _D , CT, X-ray	Nardosinane	Undetm.	Undetm.	Undetm.	<i>L. africana</i> , <i>L. laevis</i> , <i>P. thyrsoidea</i>	MLU	[100]
2-Deoxy-12α-ethoxy-7- <i>O</i> -methyllemnacarnol 8 ^β [C ₁₈ H ₃₀ O ₃]	UV, IR, MS, NMR	Nardosinane	Undetm.	Undetm.	Undetm.	<i>Nephtea</i> sp.	NSW	[101]
2-Deoxy-12α-methoxy-7- <i>O</i> -methyllemnacarnol 9 ^β [C ₁₇ H ₂₈ O ₃]	UV, IR, MS, NMR	Nardosinane	Undetm.	Undetm.	Undetm.	<i>Nephtea</i> sp.	NSW	[101]
Precapnelladiene 10a ^β [C ₁₅ H ₂₄]	IR, GCMS, NMR, Mol. Mod. (CONGEN)	Precapnellane	Undetm.	Undetm.	Undetm.	<i>C. imbricata</i>	MLU	[102]
(+)-Precapnelladiene 10b ^γ [C ₁₅ H ₂₄]	TS	Precapnellane	Undetm.	Undetm.	Undetm.			[49–52]
(+)-3α,4α-Epoxyprecapnell-10-ene 11 ^β [C ₁₅ H ₂₄ O]	MS, NMR, [α] _D	Precapnellane	Cytostatic Cytotoxic	L-929, K562 HeLa	GI ₅₀ = 227.3 ± 1.8–4.3 μM CC ₅₀ = 193.2 ± 4.3 μM	<i>D. rubeola</i>	BLI	[103]
(+)-Hydroxycolorenone 12 ^β [C ₁₅ H ₂₄ O ₂]	UV, MS, NMR, [α] _D	Guaiane	Antiinsecticidal	<i>S. littoralis</i>	EC ₅₀ = 8.8 ± 0.26 μg/mL LC ₅₀ = 453 ± 0.43 μg/mL	<i>N. chabrolia</i>	WST	[104]
(+)-Methoxycolorenone 13 ^β [C ₁₆ H ₂₆ O ₂]	UV, MS, NMR, [α] _D	Guaiane	Antiinsecticidal	<i>S. littoralis</i>	NA	<i>N. chabrolia</i>	WST	[104]

Table S1: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)- 14 ^β [C ₁₃ H ₂₂ O ₂]	IR, MS, NMR, [α] _D , QCC	Trinor-guaiane	Cytotoxic	NBT-T2	NA (10 µg/mL)	<i>Anthelia</i> sp.	BTN	[105]
(+)-Helianane 15a ^β [C ₁₅ H ₂₂ O ₂]	UV, MS, NMR, [α] _D	Helianane	Undetm.	Undetm.	Undetm.	<i>H. fascigera</i>	NSW	[106]
(+)-Curcudiol 15b ^δ [C ₁₅ H ₂₄ O ₂]	TS	Bisabolane	Undetm.	Undetm.	Undetm.			[53–57]
Boneratamide A methyl ester 16c ^ε [C ₂₄ H ₄₀ N ₂ O ₄]	MS, NMR, X-ray	Spiroaxane	Cytostatic	Antimitotic assay	NA	<i>A. aplysinoides</i>	SSW	[107]
Boneratamides B 17c /C 18c ^{α,ε} methyl esters [C ₂₄ H ₃₈ N ₂ O ₄]	MS, NMR	Spiroaxane	Cytostatic	Antimitotic assay	NA	<i>A. aplysinoides</i>	SSW	[57]
(-)-Δ ⁹⁽¹²⁾ -Capnellene 19 ^β [C ₁₅ H ₂₄]	IR, MS, NMR, [α] _D , CT	Capnellane	Undetm.	Undetm.	Undetm.	<i>C. imbricata</i>	MLU	[108]
(+) -Capnellene-8β-ol 20 ^β [C ₁₅ H ₂₄ O]	IR, MS, NMR, [α] _D	Capnellane	Cytotoxic	MCF7, HT-115 HL-60 K562, A-2780 G-402	IC ₅₀ > 4500 µM IC ₅₀ = 68 µM IC ₅₀ = 4.6 – 6.6 µM IC ₅₀ > 4.5 µM	<i>C. imbricata</i>	NMU	[109, 62]
			Anti-inflammatory	RAW 264.7 (LPS/iNOS, COX-2)	NA			
			Toxic	<i>C. septentrionalis</i> , <i>A. japonica</i> , <i>T. excentricus</i> , <i>P. micans</i> , <i>A. carterae</i>	Quan. Undetm.			
			Cytotoxic	KB, HL-60, G-402, HT-115, MCF7 K562, WIDr, A2780	IC ₅₀ = 25.6 – 93 µM IC ₅₀ = 0.7 – 9.7 µM			
			Cytostatic	HeLa L-929	CC ₅₀ = 7.6 ± 0.8 µM IC ₅₀ = 15.1 µM			
(+) -Δ ⁹⁽¹²⁾ -Capnellene-8β,10α-diol 21 ^β [C ₁₅ H ₂₄ O ₂]	IR, MS, NMR, [α] _D , CT	Capnellane	Cytostatic	RAW 264.7 (LPS/ iNOS, COX-2)	GI ₅₀ = 6.8 ± 0.8 µM 1.2 ± 0.1 – 24.8 ± 7.5% (10 µM)	<i>C. imbricata</i>	MLU	[62, 103, 109 – 112]
			Anti-inflammatory	BV2 (IFN-γ/iNOS, COX-2)	IC ₅₀ = 6.21 ± 2.5 – 17.1 ± 2.8 µM			
			Antinociception	Murine neuropathy	10 mg/kg (CCI therm. hyperalgesia behav.)			

Table S1: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)- $\Delta^{9(12)}$ -Capnellene-3 β ,8 β ,10 α -triol 22 ^{β} [C ₁₅ H ₂₄ O ₃]	UV, IR, MS, NMR, ECD, [α] _D , CT, X-ray	Capnellane	Toxic	<i>P. micans</i> , <i>A. carterae</i>	Quan. Undetm.	<i>C. imbricata</i>	MLU	[62, 111, 113]
(+)- $\Delta^{9(12)}$ -Capnellene-5 α ,8 β ,10 α -triol 23 ^{β} [C ₁₅ H ₂₄ O ₃]	IR, MS, NMR, [α] _D , CT	Capnellane	Undetm.	Undetm.	Undetm.	<i>C. imbricata</i>	MLU	[111]
$\Delta^{9(12)}$ -Capnellene-2 ξ ,8 β ,10 α -triol 24 ^{β} [C ₁₅ H ₂₄ O ₃]	IR, MS, NMR, [α] _D , CT	Capnellane	Undetm.	Undetm.	Undetm.	<i>C. imbricata</i>	MLU	[111]
(+)-3 α ,8 β -Diacetoxycapnell-9(12)-ene-10 α -ol 25 ^{β} [C ₁₉ H ₂₈ O ₅]	MS, NMR, [α] _D	Capnellane	Cytotoxic Cytostatic	HeLa K562, L-929	CC ₅₀ > 125 μ M GI ₅₀ = 62.2 $\pm$ 2.7 – 99.1 $\pm$ 1.8 μ M	<i>D. rubeola</i>	BLI	[103]
(+)- $\Delta^{9(12)}$ -Capnellene-3 β ,8 β ,10 α ,14-tetraol 26 ^{β} [C ₁₅ H ₂₄ O ₄]	IR, MS, NMR, [α] _D , CT	Capnellane	Undetm.	Undetm.	Undetm.	<i>C. imbricata</i>	MLU	[108, 114]
(+)-3 β -Acetoxycapnellene-8 β ,10 α ,14 β -triol 27 ^{β} [C ₁₇ H ₂₆ O ₅]	IR, MS, NMR, [α] _D	Capnellane	Cytotoxic	HL-60, MCF7 K562, G-402, A2780 HT-115	IC ₅₀ = 713 – 1029 μ M IC ₅₀ = 24 – 52 μ M NA	<i>C. imbricata</i>	NMU	[109]
(+)-3 α ,14-Diacetoxycapnell-9(12)-ene-8 β ,10 α -diol 28 ^{β} [C ₁₉ H ₂₈ O ₆]	MS, NMR, [α] _D	Capnellane	Cytostatic	K562	GI ₅₀ = 142.0 $\pm$ 4.7 μ M	<i>D. rubeola</i>	NSW	[103]
(+)-3 α ,8 β ,14-Triacetoxycapnell-9(12)-ene-10 α -ol 29 ^{β} [C ₂₁ H ₃₀ O ₇]	MS, NMR, [α] _D	Capnellane	Cytostatic Cytotoxic	K562 HeLa	GI ₅₀ = 126.9 $\pm$ 0.2 μ M CC ₅₀ > 125 μ M	<i>D. rubeola</i>	NSW	[103]
(-)-2 α ,8 β ,13-Triacetoxycapnell-9(12)-ene-10 α -ol 30 ^{β} [C ₂₁ H ₃₀ O ₇]	MS, NMR, [α] _D	Capnellane	Cytostatic Cytotoxic	K562 HeLa	GI ₅₀ = 126.9 $\pm$ 3.0 μ M CC ₅₀ > 125 μ M	<i>D. rubeola</i>	NSW	[103]
(-)-Hirsutanol A 31 ^{β} [C ₁₅ H ₁₈ O ₃]	IR, MS, NMR, [α] _D , X-ray, CT	Hirsutane	Cytotoxic	SW480, CNE2 SW620, LoVo, Hep3B, SUNE1 MCF7, CNE1, CNE2, A549	ED ₅₀ = 3.03 $\pm$ 0.07 μ g/mL ED ₅₀ = 0.58 $\pm$ 0.09 – 0.90 $\pm$ 0.19 μ g/mL ED ₅₀ = 2.55 $\pm$ 0.41 – 3.13 $\pm$ 0.29 μ g/mL	A fungus (symbiont) <i>Haliclona</i> sp. (host)	GTO	[58, 115 – 117]

Table S1: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Hirsutanol A 31 ^β [C ₁₅ H ₁₈ O ₃]	IR, MS, NMR, [α] _D , X-ray, CT	Hirsutane	Cytotoxic	MDA-MB-231, MDA-MB-435	ED ₅₀ = 1.34 ± 0.19 – 1.82 ± 0.37 µg/mL	A fungus (symbiont) <i>Haliclona</i> sp. (host)	GTO	[58, 115 – 117]
				HepG2	IC ₅₀ = 2.45 ± 0.13 µg/mL			
				Bel-7402	IC ₅₀ = 6.11 ± 0.41 µg/mL			
			HeLa	ED ₅₀ = 8.27 ± 0.71 µg/mL				
			B16	ED ₅₀ = 25.6 µM				
Antibacterial	<i>B. subtilis</i>	200 µg/disk	<i>B. subtilis</i> ATCC 70385, <i>S. aureus</i> 6538, <i>E. coli</i> ATCC 8739	MIC > 100 µM				
Hirsutanol B 32 ^β [C ₁₅ H ₁₈ O ₃]	NMR	Hirsutane	Undetm.	Undetm.	Undetm.	A fungus (symbiont) <i>Haliclona</i> sp. (host)	GTO	[58, 115 – 117]
(+)-Hirsutanol C 33a ^β [C ₁₅ H ₁₈ O ₃]	IR, MS, NMR, [α] _D	Hirsutane	Cytotoxic	LoVo	IC ₅₀ > 200 µM	A fungus (symbiont) <i>Haliclona</i> sp. (host)	GTO	[58, 115 – 117]
(-)-Hirsutanol C 33b ^γ [C ₁₅ H ₁₈ O ₃]	CT	Hirsutane	Cytotoxic	B16	IC ₅₀ > 200 µM	A fungus (symbiont)	JPN	[58, 115 – 117]
			Antibacterial	<i>B. subtilis</i> ATCC 70385, <i>S. aureus</i> 6538, <i>E. coli</i> ATCC 8739	MIC > 100 µM	<i>G. incarnatum</i> (host)		
<i>ent</i> -Gloeosteretriol [(-)-Hirsutanol F] 34 ^β [C ₁₅ H ₂₆ O ₃]	UV, IR, MS, NMR, [α] _D	Hirsutane	Cytotoxic	SW480, MCF7, MDA-MB-231, MDA-MB-435, MDA-MB-453, CNE1, CNE2, SUNE1, A549, Hep3B, HepG2, Bel-7402, HeLa	ED ₅₀ > 50 µg/mL	A fungus (symbiont) <i>Haliclona</i> sp. (host)	GTO	[58, 115 – 117]
			Antibacterial	<i>B. subtilis</i>	NA (200 µg/disk)			
(+)-Africanol 35 ^β [C ₁₅ H ₂₆ O]	IR, MS, NMR, [α] _D , X-ray	Africanane	Toxic	<i>L. reticulatus</i> , <i>C. septentrionalis</i> , <i>A. japonica</i> , <i>T. excentricus</i> , <i>P. micans</i> , <i>A. carterae</i>	Quan. Undetm.	<i>L. africana</i> , <i>L. nitida</i>	MLU	[59, – 62, 118]
(+)-4α,7β-Aromadendranediol 36 ^β [C ₁₅ H ₂₆ O ₂]	IR, MS, NMR, [α] _D , CT	Aromaden-drane	Undetm.	Undetm.	Undetm.	<i>S. mayi</i>	NST	[119]

Table S1: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-4 α ,7 α -Aromadendranediol 37 ^β [C ₁₅ H ₂₆ O ₂]	IR, MS, NMR, [α] _D , CT	Aromaden- drane	Undetm.	Undetm.	Undetm.	<i>S. mayi</i>	NST	[119]
(-)-Sinularane 38 ^β [C ₁₅ H ₂₄]	IR, MS, NMR, [α] _D , ORD, ECD, X-ray, CT	Sinularane	Undetm.	Undetm.	Undetm.	<i>S. mayi</i>	NST	[119, 120]
9-Thiocyanatopupukeanane 39/9-epi 40 ^{α,β} [C ₁₆ H ₂₅ NS]	MS, NMR	Pupukeanane*	Cytotoxic Antibacterial Antifungal Cytotoxic, Anti- microbial, Antiathero- sclerotic, Antiosteo- clastic, Anticancer	<i>A. salina</i> <i>B. subtilis</i> <i>C. albicans</i>	90% (39:40 = 30:70) 35% (39:40 = 50:50) W (39:40 = 30:70, 20 μg) M (39:40 = 30:70, 20 μg)	<i>P. varicosa</i> , <i>A. aculeata</i>	JSCR	[121]
(-)-Lamellodysidine A 41 ^β [C ₁₅ H ₂₀ O ₃]	UV, IR, MS, NMR, ECD, [α] _D	Lamello- dysidine*	Cytotoxic, Anti- microbial, Antiathero- sclerotic, Antiosteo- clastic, Anticancer	Several cell lines, microorganisms, enzymes	NA	<i>L. herbacea</i>	NSW	[36]
(-)-Lamellodysidine B 42 ^β [C ₁₇ H ₂₃ NO ₃]	UV, IR, MS, NMR, ECD, [α] _D	Nakafurane 8	Cytotoxic, Anti- microbial, Antiathero- sclerotic, Antiosteo- clastic, Anticancer	Several cell lines, microorganisms, enzymes	NA	<i>L. herbacea</i>	NSW	[36]

Footnote: 1. **Structure** (^αmolecule isolated as a mixture, ^βoriginal molecule, ^γrevised molecule (new), ^δrevised molecule (known), ^εmolecule isolated from its derivative using chemical reaction, skeleton names in black were already reported in the literature; those in blue were named in this review, ^{*}new skeleton, [†]rare FG, [‡]rare motif); 2. **Structure Elucidation** (UV ultraviolet spectroscopy, IR infrared spectroscopy, MS mass spectrometry, NMR nuclear magnetic resonance, ECD electronic circular dichroism, X-ray single-crystal X-ray diffraction, Mol. Mod. molecular modeling, QCC quantum chemical calculation, CT chemical transformation, TS total synthesis, [α]_D specific optical rotation); 3. **Statistic** (CC₅₀ 50% of the cytotoxic concentration, EC₅₀ 50% of the effective concentration, ED₅₀ 50% of the effective dose, GI₅₀ 50% of the growth inhibition which emphasizes the correction for the cell count at time zero, IC₅₀ 50% of the inhibitory concentration, LC₅₀ 50% of the lethal concentration, MIC minimum inhibitory concentration); 4. **Activity** (NA no activity or not active, B16 murine melanoma, BV2 murine microglia, L-929 murine fibroblast, NBT-T2 murine bladder epithelial tumor, RAW 264.7 murine macrophage, A549 human lung carcinoma, A2780 human ovarian carcinoma, Bel-7402 human hepatocarcinoma, CNE1 human nasopharyngeal carcinoma, CNE2 human nasopharyngeal carcinoma, G-402 human renal leiomyoblastoma, HeLa human cervical carcinoma, Hep3B human hepatocarcinoma, HepG2 human hepatocarcinoma, HL-60 human acute promyelocytic leukemia, HT-115 human colorectal adenocarcinoma, K562 human erythro myeloblastoid leukemia, KB human nasopharyngeal epidermoid carcinoma, LoVo human colorectal adenocarcinoma, MCF7 human breast adenocarcinoma, MDA-MB-231 human breast adenocarcinoma, MDA-MB-435 human breast adenocarcinoma, SUNE1 human nasopharyngeal carcinoma, SW480 human colorectal adenocarcinoma, SW620 human colorectal adenocarcinoma, WIDr human colorectal adenocarcinoma, COX-2 cyclooxygenase-2 enzyme, iNOS inducible nitric oxide synthase enzyme, LPS lipopolysaccharide, IFN-γ interferon gamma, CCI chronic constriction injury, **behav.** behaviour, **Quan.** Undetm quantitatively undetermined, **therm** thermal, **Undetm.** undetermined, **W** weak, **M** Moderate); 5. Geography (JPN Japan).

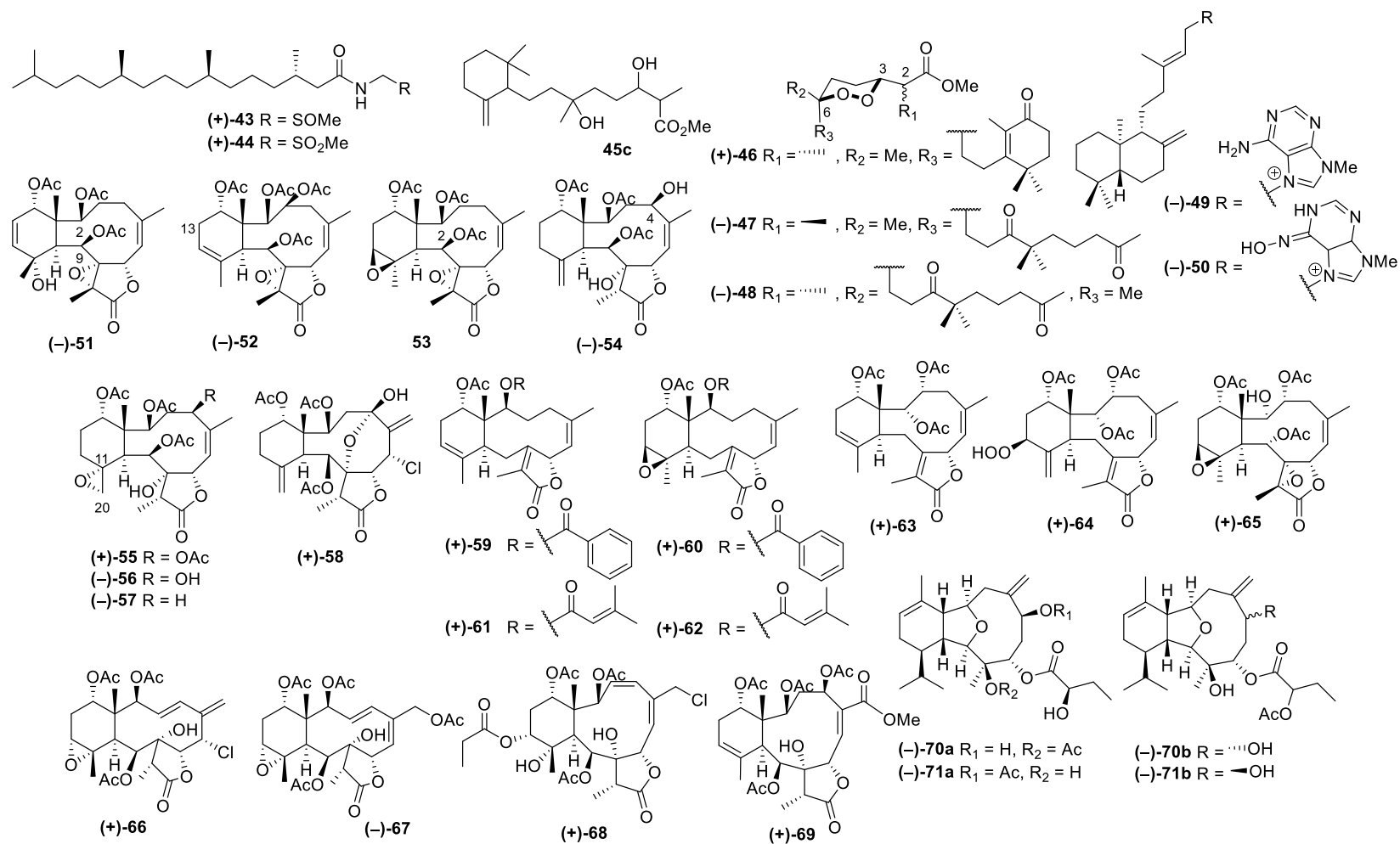


Figure S2: Cont.

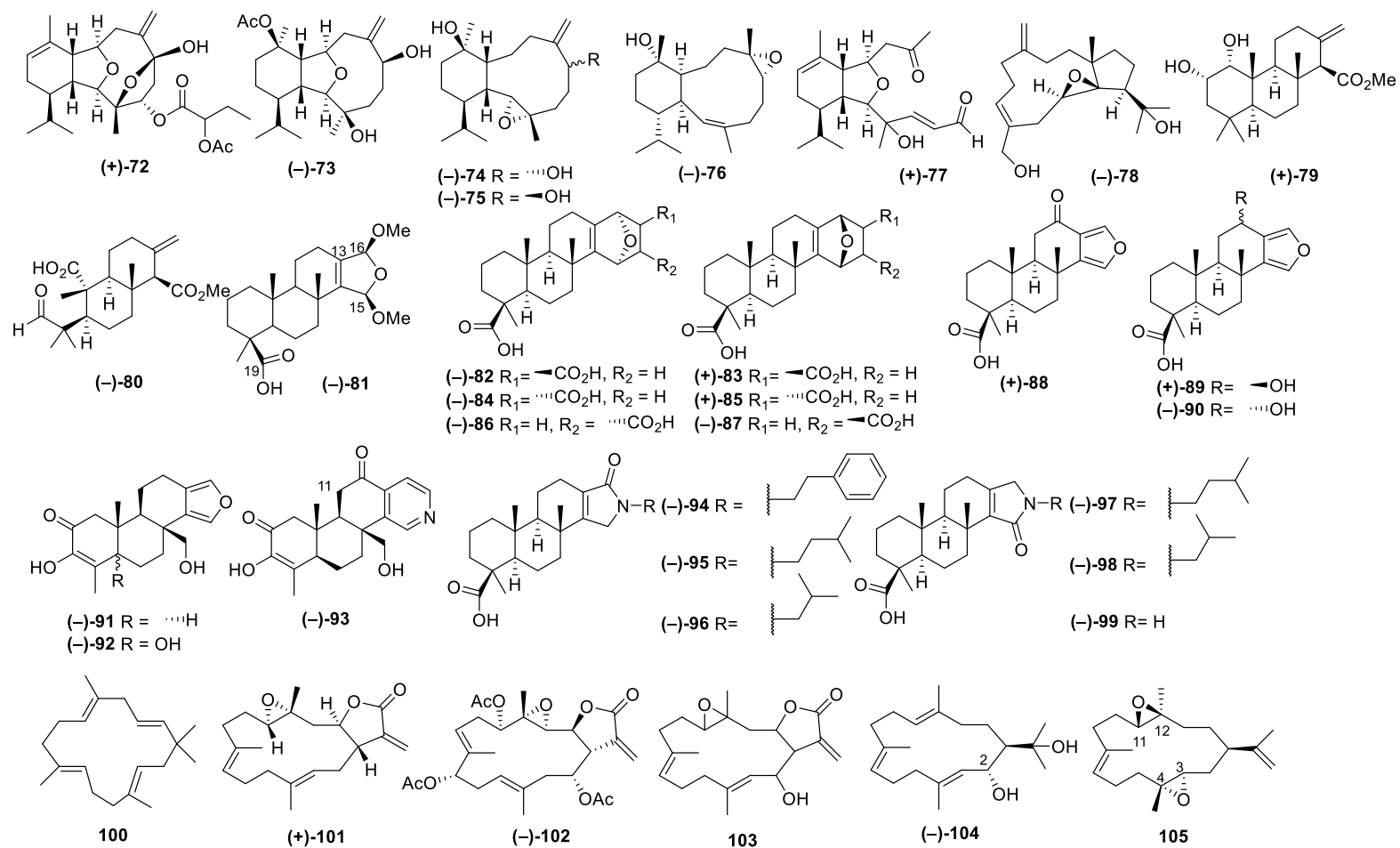


Figure S2: Cont.

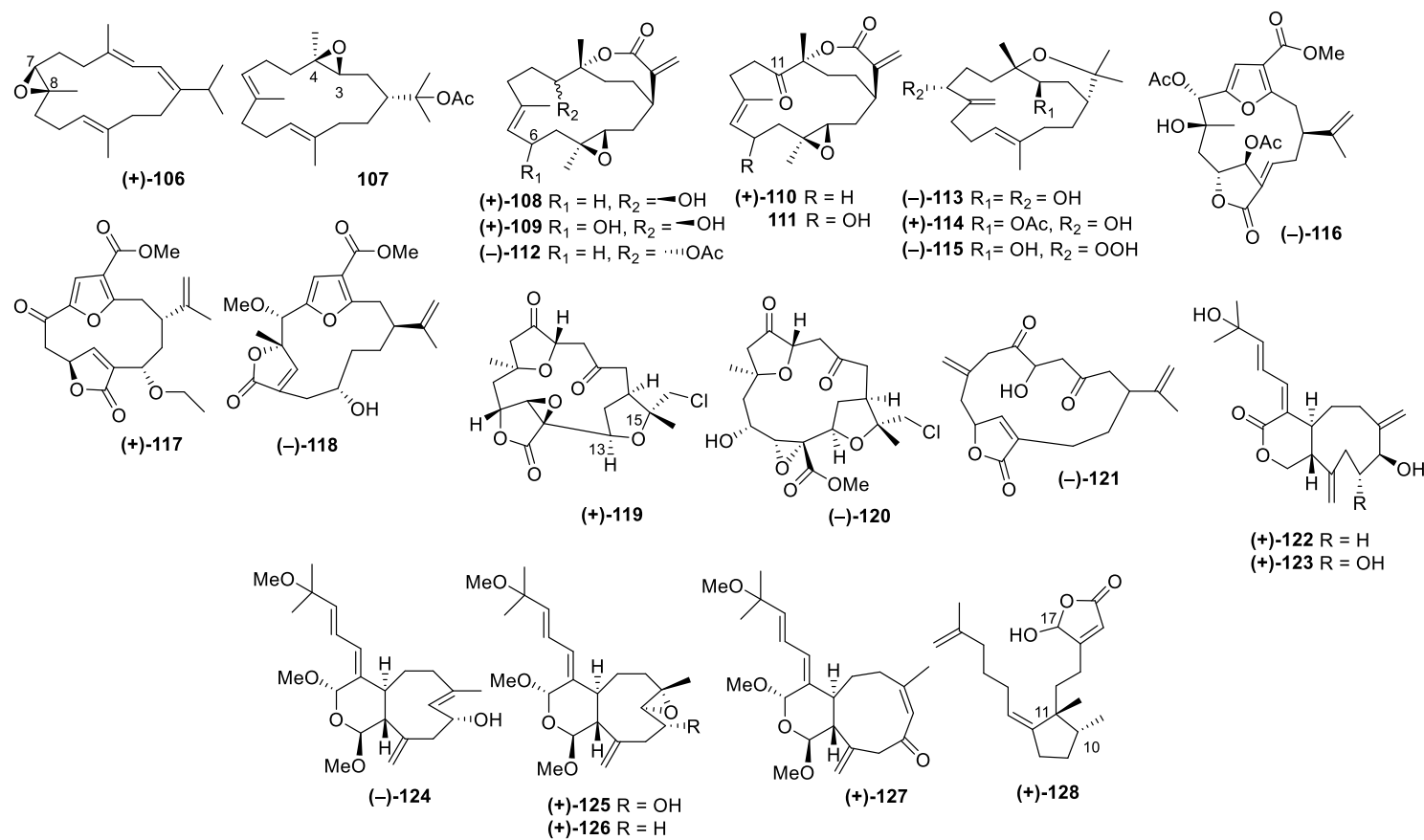


Figure S2: Structures of marine diterpenoids from Indonesian waters found in 1970–2017.

Table S2: Marine diterpenoids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Sinulasulfoxide 43 ^β [C ₂₃ H ₄₇ NO ₂ S]	IR, MS, NMR, ECD, [α] _D , CT	Acyclic diterpene [•]	Anti-inflammatory	J774 (LPS/iNOS)	22.6% (30 μM)	<i>Sinularia</i> sp.	NSW	[122]
(+)-Sinulasulfone 44 ^β [C ₂₃ H ₄₇ NO ₃ S]	NMR, MS, ECD, [α] _D , CT	Acyclic diterpene [•]	Undetm.	Undetm.	Undetm.	<i>Sinularia</i> sp.	NSW	[122]
<i>nor</i> -Diterpene 45 ^c [C ₂₀ H ₃₆ O ₄]	IR, MS, NMR, CT	Acyclic diterpene	Undetm.	Undetm.	Undetm.	<i>D. megaspi-norhabdosa</i>	SSW	[123]
(+)-Diacaperoxide A 46 ^β [C ₂₀ H ₃₂ O ₅]	MS, NMR, [α] _D	Acyclic peroxide diterpene	Cytotoxic	L5178Y HeLa, PC12 <i>A. salina</i>	EC ₅₀ > 10 μg/mL NA 35 – 55% (10 μg/mL, 24 – 48 h)	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Diacaperoxide B 47 ^β [C ₂₀ H ₃₄ O ₆]	MS, NMR, [α] _D	Acyclic peroxide diterpene	Cytotoxic	L5178Y HeLa, PC12 <i>A. salina</i>	EC ₅₀ = 10 μg/mL NA 45 – 55% (10 μg/mL, 24 – 48 h)	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Diacaperoxide C 48 ^β [C ₂₀ H ₃₄ O ₆]	MS, NMR, [α] _D	Acyclic peroxide diterpene	Cytotoxic	L5178Y HeLa, PC12 <i>A. salina</i>	EC ₅₀ > 10 μg/mL NA 45 – 55% (10 μg/mL, 24 – 48 h)	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Agelasine D 49 ^β [C ₂₆ H ₄₀ N ₅]	UV, MS, NMR, [α] _D	Copalane	Cytotoxic Antibacterial	L5178Y <i>S. epidermidis</i>	IC ₅₀ = 4.03 μM MIC < 0.0877 μM	<i>A. nakamurai</i>	JSCR	[125]
(-)-Ageloxime D 50 ^β [C ₂₆ H ₄₂ N ₅ O]	UV, MS, NMR, [α] _D	Copalane	Cytotoxic Antibacterial	L5178Y <i>S. epidermidis</i>	IC ₅₀ = 12.5 μM MIC > 45 μM	<i>A. nakamurai</i>	JSCR	[125]
(-)-2,9-Diacetyl-2-debutyrylstecholide H 51 ^β [C ₂₆ H ₃₄ O ₁₀]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ > 10 μg/mL	<i>Briareum</i> sp.	CSW	[126, 127]
(-)-13-Dehydroxystecholide J 52 ^β [C ₂₆ H ₃₈ O ₁₁]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ > 10 μg/mL	<i>Briareum</i> sp.	CSW	[126, 127]
2β-Acetoxy-2-(debutyryloxy)-stecholide E acetate 53 ^c [C ₂₆ H ₃₄ O ₁₀]	MS, NMR, [α] _D	Briarane	Cytotoxic	P-388 A549, HT-29, MEL-28	IC ₅₀ > 10 μg/mL EC ₅₀ = 1.59 μg/mL IC ₅₀ > 10 μg/mL	<i>Briareum</i> sp.	CSW	[126, 127]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-4-Deacetyljunceollolide D 54 ^β [C ₂₆ H ₃₆ O ₁₀]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	NA	<i>J. fragilis</i>	NMU	[128]
(+)-11α,20α-Epoxyjunceollolide D 55 ^β [C ₂₈ H ₃₈ O ₁₂]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	NA	<i>J. fragilis</i>	NMU	[128]
(-)-11α,20α-Epoxy-4-deacetyljunceollolide D 56 ^β [C ₂₈ H ₃₆ O ₁₁]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	NA	<i>J. fragilis</i>	NMU	[128]
(-)-11α,20α-Epoxy-4-deacetoxyjunceollolide D 57 ^β [C ₂₈ H ₃₆ O ₁₀]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	NA	<i>J. fragilis</i>	NMU	[128]
(+)-Junceollolide A 58 ^β [C ₂₆ H ₃₃ ClO ₁₀]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	P-388, A549, HT-29, MEL-28	NA	<i>J. fragilis</i>	NMU	[128]
(+)-Malayenolide A 59 ^β [C ₂₉ H ₃₄ O ₆]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	<i>A. salina</i>	LC ₅₀ = 100 µg/mL	<i>V. malayense</i>	NSW	[129]
(+)-Malayenolide B 60 ^β [C ₂₉ H ₃₄ O ₇]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	<i>A. salina</i>	LC ₅₀ < 2 µg/mL	<i>V. malayense</i>	NSW	[129]
(+)-Malayenolide C 61 ^β [C ₂₇ H ₃₆ O ₆]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	<i>A. salina</i>	LC ₅₀ = 20 µg/mL	<i>V. malayense</i>	NSW	[129]
(+)-Malayenolide D 62 ^β [C ₂₇ H ₃₆ O ₇]	UV, MS, NMR, [α] _D	Briarane	Cytotoxic	<i>A. salina</i>	LC ₅₀ = 20 µg/mL	<i>V. malayense</i>	NSW	[129]
(+)–Brianthein A 63 ^β [C ₂₆ H ₃₄ O ₈]	MS, NMR, [α] _D , CT, Mol. Mod.	Briarane	Cytotoxic	KB-3-1	11 ± 6 – 27 ± 5% (3 – 10 µg/mL)	<i>B. excavatum</i>	UEP	[130]
			Cytostatic	KB-C2, P-gp type	60 ± 2 – 84 ± 3% (0.1 µg/mL colchicine 3 – 10 µg/mL)			
			Cytotoxic	KB-3-1	5 ± 1 – 26 ± 4% (3 – 10 µg/mL)			
(+)–Brianthein B 64 ^β [C ₂₆ H ₃₄ O ₁₀]	MS, NMR, [α] _D , CT	Briarane	Cytostatic	KB-C2, P-gp type	26 ± 4 – 37 ± 6% (0.1 µg/mL colchicine 3 – 10 µg/mL)	<i>B. excavatum</i>	UEP	[130]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Brianthein C 65 ^β [C ₂₆ H ₃₄ O ₁₁]	MS, NMR, [α] _D , CT	Briarane	Cytotoxic	KB-3-1	11 ± 1 – 17 ± 6% (3 – 10 µg/mL)	<i>B. excavatum</i>	UEP	[130]
			Cytostatic	KB-C2, P-gp type	0 ± 0 – 15 ± 2% (0.1 µg/mL colchicine 3 – 10 µg/mL)			
(+)– 66 ^β [C ₂₆ H ₃₃ ClO ₁₀]	IR, MS, NMR, [α] _D	Briarane	Cytostatic	KB-C2, P-gp type KB-CV60, MRP-1 type	NA	<i>Pteroeides</i> sp	ENT	[131]
(–)– 67 ^β [C ₂₈ H ₃₆ O ₁₂]	IR, MS, NMR, [α] _D	Briarane	Cytostatic	KB-C2, P-gp type KB-CV60, MRP-1 type	NA	<i>Pteroeides</i> sp	ENT	[131]
(+)– 68 ^β [C ₂₉ H ₃₉ ClO ₁₂]	IR, MS, NMR, [α] _D	Briarane	Cytostatic	KB-C2, P-gp type KB-CV60, MRP-1 type	NA	<i>Pteroeides</i> sp	ENT	[131]
(+)– 69 ^β [C ₂₉ H ₃₈ O ₁₃]	IR, MS, NMR, [α] _D	Briarane	Cytostatic	KB-C2, P-gp type KB-CV60, MRP-1 type	NA	<i>Pteroeides</i> sp.	ENT	[131]
(–)–Cladielloide A 70a ^β [C ₂₆ H ₄₀ O ₇]	IR, MS, NMR, [α] _D , CT	Cladiellane	Cytotoxic	DLD-1, HL-60, CCRF-CEM, P388D1	IC ₅₀ > 40 µg/mL	<i>Cladiella</i> sp.	UEP	[132]
			Anti-inflammatory	Human neutrophils	20.5 ± 5.0% SA gen. FMLP/CB (10 µg/mL) 27.1 ± 4.8% elastase rel. FLMP/CB (10 µg/mL)			
(–)–Cladielloide A 70b ^γ [C ₂₆ H ₄₀ O ₇]	NMR 2D (NOESY)	Cladiellane	Undetm.	Undetm.	Undetm.	<i>Cladiella</i> sp.	UEP	[133]
(–)–Cladielloide B 71a ^β [C ₂₆ H ₄₀ O ₇]	IR, MS, NMR, [α] _D	Cladiellane	Cytotoxic	DLD-1, CCRF-CEM HL-60, P388D1	IC ₅₀ = 4.7 – 10.2 µg/mL IC ₅₀ > 40 µg/mL	<i>Cladiella</i> sp.	UEP	[132]
			Anti-inflammatory	Human neutrophils	5.9 ± 0.7% SA gen. FMLP/CB (10 µg/mL) 6.5 ± 1.9% elastase rel. FMLP/CB (10 µg/mL)			
(–)–Cladielloide B 71b ^γ [C ₂₆ H ₄₀ O ₇]	NMR 2D (NOESY)	Cladiellane	Undetm.	Undetm.	Undetm.	<i>Cladiella</i> sp.	UEP	[133]
(+)–Cladielloide C 72 ^β [C ₂₅ H ₃₈ O ₇]	IR, MS, NMR, [α] _D , Mol. Mod.	Cladiellane	Cytotoxic	DLD-1, P388D1, HL-60 CCRF-CEM	IC ₅₀ = 8.5 – 12.6 µg/mL IC ₅₀ = 3.6 µg/mL	<i>Cladiella</i> sp.	UEP	[134]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Cladielloide C 72 ^β [C ₂₅ H ₃₈ O ₇]	IR, MS, NMR, [α] _D , Mol. Mod.	Cladiellane	Anti-inflammatory	Human neutrophils	36.7 ± 7.6% SA gen. FMLP/CB (10 µg/mL) 27.2 ± 3.6% elastase rel. FMLP/CB (10 µg/mL) 1.97 ± 2.44% SA gen.	<i>Cladiella</i> sp.	UEP	[134]
(-)-Cladieunicellin G 73 ^β [C ₂₂ H ₃₆ O ₅]	IR, MS, NMR, [α] _D , Mol. Mod.	Cladiellane	Anti-inflammatory	Human neutrophils	FMLP/CB (10 µg/mL) 12.89 ± 5.03% elastase rel. FMLP/CB (10 µg/mL) 6.46 ± 1.28% SA gen.	<i>Cladiella</i> sp.	UEP	[135]
(-)-Cladieunicellin F 74 ^β [C ₂₀ H ₃₄ O ₃]	IR, MS, NMR, [α] _D	Cladiellane	Anti-inflammatory	Human neutrophils	FMLP/CB (10 µg/mL) 12.91 ± 3.56% elastase rel. FMLP/CB (10 µg/mL) 6.57 ± 0.85 % SA gen.	<i>Cladiella</i> sp.	UEP	[133]
(-)-6- <i>epi</i> -Cladieunicellin F 75 ^β [C ₂₀ H ₃₄ O ₃]	IR, MS, NMR, [α] _D , Mol. Mod.	Cladiellane	Anti-inflammatory	Human neutrophils	FMLP/CB (10 µg/mL) 41.98 ± 3.26% elastase rel. FMLP/CB (10 µg/mL) 45.82 ± 2.49% SA gen.	<i>Cladiella</i> sp.	UEP	[133]
(-)-Solenopodin C 76 ^β [C ₂₀ H ₃₄ O ₂]	IR, MS, NMR, [α] _D	Cladiellane	Anti-inflammatory	Human neutrophils	FLMP/CB (10 µg/mL) 40.45 ± 5.80% elastase rel. FLMP/CB (10 µg/mL) IC ₅₀ = 11.6 – 35.1 µg/mL IC ₅₀ > 40 µg/mL	<i>Cladiella</i> sp.	UEP	[133]
(+)-Cladielloide D 77 ^β [C ₂₀ H ₃₀ O ₄]	IR, MS, NMR, [α] _D , Mol. Mod	<i>seco</i> - cladiellane [▲]	Cytotoxic Anti-inflammatory	DLD-1, CCRF-CEM HL-60, P388D1 Human neutrophils	31.4 ± 6.9% SA gen. FMLP/CB (10 µg/mL) 10.7 ± 5.6% elastase rel. FMLP/CB (10 µg/mL)	<i>Cladiella</i> sp.	UEP	[134]
(-)- 78 ^β [C ₂₀ H ₃₂ O ₃]	IR, MS, NMR, [α] _D	Dollabellane	Cytotoxic	NBT-T2	10 µg/mL	<i>Anthelia</i> sp.	BTN	[136]
(+)-Coelodiol 79 ^β [C ₂₁ H ₃₄ O ₄]	IR, MS, NMR, [α] _D , ECD, CT	Isocopalane [▲]	Cytostatic	MKN-45	20 µg/mL	<i>C. cfr.</i> <i>singaporensis</i>	NSW	[137]
(-)-Coeloic acid 80 ^β [C ₂₀ H ₃₀ O ₅]	IR, MS, NMR, [α] _D	<i>seco</i> -nor isocopalane [▲]	Cytostatic	MKN-45	40 µg/mL	<i>C. cfr.</i> <i>singaporensis</i>	NSW	[137]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-15 α ,16-Dimethoxyspongi-13-en-19-oic acid 81 ^{β} [C ₂₂ H ₃₄ O ₅]	IR, MS, NMR, [α] _D , Mol. Mod	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ > 50 μ M 98% (5 μ M)	<i>S. ceylonensis</i>	NSW	[138]
(-)-Ceylonin A 82 ^{β} [C ₂₂ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D , ECD	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	70% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(+)-Ceylonin B 83 ^{β} [C ₂₃ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	<28% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(-)-Ceylonin C 84 ^{β} [C ₂₃ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	<28% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(+)-Ceylonin D 85 ^{β} [C ₂₃ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	28% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(-)-Ceylonin E 86 ^{β} [C ₂₃ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	47% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(-)-Ceylonin F 87 ^{β} [C ₂₃ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP)	31% (50 μ M)	<i>S. ceylonensis</i>	NSW	[139]
(+)-Ceylonin G 88 ^{β} [C ₂₀ H ₂₆ O ₄]	UV, MS, NMR, [α] _D , ECD	Spongiane	Anticancer	USP7	IC ₅₀ > 50 μ M	<i>S. ceylonensis</i>	NSW	[140]
(+)-Ceylonin H 89 ^{β} [C ₂₀ H ₂₈ O ₄]	UV, MS, NMR, [α] _D , ECD, Mol. Mod.	Spongiane	Anticancer	USP7	IC ₅₀ > 50 μ M	<i>S. ceylonensis</i>	NSW	[140]
(-)-Ceylonin I 90 ^{β} [C ₂₀ H ₂₈ O ₄]	UV, MS, NMR, [α] _D , ECD, Mol. Mod.	Spongiane	Anticancer	USP7	IC ₅₀ > 50 μ M	<i>S. ceylonensis</i>	NSW	[140]
(-)-18-nor-3,17-Dihydroxyspongia-3,13(16),14-trien-2-one 91 ^{β} [C ₁₉ H ₂₄ O ₄]	UV, IR, MS, NMR, [α] _D	Spongiane	Alzheimer	BACE1	IC ₅₀ < 100 μ M	<i>Spongia</i> sp.	NSW	[141]
(-)-18-nor-3,5,17-Trihydroxyspongia 3,13(16),14-trien-2-one 92 ^{β} [C ₁₉ H ₂₄ O ₅]	UV, IR, MS, NMR, [α] _D	Spongiane	Alzheimer	BACE1	IC ₅₀ < 100 μ M	<i>Spongia</i> sp.	NSW	[141]
(-)-Spongiapyridine 93 ^{β} [C ₂₀ H ₂₃ NO ₄]	UV, IR, MS, NMR, [α] _D	Spongiane	Alzheimer	BACE1	IC ₅₀ < 100 μ M	<i>Spongia</i> sp.	NSW	[141]
(-)-Ceylonamide A 94 ^{β} [C ₂₈ H ₃₇ NO ₃]	UV, MS, NMR, [α] _D , Mol. Mod	Spongiane	Antiestrogenic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ = 13 μ M 100% (5 μ M)	<i>S. ceylonensis</i>	NSW	[138]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Ceylonamide B 95 ^β [C ₂₅ H ₃₉ NO ₃]	UV, MS, NMR, [α] _D	Spongiane	Antisteoclastic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ = 18 μM 87% (5 μM)	<i>S. ceylonensis</i>	NSW	[138]
(-)-Ceylonamide C 96 ^β [C ₂₄ H ₃₇ NO ₃]	UV, MS, NMR, [α] _D	Spongiane	Antisteoclastic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ > 50 μM 100% (5 μM)	<i>S. ceylonensis</i>	NSW	[138]
(-)-Ceylonamide D 97 ^β [C ₂₅ H ₃₉ NO ₃]	UV, MS, NMR, [α] _D	Spongiane	Antisteoclastic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ > 50 μM 100% (5 μM)	<i>S. ceylonensis</i>	NSW	[138]
(-)-Ceylonamide E 98 ^β [C ₂₄ H ₃₇ NO ₃]	UV, MS, NMR, [α] _D	Spongiane	Antisteoclastic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ > 50 μM 100% (5 μM)	<i>S. ceylonensis</i>	NSW	[138]
(-)-Ceylonamide F 99 ^β [C ₂₀ H ₂₉ NO ₃]	UV, MS, NMR, [α] _D	Spongiane	Antisteoclastic	RAW264 (RANKL/TRAP) RAW264 cell survival ratio	IC ₅₀ > 50 μM 100% (5 μM)	<i>S. ceylonensis</i>	NSW	[138]
Flexibilene 100 ^β [C ₂₀ H ₃₂]	IR, MS, NMR, CT	Flexibilane*	Undetm.	Undetm.	Undetm.	<i>S. flexibilis</i>	MLU	[112, 143]
(+)-Lobophytolide 101 ^β [C ₂₀ H ₂₈ O ₃]	IR, MS, NMR, [α] _D , X-ray	Cembrane	Undetm.	Undetm.	Undetm.	<i>L. cristagalli</i>	MLU	[144]
(-)-Crassolide 102 ^β [C ₂₆ H ₃₄ O ₉]	UV, IR, MS, NMR, [α] _D , CT	Cembrane	Ichthyotoxic	<i>L. reticulatus</i>	LD ₅₀ = 7 mg/L	<i>L. crassum</i>	MLU	[145 – 148]
			Cytotoxic	A549, KB P-388	ED ₅₀ = 0.39 – 0.85 μg/mL ED ₅₀ = 0.04 – 0.08 μg/mL			
				HT-29	ED ₅₀ = 0.05 – 0.26 μg/mL			
2-Hydroxycrassocolide E 103 ^β [C ₂₀ H ₂₈ O ₄]	MS, NMR	Cembrane	Cytotoxic	MCF7	IC ₅₀ = 18.13 μg/mL	<i>Sarcophyton</i> <i>sp.</i>	NSW	[149]
(-)-2-Hydroxynepththenol 104 ^β [C ₂₀ H ₃₄ O ₂]	IR, MS, NMR, [α] _D , CT	Cembrane	Cytotoxic	KB, A549, HT-29, P-388	ED ₅₀ = 0.23 – 1.80 μg/mL	<i>L. viridis</i>	MLU	[150 – 152]
3,4,11,12-Diepoxyembrane A 105 ^β [C ₂₀ H ₃₂ O ₂]	IR, MS, NMR, CT	Cembrane	Undetm.	Undetm.	Undetm.	<i>S. flexibilis</i>	MLU	[153]
(+)-7,8-Epoxy-7,8 dihydrocembrene C 106 ^β [C ₂₀ H ₃₂ O]	MS, NMR, [α] _D	Cembrane	Cytostatic Cytotoxic	HUVEC, K562 HeLa	GI ₅₀ = 38.9 ± 1.8 – 52.5 ± 0.8 μM CC ₅₀ = 83.8 ± 0.8 μM	<i>S. ehrenbergi</i>	BLI	[154]
3,4-Epoxynephthenol acetate 107 ^β [C ₂₂ H ₃₆ O ₃]	MS, NMR	Cembrane	Cytostatic	SF268, MCF7, H-460	GI ₅₀ > 100 μM	<i>Nephthea</i> sp.	JSCR	[155]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Sinulariolide 108 ^B [C ₂₀ H ₃₀ O ₄]	IR, MS, NMR, [α] _D , CT, X-ray	Cembrane	Cytotoxic	KB, MCF7	ED ₅₀ = 7.6 – 16.9 µg/mL	<i>S. flexibilis</i>	MLU	[156 – 176]
				A549, P388, HT-29	ED ₅₀ = 3.0 – 3.9 µg/mL			
				HL-60	ED ₅₀ = 0.7 µg/mL			
				CCRF-CEM, DLD-1	IC ₅₀ > 20 µg/mL			
				Huh7, HepG2, Hep3B, HA22T	IC ₅₀ = 8.46 ± 0.05 – 16.52 ± 0.13 µg/mL			
				<i>B. neritina</i> , <i>B. albicostatus</i>	EC ₅₀ = 21 – 33.18 µg/mL			
				A549 (mitochondria path.)	25 µg/mL (nanoparticle hyaluronan/(+)- 116)			
				A375 (cytotoxic)	1 – 20 µg/mL			
				A375 (anti-migratory)	21 – 72% (5 – 15 µg/mL)			
				A375 (mitochondrial apop.)	15 µg/mL (caspase dependent)			
				TSGH (cytostatic)	15 – 30 µM			
			Anticancer	TSGH (anti-migratory)	24 – 71% (10 – 25 µM)			
				TSGH (mitochondrial apop.), RT4, T24, BFTC905	10 – 15 µM (Caspase-dependent)			
				TSGH (p38MAPK-ATF2 activation)	10 – 15 µM			
				HA22T (mitochondrial and ER apop.)	Active PERK/eIF2α/ATF4/CHOP			
				HA22T (anti-migratory)	50% (8 µg/mL, 24 h) 78% (8 µg/mL, 48 h)			
				HA22T (MMP-2/-9)	8 – 10 µg/mL			
				HA22T (MAPK – P13K/Akt)	10 µg/mL			
				HA22T (GRB2, FAK)	10 µg/mL			
				TSGH-8301 (anti-migratory)	7.5 – 10 µM			
				TSGH-8301 (MMP-2/-9)	7.5 – 10 µM			
				TSGH-8301 (mTOR)	10 µM			
				TSGH-8301 (P13K, MMP-2/-9)	5 µM			
				TSGH-8301 (GRB2, MKK7, MKK3)	10 µM			

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Sinulariolide 108 ^β [C ₂₀ H ₃₀ O ₄]	IR, MS, NMR, [α] _D , CT, X-ray	Cembrane	Toxic	<i>M. digitata</i> , <i>A. tenuis</i> .	> 5 µg/mL	<i>S. flexibilis</i>	MLU	[156 – 176]
			Ecology	Coral bleaching	Decreased 8%			
				(<i>S. flexibilis</i> , <i>L. compactum</i>)				
				Size and domination	Increasing (+)– 108			
				RAW 264.7 (LPS/iNOS, COX-2)	47.7 ± 6.3 – 52.2 ± 5.1% (10 µM)			
6ξ-Hydroxysinulariolide 109 ^β [C ₂₀ H ₃₀ O ₅]	UV, IR, MS, NMR, CT	Cembrane	Anti-inflammatory	DC (LPS, phenotypes, cytokine secretion, mixlympocyte, CD40, CD80, CD86, TNF-α, IL-6, IL-12, NO, nuclear-κB path.)	Conc. dependent manner.	<i>S. flexibilis</i>	MLU	[62, 177]
			Cardiovascular	Atrial muscle (rat)	EC ₅₀ = 83.1 ± 1 µM (ca.)			
			Antibacterial	<i>B. subtilis</i> , <i>S. aureus</i>	10 µg/mL			
			Antifeedant	<i>G. affinis</i>	1–10%			
			Algacidal	<i>C. codii</i>	4.4 mg/L			
(+)–11-Dehydrosinulariolide (5-Dehydrosinulariolide) 110 ^β [C ₂₀ H ₂₈ O ₄]	MS, NMR, [α] _D , CT, X-ray	Cembrane	Undetm.	Undetm.	Undetm.	<i>S. flexibilis</i>	MLU	[62, 177]
			Cytotoxic	KB	ED ₅₀ = 5.4 µg/mL			
				Hep2, P-388, HT-29, Daoy, A549	ED ₅₀ = 1.58 – 2.9 µg/mL			
				HeLa	IC ₅₀ = 3.04 – 3.14 µg/mL			
				CCRF-CEM, DLD-1	IC ₅₀ > 20 µg/mL			
11-Dehydroxysinulariolide 111 ^β [C ₂₀ H ₂₈ O ₅]	MS, NMR, CT	Cembrane	Undetm.	Undetm.	Undetm.	<i>S. flexibilis</i>	MLU	[62, 177]
			Cytotoxic	HL-60, HT-29	ED ₅₀ = 0.8 – 1.9 µg/mL			
				HA22T	54% via. (9 µg/mL)			
				HA22T (anti-migratory)	2.66 – 7.98 µM			
				Apop. (mitochondria, ER)	Conc. dependent manner			
(–)-11- <i>epi</i> -Sinulariolide acetate (5- <i>epi</i> -Sinulariolide) 112 ^β [C ₂₀ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D , X-ray	Cembrane	Anticancer	MMP-2, MMP-9, uPA, TIMP-1, TIMP-2	1.33 – 7.98 µM (24 h)	<i>S. flexibilis</i>	MLU	[181, 183 – 186, 188]
				P13K/AKT/mTOR	1.33 – 7.98 µM			
					(conc. dependent manner)			

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(–)-11- <i>epi</i> -Sinulariolide acetate (5- <i>epi</i> -Sinulariolide) 112^β [C ₂₀ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D , X-ray	Cembrane	Anti-inflammatory	RAW 264.7 (LPS/iNOS)	1.4 ± 1.74 – 84.89 ± 8.23% (1 – 50 μM)	<i>S. flexibilis</i>	MLU	[62, 177]
				RAW 264.7 (LPS/COX-2)	42.13 ± 3.25 – 82.69 ± 1.63% (10 – 50 μM)			–
				Female Lewis rat (AIA)	9 mg/kg every 2 days (day 7 – day 28)			181, 183
				Cathepsin K, MMP-9, TRAP, TNF-α (rat)	Attenuated protein expression			– 186, 188]
(–)-Decaryiol B 113^β [C ₂₀ H ₃₄ O ₃]	NMR, MS, [α] _D , CT	Cembrane [▲]	Cytostatic	C6, HeLa, H9c2	IC ₅₀ ≥ 200 μg/mL	<i>Lobophytum</i> sp.	NSW	[187]
(+)-Decaryiol C 114^β [C ₂₂ H ₃₆ O ₄]	NMR, MS, [α] _D , CT	Cembrane	Cytostatic	C6, HeLa, H9c2	IC ₅₀ ≥ 200 μg/mL	<i>Lobophytum</i> sp.	NSW	[187]
(–)-Decaryiol D 115^β [C ₂₀ H ₃₄ O ₄]	NMR, MS, [α] _D , CT	Cembrane	Cytostatic	C6 H9c2	IC ₅₀ = 40 ± 3 – 150 ± 15 μg/mL IC ₅₀ ≥ 200 μg/mL	<i>Lobophytum</i> sp.	NSW	[187]
(–)-Danielid 116^β [C ₂₅ H ₃₀ O ₁₀]	NMR, MS, [α] _D	Cembrane	Undetm.	Undetm.	Undetm.	<i>S. asterolobata</i>	BLI	[188]
(+)-Sarcofuranocembranolide A 117^β [C ₂₁ H ₂₄ O ₇]	UV, IR, NMR, MS, [α] _D	Cembrane [▲]	Cytostatic Anti-inflammatory	V79 RAW264.7 (LPS/TNF-α)	ED ₅₀ = 3.88 μg/mL (10 μM)	<i>Sarcophyton</i> sp.	NSW	[189]
(–)-Sarcofuranocembranolide B 118^β [C ₂₂ H ₂₈ O ₇]	UV, IR, NMR, MS, [α] _D	Cembrane	Cytostatic Anti-inflammatory	V79 RAW264.7 (LPS/TNF-α)	ED ₅₀ = 4.04 μg/mL NA (10 μM)	<i>Sarcophyton</i> sp.	NSW	[189]
(+)-Chloroscabrolide A 119^β [C ₁₉ H ₂₃ ClO ₇]	IR, NMR, MS, [α] _D , Mol. Mod.	<i>Nor</i> -cembrane [▲]	Anti-inflammatory	J774 (LPS/iNOS)	NA (10 μM)	<i>Sinularia</i> sp.	NSW	[190]
(–)-Chloroscabrolide B 120^β [C ₂₀ H ₂₇ ClO ₈]	IR, NMR, MS, [α] _D , Mol. Mod.	<i>Nor</i> -cembrane	Anti-inflammatory	J774 (LPS/iNOS)	NA (10 μM)	<i>Sinularia</i> sp.	NSW	[190]
(–)-Prescabrolide C 121^β [C ₁₉ H ₂₄ O ₅]	IR, NMR, MS, [α] _D , Mol. Mod.	<i>Nor</i> -cembrane	Anti-inflammatory	J774 (LPS/iNOS)	NA (10 μM)	<i>Sinularia</i> sp.	NSW	[190]
(+)-Xeniolide F 122^β [C ₂₀ H ₂₈ O ₄]	UV, IR, NMR, MS, [α] _D	Xenicane	Cytotoxic	P-388, A-549, HT-29, MEL-28	IC ₅₀ > 1 μg/mL	<i>Xenia</i> sp.	CSW	[191]
(+)-9-Hydroxyxeniolide F 123^β [C ₂₀ H ₂₈ O ₅]	UV, IR, NMR, MS, [α] _D	Xenicane	Cytotoxic	P-388, A-549, HT-29, MEL-28	IC ₅₀ > 1 μg/mL	<i>Xenia</i> sp.	CSW	[191]

Table S2: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Xenimanadin A 124 ^β [C ₂₃ H ₃₆ O ₅]	UV, IR, NMR, MS, [α] _D , CT	Xenicane [▲]	Cytotoxic	P-388	IC ₅₀ > 20 μg/mL	<i>Xenia</i> sp.	NSW	[192]
(+)-Xenimanadin B 125 ^β [C ₂₃ H ₃₆ O ₆]	UV, IR, NMR, MS, [α] _D	Xenicane	Cytotoxic	P-388	IC ₅₀ > 20 μg/mL	<i>Xenia</i> sp.	NSW	[192]
(+)-Xenimanadin C 126 ^β [C ₂₃ H ₃₆ O ₅]	UV, IR, NMR, MS, [α] _D	Xenicane	Cytotoxic	P-388	IC ₅₀ > 20 μg/mL	<i>Xenia</i> sp.	NSW	[192]
(+)-Xenimanadin D 127 ^β [C ₂₃ H ₃₄ O ₅]	UV, IR, NMR, MS, [α] _D	Xenicane	Cytotoxic	P-388	IC ₅₀ > 20 μg/mL	<i>Xenia</i> sp.	NSW	[192]
(+)-Niphatheolide A 128 ^β [C ₂₀ H ₃₀ O ₃]	UV, IR, NMR, MS, [α] _D , ECD	Niphathane [▲]	Anticancer	<i>E. coli</i> BL21 (DE3) cells transf. pGEX6P1-p53 or pGEX6P1-HDM2	IC ₅₀ = 16 μM	<i>N. olemda</i>	NSW	[63]

Footnote: **1. Structure** (◐molecule isolated as a natural product for the first time and found also in semisynthetic compound before). **2. Statistic** (LD₅₀ 50% of the lethal dose); **3. Activity** (C6 murine glioma, H9c2 murine cardiomyoblasts, J774 murine macrophage, L5178Y murine lymphoblastic leukemia, P-388 murine leukemia lymphoma, P388D1 macrophage-like murine lymphoma, PC12 murine adrenal gland pheochromocytoma, V79 Chinese hamster fibroblast, A375 human melanoma, CCRF-CEM human T cell acute lymphoblastic lymphoma, Daoy human desmoplastic cerebellar medulloblastoma, DLD-1 human colorectal adenocarcinoma, H-460 human lung carcinoma, HA22T human hepatocellular carcinoma, HeLa human cervical carcinoma, Hep2 human hepatocarcinoma, Huh7 human hepatocarcinoma, HT-29 human colorectal adenocarcinoma, HUVEC human umbilical vein endothelial, KB-3-1 human nasopharyngeal epidermoid carcinoma, KB-C2 human nasopharyngeal epidermoid carcinoma, KB-CV60 human nasopharyngeal epidermoid carcinoma, MEL-28 human melanoma, MKN-45 human gastric adenocarcinoma, SF268 human glioblastoma, TSGH, TSGH-8031, RT4, T24, BFTC human urinary bladder carcinoma, BACE1 beta-secretase 1, TRAP tartrate-resistant acid phosphatase, USP7 ubiquitin-specific protease 7, uPA urokinase plasminogen activator, CB cytochalasin B, CCI chronic constriction injury, FMLP formyl-L-methionyl-L-leucyl-L-phenylalanine, IFN-γ interferon gamma, LPS lipopolysaccharide, MRP-1 multidrug resistance protein 1, P-gp P-glycoprotein, RANKL receptor activator of nuclear factor kappa-B ligand, SA superoxide anion, ER endoplasmic reticulum, gen. generated, DC dendritic cell, AIA adjuvant-induced arthritis, path. pathway, via. viability, rel. release, transf. transformed).

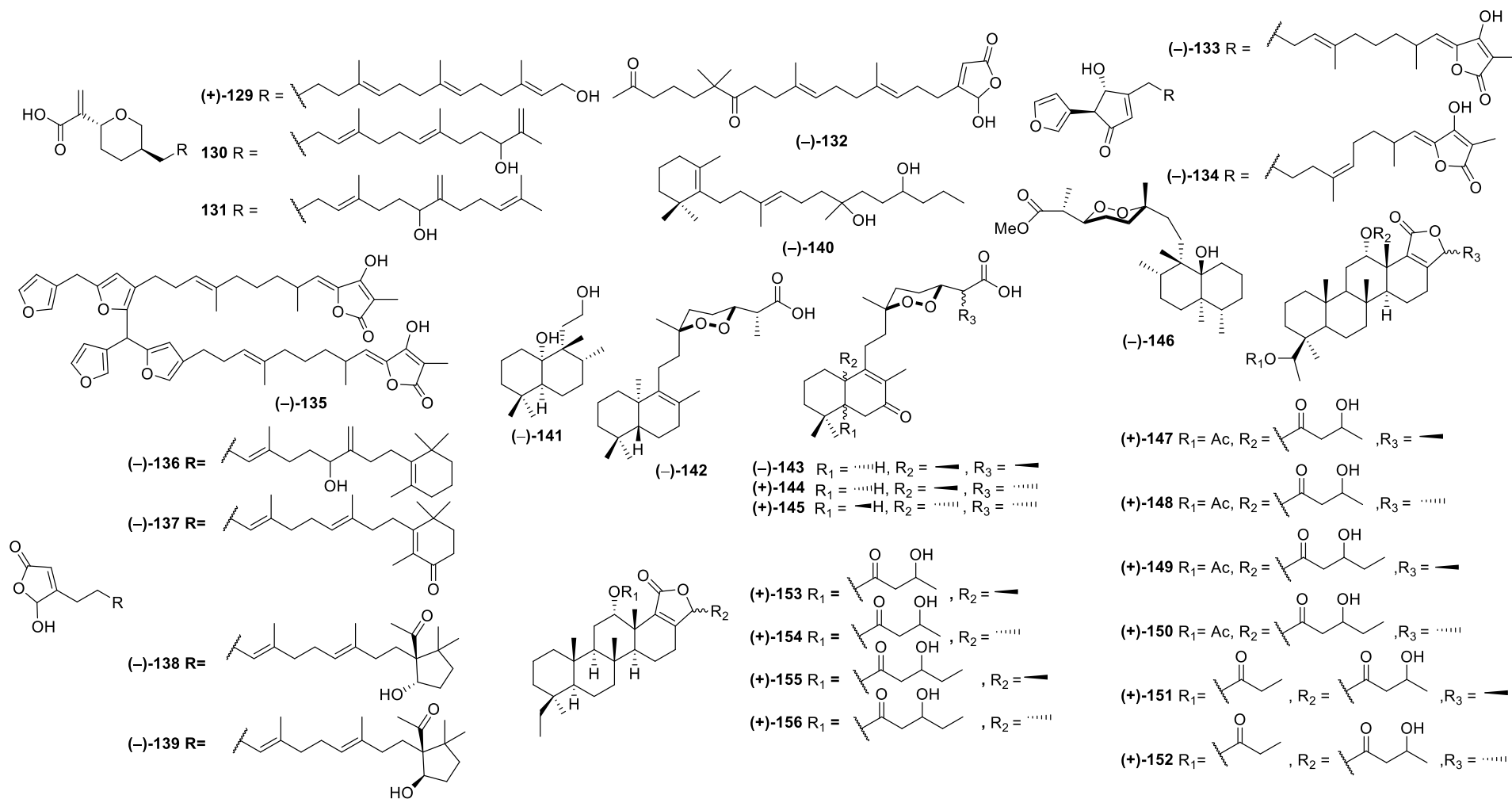


Figure S3: *Cont.*

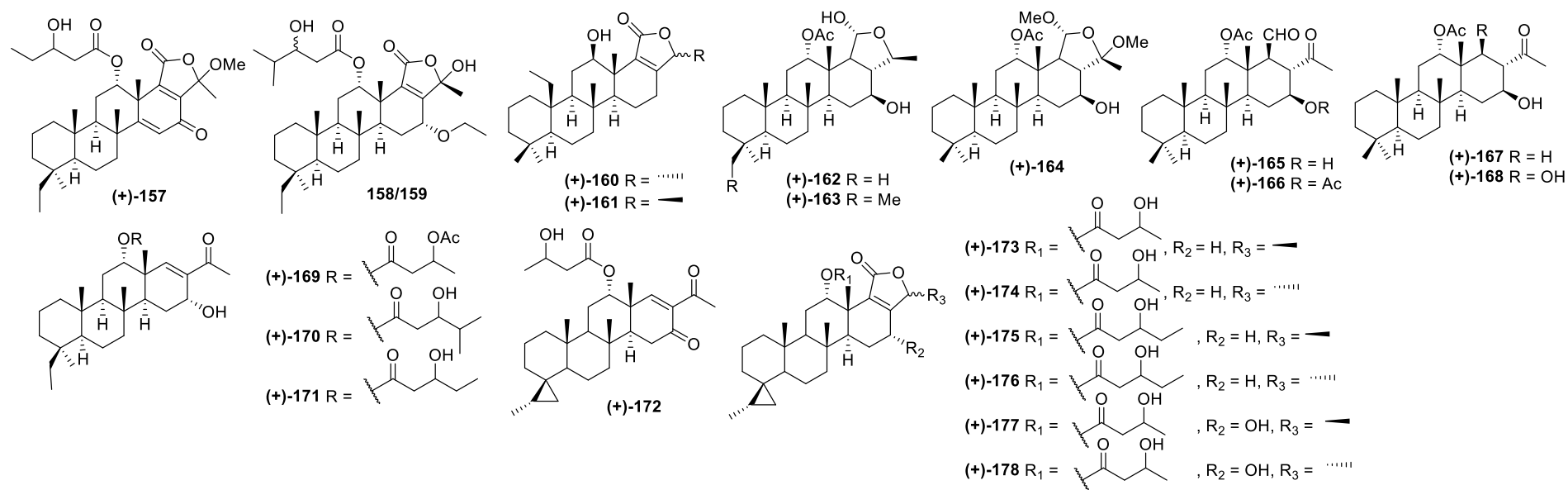


Figure S3: Structures of marine sesterterpenoids from Indonesian waters found in 1970–2017.

Table S3: Marine sesterterpenoids from Indonesian waters in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Barangcadoic acid 129 ^β [C ₂₅ H ₄₀ O ₄]	IR, MS, NMR, [α] _D	Acyclic	Cytotoxic	LoVo (mut. Ras, sens.)	IC ₅₀ ≈ 1–2 μg/mL	<i>Hippospongia</i> sp.	SSW	[193]
			Anticancer	CaCo (normal Ras, resist.) RCE	3–4-fold less activity IC ₅₀ ≈ 10 μg/mL			
Rhopaloic acid D 130 /E 131 ^{α,β} [C ₂₄ H ₃₈ O ₄]	MS, NMR	Acyclic nor-sesterterpene	Cytotoxic	LoVo (mut. Ras, sens.)	IC ₅₀ ≈ 1–2 μg/mL	<i>Hippospongia</i> sp.	SSW	[193]
			Anticancer	CaCo (normal Ras, res.) RCE	3–4-fold less activity IC ₅₀ ≈ 10 μg/mL			
(–)-Achantholide C 132 ^β [C ₂₅ H ₃₈ O ₅]	UV, MS, NMR, [α] _D	Acyclic	Undetm.	Undetm.	Undetm.	<i>Acanthodendrilla</i> sp.	SSW	[194]
(–)-Sulawesin A 133 ^{α,β} [C ₂₅ H ₃₀ O ₆]	UV, MS, NMR, [α] _D	Acyclic*	Anticancer	USP7	IC ₅₀ = 2.8 μM	<i>Psammocinia</i> sp.	NSW	[195]
(–)-Sulawesin B 134 ^{α,β} [C ₂₅ H ₃₀ O ₆]	UV, MS, NMR, [α] _D	Acyclic	Anticancer	USP7	IC ₅₀ = 4.6 μM	<i>Psammocinia</i> sp.	NSW	[195]
(–)-Sulawesin C 135 ^β [C ₅₀ H ₅₈ O ₁₀]	UV, MS, NMR, [α] _D	Acyclic	Anticancer	USP7	Undetm.	<i>Psammocinia</i> sp.	NSW	[195]
(–)-Achantholide A 136 ^β [C ₂₅ H ₃₈ O ₄]	UV, MS, NMR, [α] _D	Monocyclic	Undetm.	Undetm.	Undetm.	<i>Acanthodendrilla</i> sp.	SSW	[194]
			Cytotoxic	L5178Y <i>S. aureus</i>	ED ₅₀ > 10 μg/mL NA (5 μg) 10 mm (10 μg)			
(–)-Achantholide B 137 ^β [C ₂₅ H ₃₆ O ₄]	UV, MS, NMR, [α] _D	Monocyclic	Antibacterial	<i>B. subtilis</i>	NA (5 μg) 12 mm (10 μg)	<i>Acanthodendrilla</i> sp.	SSW	[194]
				<i>E. coli</i>	NA (5 μg) 9 mm (10 μg)			
			Antifungal	<i>C. albicans</i>	NA (5 μg) 10 mm (10 μg)			
				<i>C. herbarum</i>	NA (5 μg) 10 mm (10 μg)			
(–)-Achantholide D 138 ^β [C ₂₅ H ₃₈ O ₅]	UV, MS, NMR, [α] _D	Monocyclic	Cytotoxic	L5178Y	ED ₅₀ > 10 μg/mL	<i>Acanthodendrilla</i> sp.	SSW	[194]
(–)-Achantholide E 139 ^β [C ₂₅ H ₃₈ O ₅]	UV, MS, NMR, [α] _D	Monocyclic	Cytotoxic	L5178Y	ED ₅₀ = 7 μg/mL	<i>Acanthodendrilla</i> sp.	SSW	[194]

Table S3: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Diacardiol A 140 ^β [C ₂₄ H ₄₄ O ₂]	MS, NMR, [α] _D	Monocyclic	Cytotoxic	L5178Y HeLa, PC12 <i>A. salina</i>	EC ₅₀ = 4.80 µg/mL EC ₅₀ > 10 µg/mL 20 – 40 % (10 µg/mL, 24 – 48 h)	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Euplectellodiol 141 ^β [C ₁₆ H ₃₀ O ₂]	MS, NMR, [α] _D	Bicyclic	Undetm.	Undetm.	Undetm.	<i>M. euplec-tellioides</i>	NSW	[123]
(-)-Diacarperoxide D 142 ^β [C ₂₄ H ₄₀ O ₄]	MS, NMR, [α] _D	Bicyclic	Cytotoxic	L5178Y HeLa, PC12	EC ₅₀ < 0.10 µg/mL EC ₅₀ = 0.17 – 0.80 µg/mL	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Diacarperoxide E 143 ^β [C ₂₄ H ₃₈ O ₅]	MS, NMR, [α] _D	Bicyclic	Cytotoxic	L5178Y	EC ₅₀ < 3.0 µg/mL	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(+)-Diacarperoxide F 144 ^β [C ₂₄ H ₃₈ O ₅]	MS, NMR, [α] _D	Bicyclic	Cytotoxic	L5178Y HeLa, PC12	EC ₅₀ = 0.06 µg/mL EC ₅₀ = 0.60 – 0.80 µg/mL	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(+)-Diacarperoxide G 145 ^β [C ₂₅ H ₄₀ O ₅]	MS, NMR, [α] _D	Bicyclic	Cytotoxic	L5178Y HeLa, PC12 <i>A. salina</i>	EC ₅₀ = 2 µg/mL EC ₅₀ > 10 µg/mL 50 – 60% (10 µg/mL, 24 – 48 h)	<i>D. megaspi-norhabdosa</i>	SSW	[124]
(-)-Diacarperoxide S = (-)-Megaspinoxide A 146 ^β [C ₂₅ H ₄₄ O ₅]	IR, MS, NMR, [α] _D	Bicyclic	Antibacterial	<i>S. aureus</i> (AUMC No. B-54), <i>B. cereus</i> (AUMC No. B-52) <i>E. coli</i> (AUMC No. B-53), <i>P. aeruginosa</i> (AUMC No. B-73), <i>S. marcescens</i> (AUMC No. B-55)	29 – 35 mm (100 µg) 4 – 6 mm (100 µg)	<i>D. megaspi-norhabdosa</i>	SSW	[196, 197]
(+)-Honulactone C 147 ^β [C ₃₃ H ₅₀ O ₇]	IR, MS, NMR, [α] _D	Tetracyclic	Antifungal	<i>C. albicans</i> (AUMC No. 418)	12 – 16 mm (100 µg)			
(+)-Honulactone D 148 ^β [C ₃₃ H ₅₀ O ₇]	IR, MS, NMR, [α] _D , X-ray	Tetracyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	IC ₅₀ = 1 µg/mL	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone I 149 ^β [C ₃₄ H ₅₂ O ₇]	IR, MS, NMR, [α] _D	Tetracyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	IC ₅₀ = 1 µg/mL NA	<i>S. aliena</i>	EKM	[198]

Table S3: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Honulactone J 150 ^β [C ₃₄ H ₅₂ O ₇]	IR, MS, NMR, [α] _D	Tetracyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone K 151 ^β [C ₃₄ H ₅₂ O ₇]	IR, MS, NMR, [α] _D	Tetracyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone L 152 ^β [C ₃₄ H ₅₂ O ₇]	IR, MS, NMR, [α] _D	Tetracyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Phyllofolactone H 153 ^β [C ₃₁ H ₄₈ O ₅]	MS, NMR, [α] _D	Tetracyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
(+)-Phyllofolactone I 154 ^β [C ₃₁ H ₄₈ O ₅]	MS, NMR, [α] _D	Tetracyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
(+)-Phyllofolactone J 155 ^β [C ₃₂ H ₅₀ O ₅]	MS, NMR, [α] _D	Tetracyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
(+)-Phyllofolactone K 156 ^β [C ₃₂ H ₅₀ O ₅]	MS, NMR, [α] _D	Tetracyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
(+)-Phyllactone H 157 ^β [C ₃₃ H ₄₈ O ₇]	UV, IR, MS, NMR, [α] _D	Tetracyclic	Cytotoxic	MCF7, A549, HeLa, HL-7702	IC ₅₀ = 16 – 27 μM (ca.)	<i>P. papyracea</i>	NSW	[200]
(+)- 158 /(+)- 159 ^{α,β} [C ₃₅ H ₅₆ O ₇]	UV, MS, NMR, [α] _D	Tetracyclic	Anticancer Cytotoxic	RCE, CACO2 LoVo, PC3 MDA468	IC ₅₀ = 4.2 μg/mL IC ₅₀ = 2.9 – 3.2 μg/mL IC ₅₀ = 4.4 μg/mL	<i>C. foliascens</i>	SSW	[201]
(+)-Hyattellactone A 160 ^β [C ₂₇ H ₄₂ O ₃]	UV, IR, MS, NMR, ECD, [α] _D , Mol. Mod.	Tetracyclic	Antidiabetic Cytotoxic	PTP1B Jurkat	IC ₅₀ = 7.45 μM NA (24.2 μM)	<i>Hyattella</i> sp.	NSW	[202]
(+)-Hyattellactone B 161 ^β [C ₂₇ H ₄₂ O ₃]	UV, IR, MS, NMR, ECD, [α] _D	Tetracyclic	Antidiabetic Cytotoxic	PTP1B Jurkat	42% (24.2 μM) NA (24.2 μM)	<i>Hyattella</i> sp.	NSW	[202]
(+)- 162 ^β [C ₂₈ H ₄₆ O ₅]	MS, NMR, [α] _D	Tetracyclic	Cytotoxic	KB	30–95% (10 μg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)- 163 ^β [C ₂₉ H ₄₈ O ₅]	MS, NMR, [α] _D	Tetracyclic	Cytotoxic	KB	30–95% (10 μg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)- 164 ^β [C ₃₀ H ₅₀ O ₆]	MS, NMR, [α] _D	Tetracyclic	Cytotoxic	KB	30–95% (10 μg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)- 165 ^β [C ₂₈ H ₄₄ O ₅]	MS, NMR, [α] _D , X-ray, CT	Tetracyclic	Cytotoxic	KB	30–95% (10 μg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]

Table S3: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)- 166 ^β [C ₃₀ H ₄₆ O ₆]	MS, NMR, [α] _D , CT	Tetracyclic	Cytotoxic	KB	30–95% (10 µg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)- 167 ^β [C ₂₇ H ₄₄ O ₄]	MS, NMR, [α] _D	Tetracyclic	Cytotoxic	KB	30–95% (10 µg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)- 168 ^β [C ₂₇ H ₄₄ O ₅]	MS, NMR, [α] _D	Tetracyclic	Cytotoxic	KB	30–95% (10 µg/mL)	<i>Phyllospongia</i> sp.	SSW	[203]
(+)-Phyllofenone C 169 ^β [C ₃₂ H ₅₀ O ₆]	MS, NMR, [α] _D	Tetracyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
170 ^β [C ₃₂ H ₅₂ O ₅]	UV, MS, NMR, [α] _D	Tetracyclic	Anticancer Cytostatic	RCE LoVo CACO2, MDA468, PC3	IC ₅₀ = 38 µg/mL IC ₅₀ = 7.6 µg/mL IC ₅₀ = 3.4 – 3.8 µg/mL	<i>C. foliascens</i>	SSW	[201]
171 ^β [C ₃₁ H ₅₀ O ₅]	UV, MS, NMR, [α] _D	Tetracyclic	Anticancer Cytostatic	RCE PC3, LoVo, CACO2	IC ₅₀ > 100 µg/mL IC ₅₀ > 10 µg/mL	<i>C. foliascens</i>	SSW	[201]
(+)-Honu'enone 172 ^β [C ₃₀ H ₄₄ O ₅]	MS, NMR, [α] _D	Pentacyclic	Undetm.	Undetm.	Undetm.	<i>S. aliena</i>	EKM	[199]
(+)-Honulactone A 173 ^β [C ₃₁ H ₄₆ O ₅]	IR, MS, NMR, [α] _D	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	IC ₅₀ = 1 µg/mL	<i>S. aliena</i>	EKM	[199]
(+)-Honulactone B 174 ^β [C ₃₁ H ₄₆ O ₅]	IR, MS, NMR, [α] _D , X-ray	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	IC ₅₀ = 1 µg/mL	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone E 175 ^β [C ₃₂ H ₄₈ O ₅]	IR, MS, NMR, [α] _D	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone F 176 ^β [C ₃₂ H ₄₈ O ₅]	IR, MS, NMR, [α] _D	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone G 177 ^β [C ₃₁ H ₄₆ O ₆]	IR, MS, NMR, [α] _D	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]
(+)-Honulactone H 178 ^β [C ₃₁ H ₄₆ O ₆]	IR, MS, NMR, [α] _D	Pentacyclic	Cytotoxic	P-388 (ATCC: CCL 46), HT-29 (ATCC: CCL 8), MEL-28 (ATCC: HTB 72)	NA	<i>S. aliena</i>	EKM	[198]

Footnote: 1. **Activity** (CaCo human colorectal adenocarcinoma, CACO2 human colon carcinoma, HL-7702 human hepatocarcinoma cell, Jurkat human T lymphoma, MDA468 human breast adenocarcinoma, PC3 human prostate carcinoma, PTP1B protein tyrosine phosphatase 1B, RCE ras converting enzyme, migr. migration, mut. mutated, resist. resistant).

Table S4: Marine triterpenoids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)- 179 ^β [C ₃₀ H ₅₀ O ₂]	MS, NMR, [α] _D	Squalene	Undetm.	Undetm.	Undetm.	<i>H. erectus</i>	SSW	[204, 205]
(-)-Globostellatic acid F 180 ^β [C ₃₀ H ₄₂ O ₆]	MS, NMR, [α] _D	Isomala-baricane	Cytotoxic	L5178Y HeLa, PC12	42 – 100% (3 – 10 µg/mL) ED ₅₀ = 10.36 nmol NA (3 – 10 µg/mL)	<i>R. globostellata</i>	SSW	[206]
(+)-Globostelletin 181 ^β [C ₃₀ H ₄₄ O ₄]	MS, NMR, [α] _D	Isomala-baricane	Cytotoxic	L5178Y HeLa, PC12 HeLa L5178Y	66 – 94% (3 – 10 µg/mL) ED ₅₀ = 5.34 nmol NA (3 – 10 µg/mL) ED ₅₀ > 60 nmol 100% (3 – 10 µg/mL) ED ₅₀ = 0.39 nmol	<i>R. globostellata</i>	SSW	[206]
Globostellatic acid G 182 ^{α,β} [C ₃₁ H ₄₆ O ₆]	UV, MS, NMR	Isomala-baricane	Cytotoxic	HeLa PC12 L5178Y	8 – 25% (3 – 10 µg/mL) ED ₅₀ = 46.69 nmol 18 – 42% (3 – 10 µg/mL) ED ₅₀ = 30.33 nmol 100% (3 – 10 µg/mL) ED ₅₀ = 0.31 nmol	<i>R. globostellata</i>	SSW	[206]
Globostellatic acid I 183 ^{α,β} [C ₃₂ H ₄₈ O ₆]	UV, MS, NMR	Isomala-baricane	Cytotoxic	HeLa PC12	8 – 23% (3 – 10 µg/mL) ED ₅₀ = 46.05 nmol 16 – 41% (3 – 10 µg/mL) ED ₅₀ = 28.07 nmol	<i>R. globostellata</i>	SSW	[206]
Globostellatic acid K 184 ^{α,β} [C ₃₄ H ₅₀ O ₇]	UV, MS, NMR	Isomala-baricane	Cytotoxic	L5178Y HeLa, PC12 L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 8.28 nmol NA (3 – 10 µg/mL) 100% (3 – 10 µg/mL) ED ₅₀ = 0.92 nmol	<i>R. globostellata</i>	SSW	[206]
Globostellatic acid M 185 ^{α,β} [C ₃₀ H ₄₄ O ₆]	UV, MS, NMR	Isomala-baricane	Cytotoxic	HeLa PC12	42% (3 – 10 µg/mL) ED ₅₀ = 27.87 nmol 0 – 30% (3 – 10 µg/mL) ED ₅₀ = 27.52 nmol	<i>R. globostellata</i>	SSW	[206]

Table S4: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(–)-3- <i>O</i> -Deacetyl-13 <i>Z</i> -stelliferin riboside 186 ^β [C ₃₅ H ₅₄ O ₇]	UV, MS, NMR, [α] _D	Isomala-baricane	Cytotoxic	L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 2.40 nmol	<i>R. globostellata</i>	SSW	[206]
				HeLa	26 – 100% (3 – 10 µg/mL) ED ₅₀ = 8.14 nmol			
				PC12	33 – 54% (3 – 10 µg/mL) ED ₅₀ = 27.63 nmol			
(+)-13 <i>Z</i> , 17 <i>E</i> -Globostellatic acid X methyl ester 187 ^β [C ₃₃ H ₄₆ O ₅]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	HUVEC	IC ₅₀ = 0.064 µM	<i>R. globostellata</i>	UEP	[207]
(–)-13 <i>Z</i> , 17 <i>Z</i> -Globostellatic acid X methyl ester 188 ^β [C ₃₃ H ₄₆ O ₅]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	KB-3-1, Neuro2A	IC ₅₀ = 3.1 – 3.2 µM			
				K562	IC ₅₀ = 9 µM			
(–)-Acetyljaspiferol E 189 ^β [C ₂₄ H ₃₂ O ₆]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	HUVEC	IC ₅₀ = 0.4 µM	<i>R. globostellata</i>	UEP	[207]
Globostellatic acid H 190 ^{α,β} [C ₃₂ H ₄₈ O ₆]	UV, MS, NMR	Isomala-baricane	Cytotoxic	KB-3-1, K562, Neuro2A	IC ₅₀ = 7.9 – 22 µM			
				HUVEC	IC ₅₀ = 2.2 µM	<i>R. globostellata</i>	SSW	[206]
				KB-3-1, K562, Neuro2A	IC ₅₀ = 28–34 µM			
Globostellatic acid J 191 ^{α,β} [C ₃₄ H ₅₀ O ₇]	UV, MS, NMR	Isomala-baricane	Cytotoxic	L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 0.31 nmol	<i>R. globostellata</i>	SSW	[206]
				HeLa	8 – 23% (3 – 10 µg/mL) ED ₅₀ = 46.05 nmol			
				PC12	41% (10 µg/mL)			
Globostellatic acid L 192 ^{α,β} [C ₃₀ H ₄₄ O ₆]	UV, MS, NMR	Isomala-baricane	Cytotoxic	L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 8.28 nmol	<i>R. globostellata</i>	SSW	[206]
				HeLa, PC12	NA (3 – 10 µg/mL)			
				L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 0.92 nmol			
(+)–13 <i>E</i> -Stelliferin riboside 193 ^β [C ₃₇ H ₅₆ O ₈]	UV, NMR, MS, [α] _D	Isomala-baricane	Cytotoxic	HeLa	0 – 42% (3 – 10 µg/mL) ED ₅₀ = 27.87 nmol	<i>R. globostellata</i>	SSW	[206]
				PC12	30% (10 µg/mL)			
				L5178Y	100% (3 – 10 µg/mL) ED ₅₀ = 0.22 nmol			
				HeLa	0 – 56% (3 – 10 µg/mL) ED ₅₀ = 22.76 nmol			
				PC12	0 – 38% (10 µg/mL) ED ₅₀ = 21.54 nmol			

Table S4: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-13E, 17Z-Globostellatic acid X methyl ester 194 ^β [C ₃₃ H ₄₆ O ₅]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	HUVEC KB-3-1, K562 Neuro2A	IC ₅₀ = 0.06 μM IC ₅₀ = 18 – 22 μM IC ₅₀ = 4.7 μM	<i>R. globostellata</i>	UEP	[207]
(-)-13E, 17E-Globostellatic acid X methyl ester 195 ^β [C ₃₃ H ₄₆ O ₅]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic Anti-migratory Anticancer	HUVEC KB-3-1, K562 Neuro2A HUVEC	IC ₅₀ = 0.09 μM IC ₅₀ = 14 – 23 μM IC ₅₀ = 7.5 μM 1 μM	<i>R. globostellata</i>	UEP	[207]
(-)-Globostellatic acid F methyl ester 196 ^β [C ₃₃ H ₄₈ O ₇]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	HUVEC (apop. caspase 3/7) HUVEC KB-3-1, K562 Neuro2A	1 – 10 μM (48 h) IC ₅₀ = 0.98 μM IC ₅₀ = 8.6 – 12 μM IC ₅₀ = 5.0 μM	<i>R. globostellata</i>	UEP	[207]
(-)-13E-Globostellatic acid B methyl ester 197 ^β [C ₃₄ H ₅₀ O ₇]	MS, NMR, [α] _D	Isomala-baricane	Cytostatic	HUVEC KB-3-1, K562 Neuro2A	IC ₅₀ = 1.1 μM IC ₅₀ = 6.5 – 9.3 μM IC ₅₀ = 6.0 μM	<i>R. globostellata</i>	UEP	[207]
(-)-Vannusal A 198a ^β [C ₃₄ H ₄₈ O ₈]	UV, MS, NMR, ECD, [α] _D , Mol. Mod., CT	Vannusane ⁺	Undetm.	Undetm.	Undetm.	<i>E. vannus</i>	NSW	[36 – 43]
(-)-Vannusal A 198b ^γ [C ₃₄ H ₄₈ O ₈]	CT	Vannusane	Undetm.	Undetm.	Undetm.	<i>E. vannus</i>	NSW	[36 – 43]
(-)-Vannusal B 199a ^β [C ₃₂ H ₄₆ O ₇]	UV, MS, NMR, ECD, [α] _D , Mol. Mod., CT	Vannusane	Undetm.	Undetm.	Undetm.	<i>E. vannus</i>	NSW	[36 – 43]
(-)-Vannusal B 199b ^γ [C ₃₂ H ₄₆ O ₇]	UV, MS, NMR, [α] _D , X-ray, TS	Vannusane	Undetm.	Undetm.	Undetm.	<i>E. vannus</i>	NSW	[36 – 43]
(+)-Plakohopanoid peracetate 200c ^ε [C ₃₇ H ₅₆ O ₈]	MS, NMR, [α] _D , CT, ECD	Hopane	Undetm.	Undetm.	Undetm.	<i>P. cf lita</i>	NSW	[208]

Footnote: 1. Activity (Neuro 2A murine neuroblastoma).

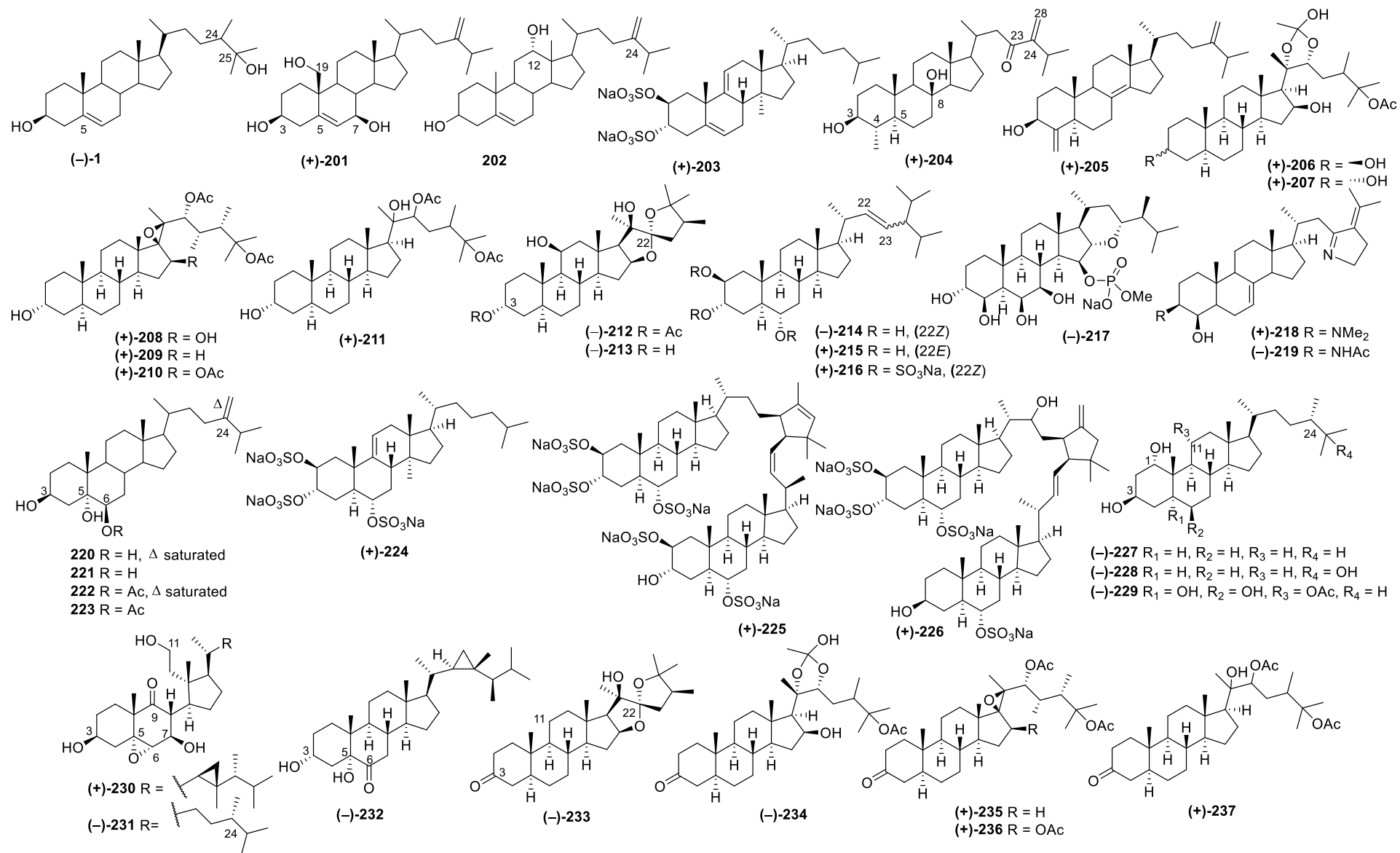


Figure S5: *Cont.*

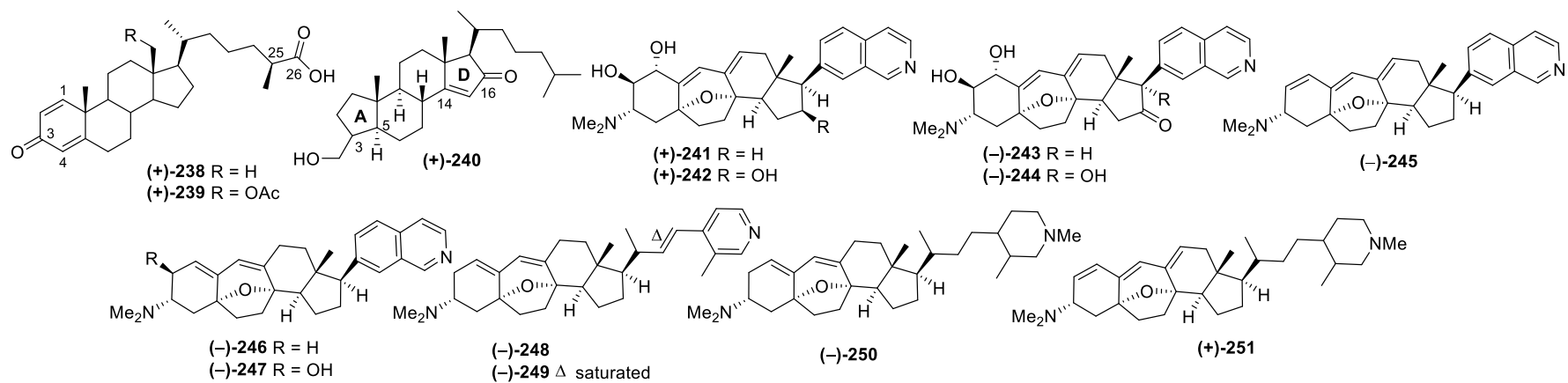


Figure S5: Structures of marine steroids from Indonesian waters found in 1970–2017.

Table S5: Marine steroids from Indonesian Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-25-Hydroxy-24 ξ -methylcholesterol 1 ^{β} [C ₂₈ H ₄₈ O ₂]	IR, MS, NMR, [α] _D , CT	Δ^5 Sterol	Undetm.	Undetm.	Undetm.	<i>Nephtea sp.</i>	NST	[28]
(+)-24-Methylenecholest-5-en-3 β ,7 β ,19-triol 201 ^{β} [C ₂₈ H ₄₆ O ₃]	IR, MS, NMR, [α] _D , X-ray, CT	Δ^5 Sterol	Undetm.	Undetm.	Undetm.	<i>L. viridis</i>	MLU	[209]
12 α -hydroxy-24-methylene cholesterol 202 ^{β} [C ₂₈ H ₄₆ O ₂]	MS, NMR, [α] _D	Δ^5 Sterol	Undetm.	Undetm.	Undetm.	<i>L. viridis</i>	MLU	[62]
(+)-Lembehsterol B 203 ^{β} [C ₂₈ H ₄₄ Na ₂ O ₈ S ₂]	IR, MS, NMR, [α] _D	Δ^5 Sterol	Anticancer	TP	IC ₅₀ = 45 μ M	<i>P. strongylata</i>	NSW	[210]
(+)-4 α -Methyl-3 β , 8 β -dihydroxy-5 α -ergost-24(28)-en-23-one 204 ^{β} [C ₂₉ H ₄₈ O ₃]	UV, IR, MS, NMR, [α] _D X-ray	Δ^5 Sterol	Undetm.	Undetm.	Undetm.	<i>L. viridis</i>	UEP	[211]
(+)-Dehydroconicasterol 205 ^{β} [C ₂₉ H ₄₆ O]	UV, MS, NMR, [α] _D	Δ^5 Sterol	Cytotoxic	C6, HeLa, H9c2	IC ₅₀ > 70 μ M	<i>T. swinhoei</i>	NSW	[212]
(+)-Orthohippurinsterol A 206 ^{β} [C ₃₂ H ₅₄ O ₇]	MS, NMR, [α] _D	Δ^5 Sterol*	Cytotoxic	P-388 A549, HT-29, MEL-28	IC ₅₀ = 2.5 μ g/mL IC ₅₀ = 5 μ g/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Orthohippurinsterol B 207 ^{β} [C ₃₂ H ₅₄ O ₇]	MS, NMR, [α] _D	Δ^5 Sterol	Cytotoxic	P-388, MEL-28 A549 HT-29	IC ₅₀ > 10 μ g/mL IC ₅₀ = 5 μ g/mL IC ₅₀ = 1 μ g/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerol A 208 ^{β} [C ₃₃ H ₅₄ O ₇]	MS, NMR, [α] _D	Δ^5 Sterol	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 1 μ g/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerol B 209 ^{β} [C ₃₃ H ₅₄ O ₆]	MS, NMR, [α] _D	Δ^5 Sterol	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 1.25 μ g/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerol C 210 ^{β} [C ₃₃ H ₅₆ O ₈]	MS, NMR, [α] _D	Δ^5 Sterol	Undetm.	Undetm.	Undetm.	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerol D 211 ^{β} [C ₃₂ H ₅₄ O ₆]	MS, NMR, [α] _D	Δ^5 Sterol	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 1 μ g/mL	<i>I. hippuris</i>	UEP	[213]

Table S5: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-3-Acetyl-22- <i>epi</i> -hippuristanol 212 ^β [C ₃₀ H ₄₈ O ₆]	MS, NMR, [α] _D	Δ ⁵ Sterol	Cytotoxic	P-388 A549, MEL-28 HT-29	IC ₅₀ = 1 μg/mL IC ₅₀ = 0.125 μg/mL IC ₅₀ = 0.5 μg/mL	<i>I. hippuris</i>	UEP	[213]
(-)-11-Dehydroxy-22- <i>epi</i> -hippuristanol 213 ^β [C ₂₈ H ₄₆ O ₄]	MS, NMR, [α] _D	Δ ⁵ Sterol	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 5 μg/mL	<i>I. hippuris</i>	UEP	[213]
(-)-Topsentinol K 214 ^β [C ₃₀ H ₅₂ O ₃]	UV, IR, MS, NMR, [α] _D	Δ ⁵ Sterol	Alzheimer	BACE1	NA	<i>Topsentia</i> sp.	EKM	[214]
(+)-Topsentinol L 215 ^β [C ₃₀ H ₅₂ O ₃]	UV, IR, MS, NMR, [α] _D	Δ ⁵ Sterol	Alzheimer	BACE1	NA	<i>Topsentia</i> sp.	EKM	[214]
(+)-Topsentinol K trisulfate 216 ^β [C ₃₀ H ₄₉ Na ₃ O ₁₂ S ₃]	UV, IR, MS, NMR, [α] _D	Δ ⁵ Sterol	Alzheimer	BACE1	IC ₅₀ = 1.2 μM	<i>Topsentia</i> sp.	EKM	[214]
(-)-Desulfohaplosamate 217 ^β [C ₃₀ H ₅₀ NaO ₉ P]	MS, NMR, [α] _D	Δ ⁵ Sterol ^Δ	Physicoactive (CB receptor ligand)	HEK-293 transf. CB ₁ and CB ₂	K _i hCB ₁ = 19.47 μM K _i hCB ₂ = 2.82 μM	<i>Dasychalina</i> sp.	NSW	[215]
(+) -Lokysterolamine A 218 ^β [C ₃₁ H ₅₀ N ₂ O]	MS, NMR, [α] _D	Δ ⁵ Oxidized sterol	Cytotoxic	P-388, A549 HT-29 MEL-28 MLR LcV	IC ₅₀ = 0.5 μg/mL IC ₅₀ = 1.0 μg/mL IC ₅₀ = 5.0 μg/mL 0.13 mm (50 μg/disk) >25.0 mm (50 μg/disk)	<i>Corticium</i> sp.	NSW	[216]
			Antibacterial Antifungal	<i>B. subtilis</i> <i>C. albicans</i>	19 mm (50 μg/disk) 11 mm (50 μg/disk)			
				P-388, HT-29 A549 MEL-28 MLR LcV	IC ₅₀ = 1.0 μg/mL IC ₅₀ = 0.5 μg/mL IC ₅₀ > 2 μg/mL 0.48 mm (50 μg/disk) >12.5 mm (50 μg/disk)			
(-) -Lokysterolamine B 219 ^β [C ₃₁ H ₄₈ N ₂ O ₂]	MS, NMR, [α] _D	Δ ⁵ Oxidized sterol	Cytotoxic	MEL-28 MLR LcV	IC ₅₀ > 2 μg/mL 0.48 mm (50 μg/disk) >12.5 mm (50 μg/disk)	<i>Corticium</i> sp.	NSW	[216]
			Antibacterial Antifungal	<i>B. subtilis</i> <i>C. albicans</i>	8 mm (50 μg/disk) 0 mm (50 μg/disk)			
24ξ-Methylcholestane-3β,5α,6β-triol 220 [C ₂₈ H ₅₀ O ₃]/ 221 ^{α,β} [C ₂₈ H ₄₈ O ₃]	IR, MS, NMR, CT	Δ ⁵ Oxidized sterol	Undetm.	Undetm.	Undetm.	<i>S. dissecta</i>	MLU	[217]

Table S5: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
24ξ-Methylcholestane-3β,5α,6β-triol-6-monoacetate 222 [C ₃₀ H ₅₂ O ₄]/ 223 ^{α,β} [C ₃₀ H ₅₀ O ₄]	IR, MS, NMR, CT	Δ ⁵ Oxidized steroid	Undetm.	Undetm.	Undetm.	<i>S. dissecta</i>	MLU	[217]
(+)-Lembesterol A 224 ^β [C ₂₈ H ₄₅ Na ₃ O ₁₂ S ₃]	IR, MS, NMR, [α] _D , CT	Δ ⁵ Oxidized steroid	Anticancer	TP	IC ₅₀ = 41 μM	<i>P. strongylata</i>	NSW	[210]
(+)-Manadosterol A 225 ^β [C ₅₄ H ₈₃ Na ₅ O ₂₁ S ₅]	IR, MS, NMR, [α] _D	Δ ⁵ Oxidized steroid [▲]	Anticancer	Ubc13-Uev1A	IC ₅₀ = 0.09 μM	<i>L. fibrosa</i>	NSW	[218]
(+)-Manadosterol B 226 ^β [C ₅₄ H ₈₄ Na ₄ O ₁₈ S ₄]	IR, MS, NMR, [α] _D	Δ ⁵ Oxidized steroid	Anticancer	Ubc13-Uev1A	IC ₅₀ = 0.13 μM	<i>L. fibrosa</i>	NSW	[218]
(-)-24S-Methylcholestan-1α, 3β-diol 227 ^β [C ₂₈ H ₅₀ O ₂]	MS, NMR, [α] _D	Δ ⁵ Oxidized steroid	Anti-inflammatory	HepG2 transf. pCMV-FXR, pSG5-RXR, p(hsp27)TKLUC, pCMV-β-gal	NA as antagonized FXR (10 μM 234 + 50 μM CDCA)	<i>Sinularia</i> sp.	NSW	[219]
(-)-24S-Methylcholestan-1α, 3β, 25-triol 228 ^β [C ₂₈ H ₅₀ O ₃]	MS, NMR, [α] _D	Δ ⁵ Oxidized steroid	Anti-inflammatory	HepG2 transf. pCMV-FXR, pSG5-RXR, p(hsp27)TKLUC, pCMV-β-gal	NA as antagonized FXR (10 μM 235 + 50 μM CDCA)	<i>Sinularia</i> sp.	NSW	[219]
(-)-24S-Methylcholestan-11-acetoxy-1α, 3β, 5α, 6β-tetraol 229 ^β [C ₃₀ H ₅₂ O ₆]	MS, NMR, [α] _D	Δ ⁵ Oxidized steroid	Anti-inflammatory	HepG2 transf. with pCMV-FXR, pSG5-RXR, p(hsp27)TKLUC, pCMV-β-gal	NA as antagonized FXR (10 μM 236 + 50 μM CDCA)	<i>Sinularia</i> sp.	NSW	[219]
(+)-3β,7β,11-Trihydroxy-5α,6α-epoxy-9,11-secogorgostan-9-one 230 ^β [C ₃₀ H ₅₀ O ₅]	IR, MS, NMR, [α] _D	Δ ⁵ Oxidized steroid [▲]	Cytotoxic	A2780 K562	IC ₅₀ = 6.3 μM IC ₅₀ = 7.1 μM	<i>Lobophytum</i> sp.	NMU	[220]
(-)-24S*,3β,11-Dihydroxy-5β,6β-epoxy-24-methyl-9,11-secocholestan-9-one 231 ^β [C ₂₈ H ₄₈ O ₄]	IR, MS, NMR, [α] _D	Δ ⁵ Oxidized steroid [▲]	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ > 1 μg/mL	<i>P. violacea</i>	CSW	[221]
(-)-3α,5β-Dihydroxy-gorgostan-6-one 232 ^β [C ₃₀ H ₅₀ O ₃]	MS, NMR, [α] _D	Δ ⁵ Oxidized steroid	Undetm.	Undetm.	Undetm.	<i>Sinularia</i> sp.	NSW	[219]
(-)-11-Dehydroxy-22- <i>epi</i> -hippuristan-3-one 233 ^β [C ₂₈ H ₄₄ O ₄]	MS, NMR, [α] _D	C(3) ketone steroid [▲]	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 5 μg/mL	<i>I. hippuris</i>	UEP	[213]

Table S5: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Orthohippuristerone A 234 ^β [C ₃₂ H ₅₂ O ₇]	MS, NMR, [α] _D	C(3) ketone steroid	Cytotoxic	P-388 A549, HT-29, MEL-28	IC ₅₀ = 2.5 µg/mL IC ₅₀ = 5 µg/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerone B 235 ^β [C ₃₃ H ₅₂ O ₆]	MS, NMR, [α] _D	C(3) ketone steroid	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ > 10 µg/mL	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerone C 236 ^β [C ₃₅ H ₅₄ O ₈]	MS, NMR, [α] _D	C(3) ketone steroid	Undetm.	Undetm.	Undetm.	<i>I. hippuris</i>	UEP	[213]
(+)-Hippuristerone D 237 ^β [C ₃₂ H ₅₂ O ₆]	MS, NMR, [α] _D	C(3) ketone steroid	Cytotoxic	P-388, A549, HT-29, MEL-28	IC ₅₀ = 1 µg/mL	<i>I. hippuris</i>	UEP	[213]
(+) -25S-3-Oxocholesta-1,4-dien-26-oic acid 238 ^β [C ₂₇ H ₄₀ O ₃]	MS, NMR, [α] _D	C(3) ketone steroid	Antibacterial	<i>S. aureus</i> IAM 12544T, <i>E. coli</i> IAM 12119T	NA (100 µg/disk)	<i>Minabea</i> sp.	NSW	[222]
			Antifungal	<i>S. cerevisiae</i> IAM 14383T, <i>M. hiemalis</i> IAM 6088	NA (100 µg/disk)			
			Cytostatic	V79	NA (10 µM)			
			Cytotoxic	L1210	NA (50 µg/mL)			
(+) -25S-18-Acetoxy-3-oxocholesta-1,4-dien-26-oic acid 239 ^β [C ₂₉ H ₄₂ O ₅]	MS, NMR, [α] _D , CT	C(3) ketone steroid	Antibacterial	<i>S. aureus</i> IAM 12544T, <i>E. coli</i> IAM 12119T	NA (100 µg/disk)	<i>Minabea</i> sp.	NSW	[222]
			Antifungal	<i>S. cerevisiae</i> IAM 14383T, <i>M. hiemalis</i> IAM 6088	NA (100 µg/disk)			
			Cytostatic	V79	NA (10 µM)			
(+) -3-(Hydroxymethyl)-A-nor-5α-cholest-14-en-16-one 240 ^β [C ₂₇ H ₄₄ O ₂]	MS, NMR, [α] _D	Degraded/contracted steroid	Undetm.	Undetm.	Undetm.	<i>A. carteri</i>	EKM	[223]
(+) -Cortistatin A 241 ^β [C ₃₀ H ₃₆ N ₂ O ₃]	UV, MS, NMR, ECD, [α] _D , X-ray	Degraded/contracted steroid*	Cytostatic	HUVEC	IC ₅₀ = 0.0018 µM GI ₅₀ = 0.002 ± 0.001 µM	<i>C. simplex</i>	ENT	[64 – 72, 224]
				KB-3-1, K562, Neuro2A, NHDF	IC ₅₀ = 6.0 – 7.0 µM			
				MCF7	GI ₅₀ > 10 µM			
			Antiangiogenic	SF268, IA9, PTX22, A8, NCI-H460 bFGF/VEGF-induced HUVEC migr. and tube form.	GI ₅₀ = 4.429 ± 0.664 – 7.786 ± 0.001 µM 2 nM			

Table S5: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -Cortistatin B 242 ^β [C ₃₀ H ₃₆ N ₂ O ₄]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 1.1 μM	<i>C. simplex</i>	ENT	[64
				KB-3-1, K562, Neuro2A	IC ₅₀ = 120 – 200 μM			–
				NHDF	IC ₅₀ > 300 μM			72, 224]
(–) -Cortistatin C 243 ^β [C ₃₀ H ₃₄ N ₂ O ₄]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.019 μM	<i>C. simplex</i>	ENT	[64
				KB-3-1, Neuro2A	IC ₅₀ = 150 – 180 μM			–
				K562, NHDF	IC ₅₀ > 300 μM			72, 224]
(–) -Cortistatin D 244 ^β [C ₃₀ H ₃₄ N ₂ O ₅]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.15 μM	<i>C. simplex</i>	ENT	[64
				KB-3-1	IC ₅₀ = 55 μM			–
				K562, Neuro2A, NHDF	IC ₅₀ > 300 μM			72, 224]
(–) -Cortistatin J 245 ^β [C ₃₀ H ₃₄ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 8 nM	<i>C. simplex</i>	ENT	[64
				MCF7, IA9	GI ₅₀ = 0.070 ± 0.013 μM			–
				SF268, NCI-H460, PTX22, A8	GI ₅₀ = 27.439 – 43.22 μM			72, 224]
(–) -Cortistatin K 246 ^β [C ₃₀ H ₃₆ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic		GI ₅₀ = 4.340 ± 0.161 – 8.538 ± 0.588 μM	<i>C. simplex</i>	ENT	[64
				HUVEC	IC ₅₀ = 40 nM			–
								72, 224]
(–) -Cortistatin L 247 ^β [C ₃₀ H ₃₆ N ₂ O ₂]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 23 nM	<i>C. simplex</i>	ENT	[64
								–
								72, 224]
(–) -Cortistatin G 248 ^β [C ₃₁ H ₄₂ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.35 – 1.9 μM	<i>C. simplex</i>	ENT	[64
								–
								72, 224]

Table S5: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Cortistatin H 249 ^β [C ₃₁ H ₄₄ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.35 – 1.9 μM	<i>C. simplex</i>	ENT	[64 – 72, 224]
(-)-Cortistatin E 250 ^β [C ₃₂ H ₅₂ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.35 – 1.9 μM	<i>C. simplex</i>	ENT	[64 – 72, 224]
(+)-Cortistatin F 251 ^β [C ₃₀ H ₅₂ N ₂ O]	UV, MS, NMR, [α] _D	Degraded/contracted steroid	Cytostatic	HUVEC	IC ₅₀ = 0.35 – 1.9 μM	<i>C. simplex</i>	ENT	[64 – 72, 224]

Footnote: 1. **Statistic** (*K_i* inhibition constant); 2. **Activity** (**L1210** murine lymphocytic leukemia, **MLR** murine mixed lymphocyte reaction, **A8** epothilone-resistant human ovarian carcinoma, **HEK-293** human embryonic kidney, **LcV** human leukocytoclastic vasculities, **NCI-H460** human nonsmall cell lung cancer, **NHDF** normal human dermal fibroblast, **PTX22** paclitaxel-resistant human ovarian carcinoma, **CB₁** cannabinoid receptor type 1, **CB₂** Cannabinoid receptor type 2, **FXR** farnesoid X receptor, **RXR** retinoid X receptor, **Ubc13** E2 ubiquitin-conjugating protein ubiquitin-conjugating enzyme complex, **bFGF** basic fibroblast growth factor, **TP** Thymidine phosphorylase, **CDCA** chenodeoxycholic acid, **VEGF** Vascular endothelial growth factor).

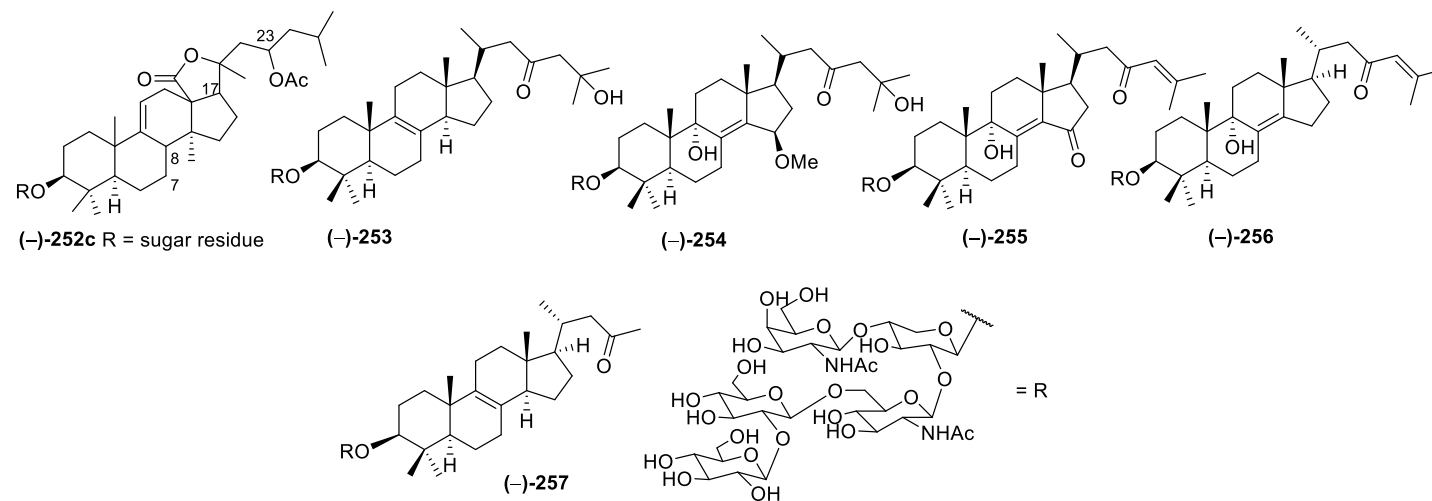


Figure S6: Structures of marine saponins from Indonesian waters found in 1970–2017.

Table S6: Marine saponins from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-23ξ-Acetoxy-17-deoxy-7,8-dihydroholothurinogenin 252c ^e [C ₃₂ H ₄₉ O ₅ R]	IR, MS, NMR, [α] _D , ORD, ECD, CT	Saponin	Undetm.	Undetm.	Undetm.	<i>S. chloronotus</i>	NST	[225]
(-)-Sarasinoides J 253 ^β [C ₆₂ H ₁₀₂ N ₂ O ₂₇]	MS, NMR, [α] _D	Saponin	Antifungal	<i>S. cerevisiae</i>	13 mm (10 µg/disk)	<i>M. sarassinorum</i>	SSW	[226 – 228]
			Antibacterial	<i>B. subtilis</i> DSM2109	9 mm (10 µg/disk)			
			Cytotoxic	K562, A549	LC ₅₀ = 10.3 – 20.8 µM			
(-)-Sarasinoides K 254 ^β [C ₆₃ H ₁₀₄ N ₂ O ₂₉]	MS, NMR, [α] _D	Saponin	Cardiovascular	Na ⁺ /K ⁺ -ATPase	IC ₅₀ > 100 µg/mL	<i>M. sarassinorum</i>	SSW	[226]
			Antidiabetic	PTP1B	NA (IC ₅₀ = 15.2 – 16.0 µM)			
			Undetm.	Undetm.	Undetm.			
(-)-Sarasinoides L 255 ^β [C ₆₂ H ₉₈ N ₂ O ₂₈]	MS, NMR, [α] _D	Saponin	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[226]
(-)-Sarasinoides M 256 ^β [C ₆₂ H ₁₀₀ N ₂ O ₂₇]	MS, NMR, [α] _D	Saponin	Cytotoxic	K562, A549	LC ₅₀ = 7.7 – 12.1 µM	<i>M. sarassinorum</i>	SSW	[226, 227]
(-)-Sarasinoides S 257 ^β [C ₆₂ H ₉₆ N ₂ O ₂₆]	UV, IR, MS, NMR, [α] _D	Saponin	Cardiovascular	Na ⁺ /K ⁺ -ATPase	LC ₅₀ > 100 µg/mL	<i>M. sarassinorum</i>	SSW	[226, 227]
			Antidiabetic	PTP1B	NA (IC ₅₀ = 15.2 – 16.0 µM)	<i>Petrosia</i> sp.	NSW	[228]

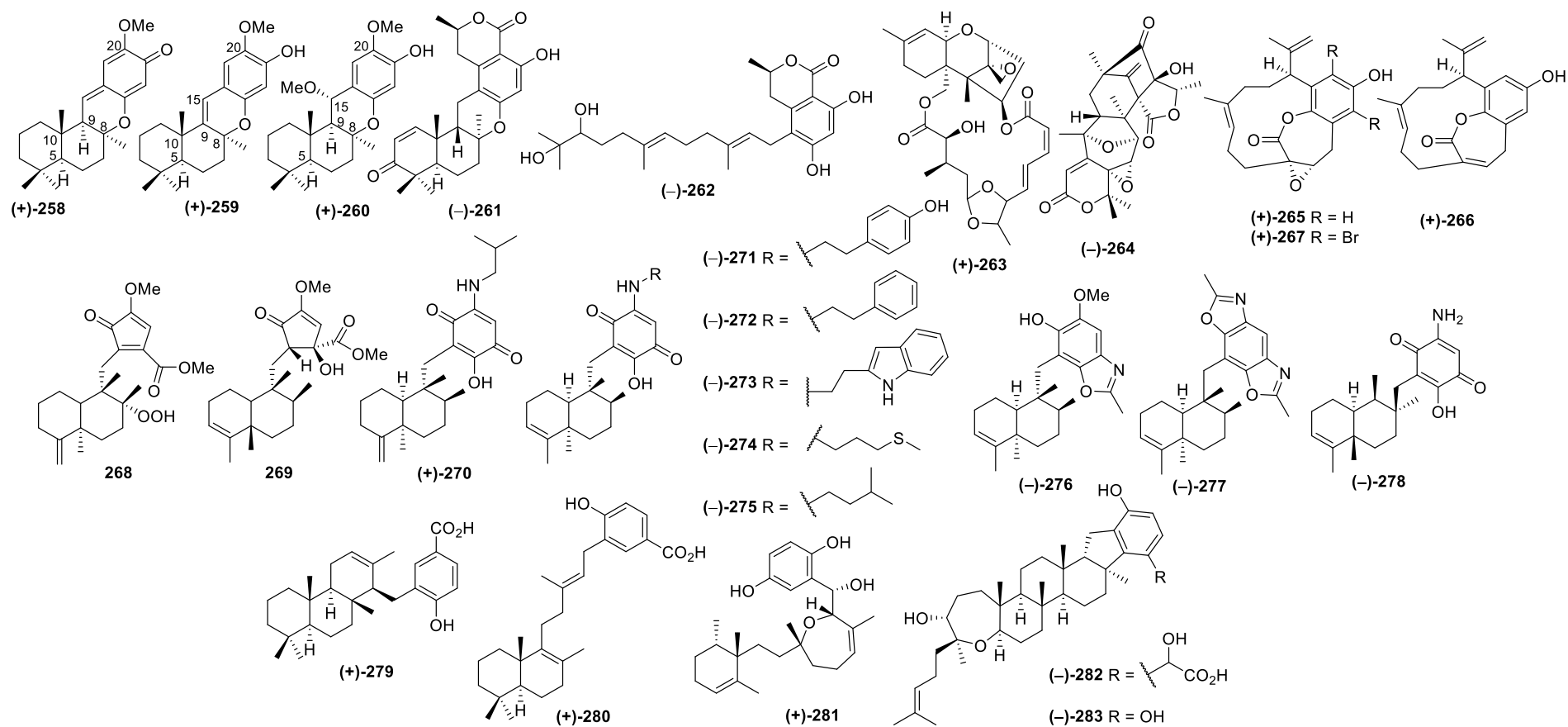


Figure S7: Structures of marine meroterpenoids from Indonesian waters found in 1970–2017.

Table S7: Marine meroterpenoids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-5 <i>S</i> ,8 <i>S</i> ,9 <i>R</i> ,10 <i>S</i> -20-methoxypuuphenone 258 ^β [C ₂₂ H ₃₀ O ₃]	MS, NMR, [α] _D	Merosesqui-terpene	Undetm.	Undetm.	Undetm.	<i>Hyrtios</i> sp.	GTO	[229]
(+)-5 <i>S</i> ,8 <i>S</i> , 10 <i>S</i> -20-methoxy-9,15-ene-puuphenol 259 ^β [C ₂₂ H ₃₀ O ₃]	MS, NMR, [α] _D	Merosesqui-terpene	Antiatherosclerotic	HepG2 transf. SR-B1	EC ₅₀ = 3.05 μM	<i>Hyrtios</i> sp.	GTO	[229, 230]
(+)-5 <i>S</i> ,8 <i>S</i> ,9 <i>R</i> ,10 <i>S</i> -15,20-dimethoxy puuphenol 260 ^β [C ₂₃ H ₃₄ O ₄]	MS, NMR, [α] _D	Merosesqui-terpene	Undetm.	Undetm.	Undetm.	<i>Hyrtios</i> sp.	GTO	[229]
(–)-Verruculide A 261 ^β [C ₂₅ H ₃₀ O ₅]	UV, IR, MS, NMR, ECD, [α] _D , Mol. Mod.	Merosesqui-terpene	Antidiabetic	PTP1B	IC ₅₀ = 8.4 μM	<i>P. verruculosum</i> TPU1311 (symbiont) <i>P. aurata</i> (host)	NSW	[231]
(–)-Verruculide B 262 ^β [C ₂₅ H ₃₆ O ₆]	UV, IR, MS, NMR, ECD, [α] _D , Mol. Mod.	Merosesqui-terpene	Antidiabetic	PTP1B	40% (23.1 μM)	<i>P. verruculosum</i> TPU1311 (symbiont) <i>P. aurata</i> (host)	NSW	[231]
(+)-Roridin R 263 ^β [C ₂₉ H ₃₈ O ₉]	UV, IR, MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	L1210	IC ₅₀ = 0.45 μM	<i>Myrothecium</i> sp. TUF 02F6 (symbiont) a sponge (host)	NSW	[232]
(–)-Citreonigrin A 264 ^β [C ₂₅ H ₂₈ O ₈]	UV, MS, NMR, [α] _D , X-ray	Merosesqui-terpene*	Anticancer	Various protein kinases enzyme	NA	<i>P. citreonigrum</i> (symbiont) <i>P. purpurea</i> (host)	BLI	[233, 234]
(+)-Floresolide A 265 ^β [C ₂₁ H ₂₄ O ₄]	MS, NMR, [α] _D , X-ray	Merosesqui-terpene*	Cytotoxic	KB		<i>Aplidium</i> sp.	ENT	[235, 236]
(+)-Floresolide B 266 ^β [C ₂₁ H ₂₄ O ₃]	MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	KB	IC ₅₀ = 1 – 10 μg/mL	<i>Aplidium</i> sp.	ENT	[235, 236]
(+)-Floresolide C 267 ^β [C ₂₁ H ₂₂ Br ₂ O ₅]	MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	KB		<i>Aplidium</i> sp.	ENT	[235, 236]

Table S7: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Dactylospongenone G 268 ^{α,β} [C ₂₃ H ₃₂ O ₆]	UV, MS, NMR	Merosesqui-terpene	Undetm.	Undetm.	Undetm.	<i>D. elegans</i>	MLU	[237]
Dactylospongenone H 269 ^{α,β} [C ₂₅ H ₃₄ O ₅]	UV, MS, NMR	Merosesqui-terpene	Undetm.	Undetm.	Undetm.	<i>D. elegans</i>	MLU	[237]
(+)-5- <i>epi</i> -Smenospongorine 270 ^β [C ₂₅ H ₃₇ NO ₃]	UV, IR, MS, NMR, [α] _D	Merosesqui-terpene	Anticancer	Pseudo-peroxidase hemoglobin (K562)	2 μM	<i>D. elegans</i>	ENT	[238]
(–)-5- <i>epi</i> -Nakijiquinone S 271 ^β [C ₂₉ H ₃₇ NO ₄]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	L5178Y	IC ₅₀ = 1.7 μM	<i>D. metachromia</i>	MLU	[239]
			Cytotoxic	L5178Y	IC ₅₀ = 1.1 μM			
				<i>S. aureus</i> ATCC 25923 <i>S. aureus</i> ATCC 700699, <i>E. faecium</i> ATCC 35667, <i>E. faecium</i> ATCC 700221 <i>E. faecalis</i> ATCC 29212, <i>E. faecalis</i> ATCC 51299	MIC = 25 μM MIC = 50 μM MIC = 100 μM			
(–)-5- <i>epi</i> -Nakijiquinone Q 272 ^β [C ₂₉ H ₃₇ NO ₃]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Antibacterial			<i>D. metachromia</i>	MLU	[237, 239]
(–)-5- <i>epi</i> -Nakijiquinone T 273 ^β [C ₃₁ H ₃₈ N ₂ O ₃]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	L5178Y	IC ₅₀ = 3.7 μM	<i>D. metachromia</i>	MLU	[239]
(–)-5- <i>epi</i> -Nakijiquinone U 274 ^β [C ₂₅ H ₃₇ NO ₃ S]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	L5178Y	IC ₅₀ = 1.8 μM	<i>D. metachromia</i>	MLU	[239]
			Cytotoxic	L5178Y	IC ₅₀ = 1.3 μM			
(–)-5- <i>epi</i> -Nakijiquinone N 275 ^β [C ₂₆ H ₃₉ NO ₃]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Anticancer	AKTK1, Aurora-B, METwt, NEK2, NEK6, PIM1, PLK1	IC ₅₀ = 29.9 – 73.2 μM	<i>D. metachromia</i>	MLU	[239]
				ALK	IC ₅₀ = 0.97 μM			
				ARK5, AXL, MEK1 wt, PRK1 FAK, IGFI-R, SRC, VEGF-R2	IC ₅₀ > 100 μM IC ₅₀ = 1.94 – 3.03 μM			
(–)-5- <i>epi</i> -Nakijinol C 276 ^β [C ₂₄ H ₃₃ NO ₃]	UV, MS, NMR, [α] _D	Merosesqui-terpene [▲]	Anticancer	L5178Y	IC ₅₀ > 10.0 μM	<i>D. metachromia</i>	MLU	[239]
				AKT1, ARK5, Aurora-B, MEK1 wt, MET wt, NEK6, PIM1, PLK1, PRK1	IC ₅₀ > 100 μM			
				ALK	IC ₅₀ = 3.38 μM			
				AXL, FAK, SRC IGF1-R, VEGF-R2	IC ₅₀ = 7.78 – 19.7 μM IC ₅₀ = 3.31 – 3.66 μM			

Table S7: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-5- <i>epi</i> -Nakijinol D 277 ^β [C ₂₅ H ₃₂ N ₂ O ₂]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Cytotoxic	L5178Y	IC ₅₀ > 10.0 μM	<i>D. metachromia</i>	MLU	[239]
(-)-Dysideamine 278 ^β [C ₂₁ H ₂₉ NO ₃]	UV, MS, NMR, [α] _D	Merosesqui-terpene	Neuro disease	HT22	43% (HT22 surv. 10 μM)	<i>Dysidea</i> sp.	UEP	[240]
				Neuro 2A	(IAA ind.) 40%, 3.0 μM; 25%, 10 μM (AChE incr.)			
(+)-Makassaric acid 279 ^β [C ₂₇ H ₃₈ O ₃]	UV, MS, NMR, [α] _D	Mero diterpene	Anti-inflammatory	MK-2	IC ₅₀ = 20 μM	<i>Acantho-dendrilla</i> sp.	SSW	[241, 242]
(+)-Subersic acid 280 ^β [C ₂₇ H ₃₈ O ₃]	UV, MS, NMR, [α] _D	Mero diterpene	Anti-inflammatory	MK-2	IC ₅₀ = 9.6 μM	<i>Acantho-dendrilla</i> sp.	SSW	[241, 242]
(+) -Halioxepine 281 ^β [C ₂₆ H ₃₈ O ₄]	UV, IR, MS, NMR, [α] _D , CT	Mero diterpene*	Cytotoxic	NBT-T2	IC ₅₀ = 4.8 μg/mL	<i>Haliclona</i> sp.	SES	[243]
			Antioxidant	DPPH	IC ₅₀ = 3.2 μg/mL			
(-) -Haliclotriol A 282 ^β [C ₃₈ H ₅₆ O ₆]	UV, IR, MS, NMR, [α] _D	Mero triterpene*	Cytotoxic	Various murine and human cancer	NA	<i>Haliclona</i> sp.	NMU	[244]
(-) -Haliclotriol B 283 ^β [C ₃₆ H ₅₄ O ₄]	UV, IR, MS, NMR, [α] _D	Mero triterpene	Antibacterial	<i>B. subtilis</i> , <i>S. aureus</i>	1 mg/disk	<i>Haliclona</i> sp.	NMU	[244]

Footnote: 1. Activity (HT22 murine hippocampal neuronal, **AChE** acetylcholinesterase, **AKT1** protein kinase B, **ALK** anaplastic lymphoma kinase, **ARK5** AMPK-related protein kinase 5, **AXL** tyrosine-protein kinase, **FAK** focal adhesion kinase, **IGF1-R** insulin like growth factor 1 receptor, **MEK1** mitogen-activated protein kinase kinase, **MET** tyrosine-protein kinase, **MK-2** mitogen-activated protein kinase-activated protein kinase 2, **NEK2** serine/threonine-protein kinase, **NEK6** serine/threonine-protein kinase, **PIM1** serine/threonine-protein kinase, **PLK1** serine/threonine-protein kinase, **PRK1** serine/threonine-protein kinase, **SRC** non-receptor tyrosine kinase, **VEGF-R2** vascular endothelial growth factor receptor 2, **DPPH** 1,1-diphenyl-2-picrylhydrazyl, **IAA** iodoacetic acid, **surv.** survived, **incr.** Increased, **ind.** induced).

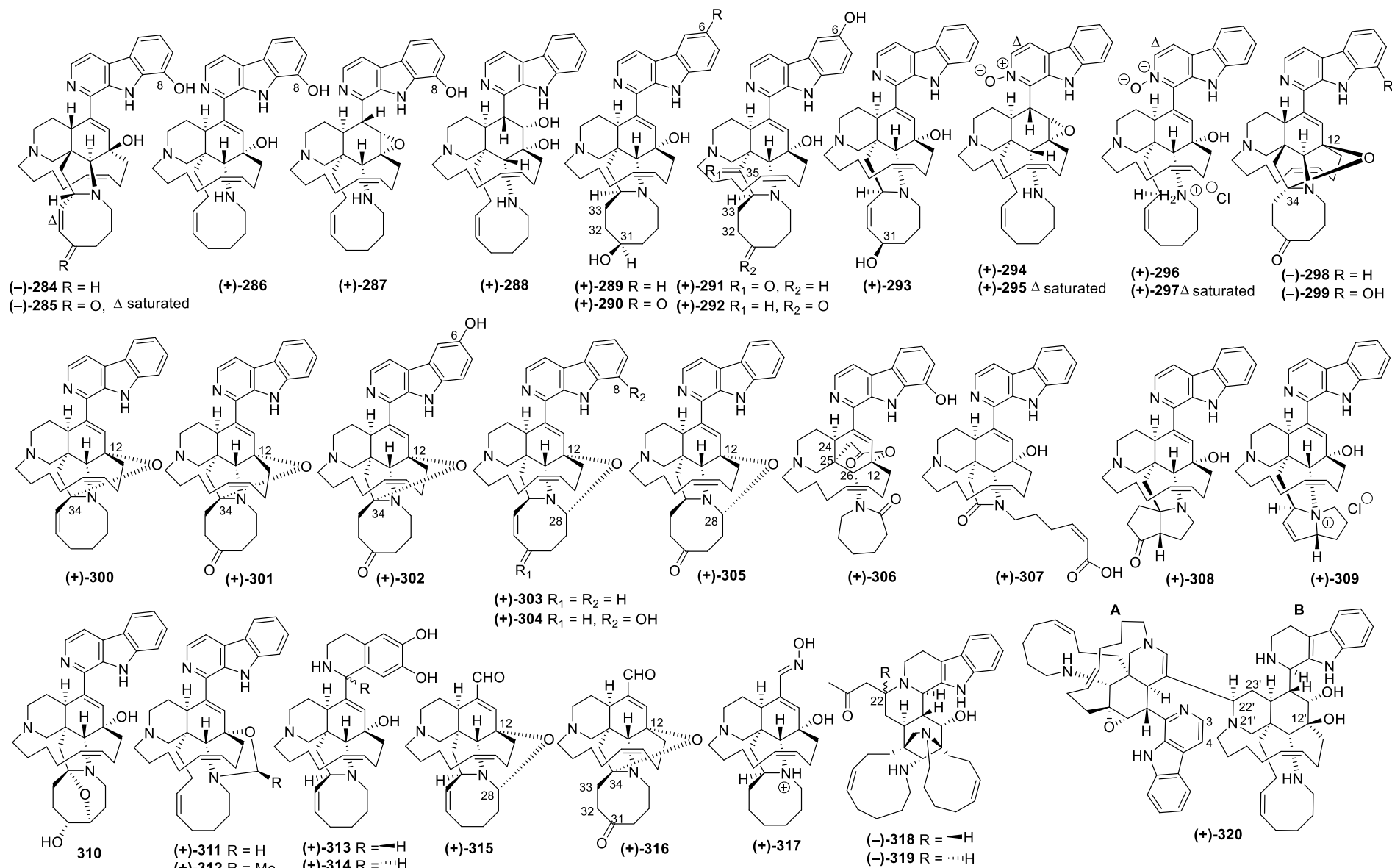


Figure S8: *Cont.*

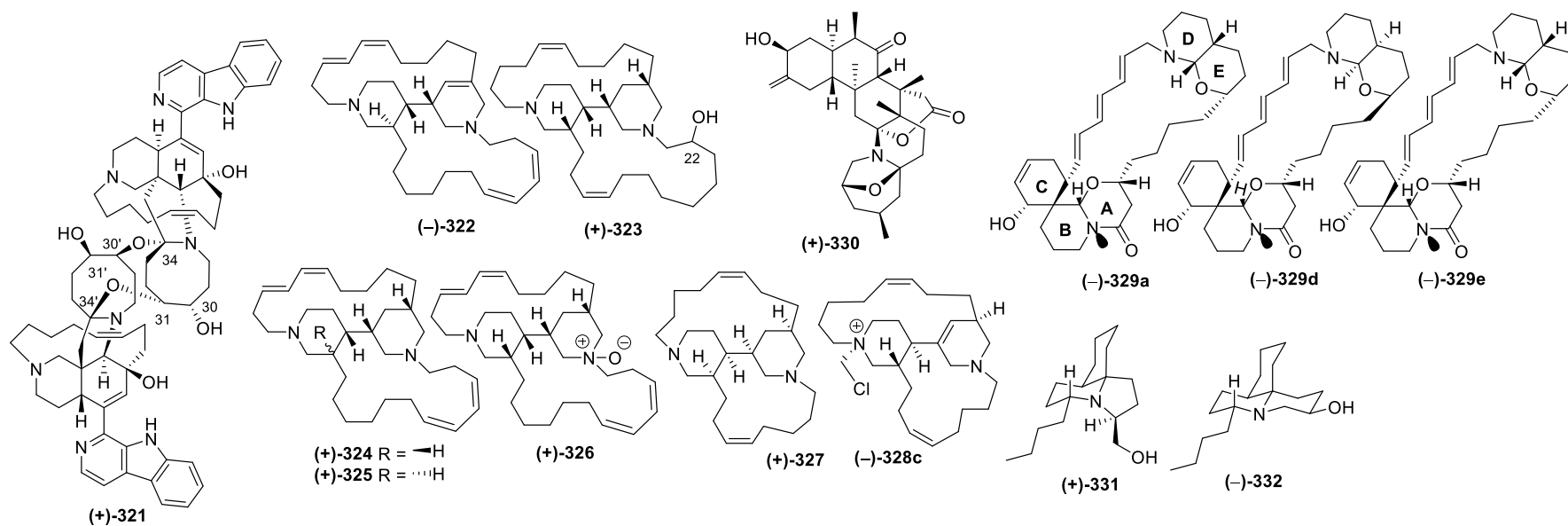


Figure S8: Structures of marine piperidine alkaloids from Indonesian waters found in 1970–2017.

Table S8: Marine piperidine alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-8-Hydroxymanzamine A 284 ^{β,η} [C ₃₆ H ₄₄ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antiparasite	<i>T. gondii</i>	71% (1 μM) [38% hc.] 9 – 12 days	A sponge (<i>Petrosiidae</i>)	NSW	[245 – 248]
			Antibacterial	<i>P. berghei</i>	(100 μmoles/kg, wo. tox.)			
			Antiinsecticidal	<i>M. tuberculosis</i> H37Rv	MIC = 3.13 μg/mL			
				WCR	100% (2 mM)			
				WTPB	25% (2 mM)			
			Antifungal	<i>S. nodorum</i>	92% (10 μg/mL)			
				<i>F. culmorum</i> , <i>P. recondita</i>	0% (10 μg/mL)			
(-)-Manzamine F 285 ^{β,η} [C ₃₆ H ₄₄ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Anti-inflammatory	<i>P. infestans</i> , <i>P. grisei</i>	22 – 41% (10 μg/mL)	A sponge (<i>Petrosiidae</i>)	NSW	[245, 248]
				B ₂	IC ₅₀ > 30 μM (SA)			
			Antiparasite	<i>T. gondii</i>	37% (10 μM, wo. tox.)			
				<i>P. berghei</i>	NA			
			Antibacterial		(100 μmoles/kg, wo. tox.) 98 – 99%			
				<i>M. tuberculosis</i> H37Rv	(MIC < 12.5 μg/mL)			
			Antiinsecticidal	WCR	100% (3 mM)			
(+) -8-Hydroxymanzamine J 286 ^{β,η} [C ₃₆ H ₄₆ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid		WTPB	0% (3 mM)	Acanthostro- ngylophora sp.	NSW	[249]
			Antifungal	<i>S. nodorum</i> , <i>P. infestans</i>	79 – 80 % (10 μg/mL)			
				<i>P. grisei</i> , <i>F. culmorum</i>	19 – 31% (10 μg/mL)			
				<i>P. recondita</i>	0% (10 μg/mL)			
			Antibacterial	<i>S. aureus</i> , MRSA	IC ₅₀ = 3.0 – 5.0 μg/mL			
			Antifungal	<i>M. intracellulare</i>	IC ₅₀ = 0.45 μg/mL			
				<i>C. neoformans</i>	IC ₅₀ = 3.5 μg/mL			
(+) -8-Hydroxymanzamine B 287 ^{β,η} [C ₃₆ H ₄₆ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Undetm.	Undetm.	Undetm.	Acanthostro- ngylophora sp.	NSW	[250]
			Cytotoxic	A549, K562	LC ₅₀ = 6.2 – 8.2 μM			
(+) -11-Hydroxymanzamine J 288 ^{β,θ} [C ₃₆ H ₄₈ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>P. hauseri</i> (NBRC 3851), <i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708)	MIC = 2.0 – 4.0 ng/mL	Acanthostro- ngylophora sp.	JSCR	[251]

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-11-Hydroxymanzamine J 288 ^{β,θ} [C ₃₆ H ₄₈ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial Hypercholesterolemic Cardiovascular	<i>S. enterica</i> (ATCC 14028), <i>E. coli</i> (ATCC 35270) Isocytate lyase Na ⁺ /K ⁺ -ATPase	MIC = 8.0 – 16.0 ng/mL IC ₅₀ = 27 μM IC ₅₀ > 150 μM	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
(+)-32,33-Dihydro-31-hydroxymanzamine A 289 ^{β,θ} [C ₃₆ H ₄₆ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT, X-ray	Piperidine Alkaloid	Antiparasite Cytostatic	<i>P. falciparum</i> (D6 clone), (W2 clone), <i>L. donovani</i> V79	NA NA (4.76 μg/mL)	A sponge (<i>Petrosiidae</i>)	NSW	[252]
(+)-32,33-Dihydro-6,31-dihydroxymanzamine A 290 ^{β,θ} [C ₃₆ H ₄₆ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D , CT	Piperidine Alkaloid	Undetm.	Undetm.	Undetm.	A sponge (<i>Petrosiidae</i>)	NSW	[252]
(+)-32,33-dihydro-6-hydroxymanzamine A-35-one 291 ^{β,θ} [C ₃₆ H ₄₄ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antiparasite Cytostatic Antibacterial	<i>P. falciparum</i> (D6 clone), (W2 clone), <i>L. donovani</i> V79 <i>M. tuberculosis</i> H37Rv <i>M. intracellulare</i> <i>C. neoformans</i>	NA NA (4.76 μg/mL) IC ₅₀ = 0.4 μg/mL IC ₅₀ = 3.5 μg/mL IC ₅₀ = 5.5 μg/mL	A sponge (<i>Petrosiidae</i>)	NSW	[252]
(+)-6-Hydroxymanzamine E 292 ^{β,η} [C ₃₆ H ₄₄ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antifungal Antiparasite Cytostatic Cytotoxic	<i>P. falciparum</i> (D6 clone), (W2 clone) <i>L. donovani</i> V79 A549, K562	IC ₅₀ = 0.78 – 0.87 μg/mL IC ₅₀ = 2.5 – 4.3 μg/mL IC ₅₀ = 4.3 μg/mL LC ₅₀ = 5.8 – 7.2 μM	<i>Acanthostro- ngylophora</i> sp.	NSW	[249, 250]
(+)-31-Hydroxymanzamine A 293 ^{β,θ} [C ₃₆ H ₄₄ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D , CT	Piperidine Alkaloid	Antibacterial Hypercholesterolemic Cardiovascular	<i>S. aureus</i> (ATCC 6538p), <i>P. hauseri</i> (NBRC 3851) <i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708) <i>S. enterica</i> (ATCC 14028) <i>E. coli</i> (ATCC 35270) Isocytate lyase Na ⁺ /K ⁺ -ATPase	MIC = 13 – 25 ng/mL MIC = 6.3 ng/mL MIC = 1.6 ng/mL MIC > 100 ng/mL IC ₅₀ = 140 μM IC ₅₀ > 150 μM	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
(+)-Manzamine B N-oxide 294 ^{β,η} [C ₃₆ H ₄₆ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Cytotoxic	A549, K562	LC ₅₀ = 9.8 – 12.0 μM	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Manzamine B N-oxide 294 ^{β,η} [C ₃₆ H ₄₆ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>P. hauseri</i> (NBRC 3851), <i>E. coli</i> (ATCC 35270)	MIC > 100 ng/mL	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
				<i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708)	MIC = 100 ng/mL			
			Hypercholesterolemic	<i>S. enterica</i> (ATCC 14028)	MIC = 50 ng/mL			
			Cardiovascular Cytotoxic	Isocytate lyase Na ⁺ /K ⁺ -ATPase A549, K562	IC ₅₀ > 150 μM LC ₅₀ = 5.2 – 5.8 μM			
(+)–3,4-Dihydromanizamine B N-oxide 295 ^{β,η} [C ₃₆ H ₄₈ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>P. hauseri</i> (NBRC 3851)	MIC = 13 – 25 ng/mL	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
				<i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708), <i>S. enterica</i> (ATCC 14028)	MIC = 3.1 – 6.3 ng/mL			
			Hypercholesterolemic	<i>E. coli</i> (ATCC 35270)	MIC > 100 ng/mL			
			Cardiovascular Cytotoxic	Isocytate lyase Na ⁺ /K ⁺ -ATPase A549, K562	IC ₅₀ = 70 μM IC ₅₀ > 150 μM LC ₅₀ = 4.7 – 8.1 μM			
(+)–Manzamine J N-oxide-HCl 296 ^{β,ι} [C ₃₆ H ₄₇ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>S. enterica</i> (ATCC 14028)	MIC = 32 ng/mL	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
				<i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708), <i>P. hauseri</i> (NBRC 3851), <i>E. coli</i> (ATCC 35270)	MIC > 100 ng/mL			
			Hypercholesterolemic	Isocytate lyase Na ⁺ /K ⁺ -ATPase A549, K562	IC ₅₀ = 26 μM IC ₅₀ > 150 μM LC ₅₀ = 5.7 – 9.5 μM			
			Cardiovascular Cytotoxic					
(+)–3,4-Dihydromanizamine J N-oxide-HCl 297 ^{β,ι} [C ₃₆ H ₄₉ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>B. subtilis</i> (ATCC 6633), <i>K. rhizophila</i> (NBRC 12708), <i>P. hauseri</i> (NBRC 3851)	MIC = 32 – 64 ng/mL	<i>Acanthostro- ngylophora</i> sp.	JSCR	[251]
				<i>S. enterica</i> (ATCC 14028), <i>E. coli</i> (ATCC 35270)	MIC > 100 ng/mL			

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -3,4-Dihydromanzamine J N-oxide-HCl 297 ^{β,ι} [C ₃₆ H ₄₉ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Hypercholesterolemic	Isocitrate lyase	IC ₅₀ > 150 μM	<i>Acanthostromylophora</i> sp.	JSCR	[251]
			Cardiovascular	Na ⁺ /K ⁺ -ATPase	IC ₅₀ = 110 μM			
(–)-12,34-Oxamanzamine E 298 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>M. tuberculosis</i> H37Rv	MIC = 128 μg/mL	A sponge (<i>Petrosiidae</i>)	NSW	[246, 252]
			Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone)	NA			
			Antifungal	<i>C. neoformans</i>	NA			
			Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone)	IC ₅₀ = 0.84 – 1.10 μg/mL			
(–)-12,34-Oxamanzamine F 299 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Cytostatic	<i>L. donovani</i> V79	NA (4.76 μg/mL)	A sponge (<i>Petrosiidae</i>)	NSW	[246]
			Antibacterial	<i>S. aureus</i> , MRSA, <i>M. intracellulare</i>	NA			
				<i>M. tuberculosis</i> H37Rv	MIC = 12.5 μg/mL			
			Antiviral	HIV-1	IC ₅₀ = 14.9 μM			
(+) -12,34-Oxamanzamine A 300 ^{β,θ} [C ₃₆ H ₄₂ N ₄ O]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antiparasite	<i>P. falciparum</i> (D6 clone) (W2 clone)	IC ₅₀ = 4.76 μg/mL NA	A sponge (<i>Petrosiidae</i>)	NSW	[246]
			Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone), <i>L. donovani</i> V79	NA			
			Cytostatic		NA (4.76 μg/mL)			
(+) -12,34-Oxamanzamine E 301 ^{β,θ} [C ₃₆ H ₄₂ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> , MRSA, <i>M. intracellulare</i>	NA	<i>Acanthostromylophora</i> sp.	NSW	[249, 250, 252]
			Antifungal	<i>C. neoformans</i>	NA			
			Antiviral	HIV-1	EC ₅₀ = 17.5 μM			
			Antiatherosclerotic	Human monocyte-derived macrophage	NA			
(+) -12,34-Oxa-6-hydroxymanzamine E 302 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone), <i>L. donovani</i>	NA	<i>Acanthostromylophora</i> sp.	NSW	[250]

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -12,28-Oxamanzamine A 303 ^{β,θ} [C ₃₆ H ₄₂ N ₄ O]	UV, IR, MS, NMR, [α] _D , Mol. Mod.	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> , MRSA, <i>M. intracellulare</i>	NA	A sponge (<i>Petrosiidae</i>)	NSW	[250, 253]
			Antiviral	HIV	EC ₅₀ = 22.2 μM			
			Antifungal	<i>C. neoformans</i> <i>P. falciparum</i> (D6 clone), (W2 clone)	NA			
			Antiparasite	<i>L. donovani</i>	IC ₅₀ = 7.8 μg/mL IC ₉₀ = 50 μg/mL			
(+) -12,28-Oxa-8-hydroxymanzamine A 304 ^{β,θ} [C ₃₆ H ₄₂ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Cytostatic	V79	NA (4.76 μg/mL)	A sponge (<i>Petrosiidae</i>)	NSW	[253]
			Antibacterial	<i>S. aureus</i> , MRSA, <i>M. intracellulare</i>	NA			
			Antifungal	<i>C. neoformans</i> <i>P. falciparum</i> (D6 clone), (W2 clone)	NA			
			Antiparasite	<i>L. donovani</i>	IC ₅₀ = 18 μg/mL IC ₉₀ = 40 μg/mL			
(+) -12,28-Oxamanzamine E 305 ^{β,θ} [C ₃₆ H ₄₂ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	NA (4.76 μg/mL)	V79	NA (4.76 μg/mL)	<i>Acanthostromyngylophora</i> sp.	NSW	[250]
			Cytostatic	V79	NA (4.76 μg/mL)			
			Antibacterial	<i>S. aureus</i> , MRSA, <i>M. intracellulare</i>	NA			
			Antifungal	<i>C. neoformans</i>	NA			
(+) -Acantholactone 306 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₄]	UV, MS, NMR, [α] _D , Mol. Mod, ECD	Piperidine Alkaloid ^Δ	Undetm.	Undetm.	Undetm.	<i>Acanthostromyngylophora</i> sp.	NSW	[254]
(+) -Acantholactam 307 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₄]	UV, MS, NMR, [α] _D , ECD	Piperidine Alkaloid ^Δ	Cytotoxic	HeLa	IC ₅₀ > 50 μM	<i>A. ingens</i>	NSW	[255]
			Anticancer	Chymotrypsin-like activity	IC ₅₀ = 33 μM			
(+) -Acanthomanzamine C 308 ^{β,η} [C ₃₆ H ₄₂ N ₄ O ₂]	UV, MS, NMR, [α] _D	Piperidine Alkaloid ^Δ	Antiatherosclerotic	Human monocyte-derived macrophage	NA (20 μM)	<i>A. ingens</i>	NSW	[256]
			Undetm.	Undetm.	Undetm.			
(+) -Kepulauamine A 309 ^{β,ι} [C ₃₆ H ₄₃ N ₄ O]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid ^Δ	Cytotoxic	A549, K562	LC ₅₀ = 4.6 – 7.2 μM	<i>Acanthostromyngylophora</i> sp.	JSCR	[251]

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Kepulauamine A 309 ^{β,4} [C ₃₆ H ₄₃ N ₄ O]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> (ATCC 6538p), <i>P. hauseri</i> (NBRC 3851) <i>B. subtilis</i> (ATCC 6633), <i>E. coli</i> (ATCC 35270) <i>K. rhizophila</i> (NBRC 12708), <i>S. enterica</i> (ATCC 14028)	MIC = 8.0 ng/mL MIC = 32 – 64 ng/mL MIC = 16 ng/mL	<i>Acanthostromylophora</i> sp.	JSCR	[251]
			Hypercholesterolemic	Isocitrate lyase	IC ₅₀ > 150 μM			
			Cardiovascular	Na ⁺ /K ⁺ -ATPase	IC ₅₀ > 150 μM			
			Cytotoxic	HeLa	IC ₅₀ = 16 μM			
<i>Pre-neo</i> -Kauluamine 310 ^{β,η} [C ₃₆ H ₄₄ N ₄ O ₃]	MS, NMR	Piperidine Alkaloid	Anticancer	Chymotrypsin-like activity	IC ₅₀ = 0.34 μM	<i>Acanthostromylophora</i> sp.	NSW	[255]
			Antiatherosclerotic	Human monocyte-derived macrophage	91% (20 μM)			
(+)–Acanthomanzamine D 311 ^{β,η} [C ₃₇ H ₄₆ N ₄ O]	UV, MS, NMR, [α] _D	Piperidine Alkaloid*	Cytotoxic	HeLa	IC ₅₀ = 15 μM	<i>A. ingens</i>	NSW	[256]
			Anticancer	Chymotrypsin-like activity	IC ₅₀ = 0.63 μM			
(+)–Acanthomanzamine E 312 ^{β,η} [C ₃₈ H ₄₈ N ₄ O]	UV, MS, NMR, [α] _D	Piperidine Alkaloid	Antiatherosclerotic	Human monocyte-derived macrophage	73% (20 μM)	<i>A. ingens</i>	NSW	[256]
			Cytotoxic	HeLa	IC ₅₀ > 20 μM			
(–)-Acanthomanzamine A 313 ^{β,η} [C ₃₄ H ₄₇ N ₃ O ₃]	UV, MS, NMR, [α] _D , ECD, Mol. Mod.	Piperidine Alkaloid*	Anticancer	Chymotrypsin-like activity	IC ₅₀ = 4.2 μM	<i>A. ingens</i>	NSW	[256]
			Antiatherosclerotic	Human monocyte-derived macrophage	IC ₅₀ = 4.1 μM			
(+)–Acanthomanzamine B 314 ^{β,η} [C ₃₄ H ₄₇ N ₃ O ₃]	UV, MS, NMR, [α] _D , ECD, Mol. Mod.	Piperidine Alkaloid	Cytotoxic	HeLa	IC ₅₀ = 5.7 μM	<i>A. ingens</i>	NSW	[256]
			Anticancer	Chymotrypsin-like activity	IC ₅₀ = 7.8 μM			
(+)–12,28-Oxaircinal A 315 ^{β,η} [C ₂₆ H ₃₆ N ₂ O ₂]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antiatherosclerotic	Human monocyte-derived macrophage	73% (20 μM)	<i>Acanthostromylophora</i> sp.	NSW	[250]
			Undetm.	Undetm.	Undetm.			

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -31-Keto-12,34-oxa-32,33-dihydroircinal A 316 ^{β,η,θ} [C ₂₆ H ₃₆ N ₂ O ₃]	UV, IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>S. aureus</i> , MRSA,	NA	A sponge (Petrosiidae)	NSW	[253]
			Antifungal	<i>M. intracellulare</i> <i>C. neoformans</i>	NA			
(+) -Ircinal E 317 ^{β,θ} [C ₂₆ H ₄₂ N ₃ O ₂]	MS, NMR, [α] _D	Piperidine Alkaloid	Cytotoxic	L5178Y	IC ₅₀ = 21.7 μM	<i>A. ingens</i>	MLU	[257]
(–)-Manadomanzamine A 318 ^{β,η} [C ₃₉ H ₅₄ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D , ECD, Mol. Mod.	Piperidine Alkaloid*	Antibacterial	<i>M. tuberculosis</i> H37Rv (ATCC 27294)	MIC = 1.9 μg/mL	<i>Acanthostromylophora</i> sp.	NSW	[73]
			Antiviral	HIV-1 LAV	EC ₅₀ = 7.0 μg/mL			
			Antifungal	<i>C. albicans</i> <i>C. neoformans</i>	IC ₅₀ = 20 μg/mL NA			
			Cytotoxic	A549, H-116	IC ₅₀ = 2.5 – 5.0 μg/mL			
(–)-Manadomanzamine B 319 ^{β,η} [C ₃₉ H ₅₄ N ₄ O ₂]	UV, IR, MS, NMR, [α] _D , ECD, Mol. Mod.	Piperidine Alkaloid	Antibacterial	<i>M. tuberculosis</i> H37Rv (ATCC 27294)	MIC = 1.5 μg/mL	<i>Acanthostromylophora</i> sp.	NSW	[73]
			Antiviral	HIV-1 LAV	EC ₅₀ = 16.5 μg/mL			
			Antifungal	<i>C. albicans</i> <i>C. neoformans</i>	NA IC ₅₀ = 3.5 μg/mL			
			Cytotoxic	A549, H-116	IC ₅₀ ≥ 5.0 μg/mL			
(+) -Kauluamine 320 ^{β,η} [C ₇₂ H ₉₄ N ₈ O ₃]	IR, MS, NMR, [α] _D	Piperidine Alkaloid*	Immuno suppressive activity	MLR LcV, LcV/MLR	IC ₅₀ = 1.57 μg/mL IC ₅₀ > 16.0 μg/mL	<i>Prianos</i> sp.	NSW	[258]
				Human lung cancer, human colon carcinoma	IC ₅₀ = 1.0 μg/mL			
			Cytotoxic	A549, HeLa, K562	IC ₅₀ = 5.4 – 13 μM			
				V79	NA (4.7 μg/mL)			
(+) - <i>neo</i> -Kauluamine 321 ^{β,η} [C ₇₂ H ₈₈ N ₈ O ₆]	UV, IR, MS, NMR, [α] _D , Mol. Mod.	Piperidine Alkaloid*	Anticancer	Chymotrypsin-like activity	IC ₅₀ = 0.13 μM	A sponge (Petrosiidae)	NSW	[245, 249, 250, 255]
			Antiatherosclerotic	Human monocyte-derived macrophage	92% (20 μM)			
				<i>M. tuberculosis</i> H37Rv (ATCC 27294)	MIC = 2.0 μg/mL			
			Antibacterial	<i>P. falciparum</i> (D6 clone), (W2 clone) <i>L. donovani</i>	IC ₅₀ = 1.70 – 2.80 μg/mL IC ₅₀ = 4.2 – 8.2 μg/mL			

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(–)-Halicyclamine A 322 ^{β,η} [C ₃₂ H ₅₀ N ₂]	IR, MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial	<i>M. smegmatis</i> (aerobic), <i>M. smegmatis</i> (hypoxic)	MIC = 2.5 µg/mL	<i>Haliclona</i> sp.	PUA	[259 – 262]
				<i>M. bovis</i> BCG (aerobic), <i>M. bovis</i> BCG (hypoxic)	MIC = 1.0 µg/mL			
				<i>M. tuberculosis</i> (aerobic), <i>M. tuberculosis</i> (hypoxic)	MIC = 5.0 µg/mL			
				<i>M. tuberculosis</i> H37Rv ATCC25618, <i>M. tuberculosis</i> H37Rv STR ATCC35820 (streptomycin resist.), <i>M. avium</i> ATCC35712, <i>M. aurum</i> ATCC23366	MIC = 6.25 µg/mL			
				<i>M. tuberculosis</i> H37Rv ETH ATCC35837 (ethambutol resist.), <i>M. tuberculosis</i> H37Rv INH ATCC35822 (isoniazid resist.), <i>M. tuberculosis</i> H37Rv RIF ATCC35838 (rifampicin), <i>M. tuberculosis</i> Kurono RIF ATCC35761, <i>M. kansasii</i> ATCC35775	MIC = 3.13 µg/mL			
				<i>M. fortuitum</i> ATCC9820 P-388	MIC = 25 µg/mL IC ₅₀ = 0.45 µg/mL			
			Cytotoxic	<i>M. smegmatis</i> (aerobic)	MIC = 12.5 µg/mL			
			Antibacterial	<i>M. smegmatis</i> (hypoxic), <i>M. bovis</i> BCG (aerobic) <i>M. bovis</i> BCG (hypoxic)	MIC = 25 µg/mL MIC = 50 µg/mL		ENT	[263]
(+)-22-Hydroxyhaliclona cyclamine B 323 ^{β,η} [C ₃₂ H ₅₆ N ₂ O]	MS, NMR, [α] _D	Piperidine Alkaloid	Antibacterial			<i>Haliclona</i> sp.	ENT	[263]
(+)-Tetradehydro haliclona cyclamine A 324 ^{β,η} [C ₃₂ H ₅₂ N ₂]	MS, NMR, [α] _D , X-ray	Piperidine Alkaloid	Cytotoxic	P-388	IC ₅₀ = 1.80 µg/mL	<i>Halichondria</i> sp.	BLI	[264]
(+)-2- <i>epi</i> -Tetradehydro haliclona cyclamine 325 ^{β,η} [C ₃₂ H ₅₂ N ₂]	MS, NMR, [α] _D	Piperidine Alkaloid	Undetm.	Undetm.	Undetm.	<i>Halichondria</i> sp.	BLI	[264]

Table S8: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Tetradehydro haliclonyclamine A mono-N-oxide 326 ^{β,η} [C ₃₂ H ₅₂ N ₂ O]	MS, NMR, [α] _D	Piperidine Alkaloid	Undetm.	Undetm.	Undetm.	<i>Halichondria</i> sp.	BLI	[264]
(-)-Acanthocyclamine A 327 ^{β,η} [C ₂₆ H ₄₄ N ₂]	MS, NMR, [α] _D , X-ray	Piperidine-Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. ingens</i>	SES	[265]
(-)-Chloromethyl halicyclamine B 328 ^{c,η} [C ₂₇ H ₄₄ ClN ₂]	MS, NMR, [α] _D , ECD, Mol. Mod.	Piperidine-Alkaloid	Anticancer	CK1δ/ε	IC ₅₀ = 6 μM	<i>A. ingens</i>	SSW	[266]
(-)-Upenamamide 329a ^{β,η} [C ₃₂ H ₄₆ N ₂ O ₄]	MS, NMR, [α] _D	Piperidine-Alkaloid [†]	Undetm.	Undetm.	Undetm.	<i>Echinochalina</i> sp.	EKM	[267]
(-)-Upenamamide 329d ^η [C ₃₂ H ₄₆ N ₂ O ₄]	TS	Piperidine-Alkaloid	Undetm.	Undetm.	Undetm.			[268]
(-)-Upenamamide 329e ^η [C ₃₂ H ₄₆ N ₂ O ₄]	TS	Piperidine-Alkaloid	Undetm.	Undetm.	Undetm.			[268]
(+)-Loboanthamine 330 ^{β,η} [C ₃₀ H ₄₃ NO ₅]	IR, MS, NMR, [α] _D , CT	Piperidine-Alkaloid	Cytotoxic	AGS, C6	IC ₅₀ > 50 μM	<i>Lobophytum</i> sp.	NSW	[269]
(+)-Polycitorol A 331 ^{β,θ} [C ₁₇ H ₃₁ NO]	IR, MS, NMR, [α] _D	Piperidine-Alkaloid	Undetm.	Undetm.	Undetm.	An ascidian (Polycitoridae)	ENT	[270]
(+)-Polycitorol B 332 ^{β,θ} [C ₁₇ H ₃₁ NO]	IR, MS, NMR, [α] _D	Piperidine-Alkaloid	Undetm.	Undetm.	Undetm.	An ascidian (Polycitoridae)	ENT	[270]

Footnote: 1. **Structure** (^ηmolecule isolated as freebase, ^θmolecule isolated as TFA salt, [†]molecule isolated as HCl salt); 2. **Activity** (**B2** murine neonatal brain microglia, **AGS** human stomach adenocarcinoma, **H-116** human colorectal adenocarcinoma, **HIV** human immunodeficiency virus, **D6 clone** *Plasmodium falciparum* chloroquine-sensitive, **W2 clone** *Plasmodium falciparum* chlorine-resistant, **MRSA** methicillin-resistant *Staphylococcus aureus*, **CK1δ/ε** protein kinase, **hc.** host cell, **wo.** without, **tox.** toxicity).

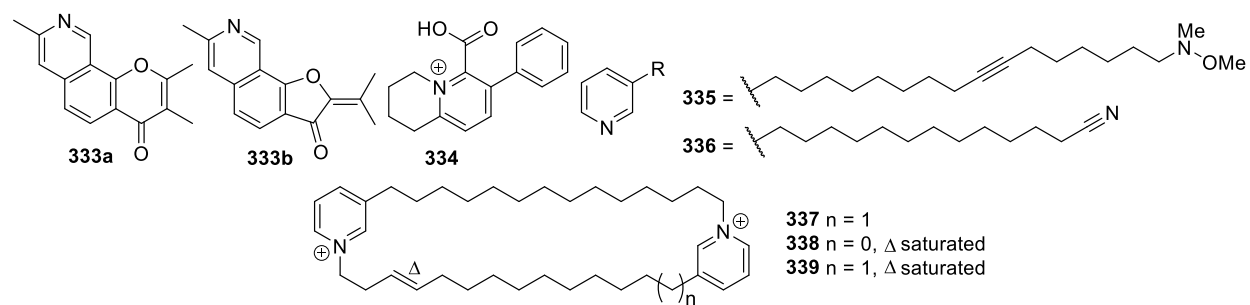


Figure S9: Structures of marine pyridine alkaloids from Indonesian waters found in 1970–2017.

Table S9: Marine pyridine alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Aspergillitine 333a ^{β,η} [C ₁₅ H ₁₃ NO ₂]	UV, MS, NMR	Pyridine Alkaloid	Antibacterial	<i>B. subtilis</i>	7 – 8 mm (5 – 10 µg/disk)	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[74, 271, 272]
			Antifungal	<i>E. coli</i>	NA			
				<i>S. cerevisiae</i>	NA			
TMC-120B 333b ^{δ,η} [C ₁₅ H ₁₃ NO ₂]	TS	Pyridine Alkaloid	Undetm.	Undetm.	Undetm.			[74, 271, 272]
Clathryimine A 334 ^{β,κ} [C ₁₆ H ₁₆ NO ₂]	UV, IR, MS, NMR, CT	Pyridine Alkaloid*	Undetm.	Undetm.	Undetm.	<i>C. basilana</i>	UEP	[273]
<i>N</i> -methylniphyatyne A 335 ^{β,η} [C ₂₃ H ₃₈ N ₂ O]	UV, IR, MS, NMR	Pyridine Alkaloid	Anticancer	PANC-1	IC ₅₀ = 16 µM (natural, Glu-def. Med.) IC ₅₀ = 17 µM (synthetic, Glu-def. Med.) IC ₅₀ > 100 µM (natural, Gen. Glu. Med.) IC ₅₀ > 100 µM (synthetic, Gen. Glu. Med.)	<i>Xestospongia</i> sp.	UEP	[274]
3-Dodecyl pyridine 336 ^{β,η} [C ₁₈ H ₂₈ N ₂]	MS, NMR	Pyridine Alkaloid	Cytotoxic	HeLa, A549, MCF7	IC ₅₀ = 33.2 – 48.4 µM	<i>Haliclona</i> sp.	UEP	[275]
Halilocyclamine A 337 ^{β,κ} [C ₃₅ H ₅₆ N ₂]	UV, IR, MS, NMR	Pyridine Alkaloid	Antibacterial	<i>M. smegmatis</i>	10 – 17 mm (5 – 10 µg/disk)	<i>Haliclona</i> sp.	NSW	[276]
Halilocyclamine B 338 ^{β,κ} [C ₃₈ H ₆₄ N ₂]	UV, IR, MS, NMR	Pyridine Alkaloid	Antibacterial	<i>M. smegmatis</i>	7 – 10 mm (5 – 10 µg/disk)	<i>Haliclona</i> sp.	NSW	[276]
Halilocyclamine C 339 ^{β,κ} [C ₃₉ H ₆₆ N ₂]	UV, IR, MS, NMR	Pyridine Alkaloid	Antibacterial	<i>M. smegmatis</i>	9 – 13 mm (5 – 10 µg/disk)	<i>Haliclona</i> sp.	NSW	[276]

Footnote: 1. Activity (PANC-1 human pancreatic carcinoma, Glu-def. Med Glucose-deficient Medium, Gen. General).

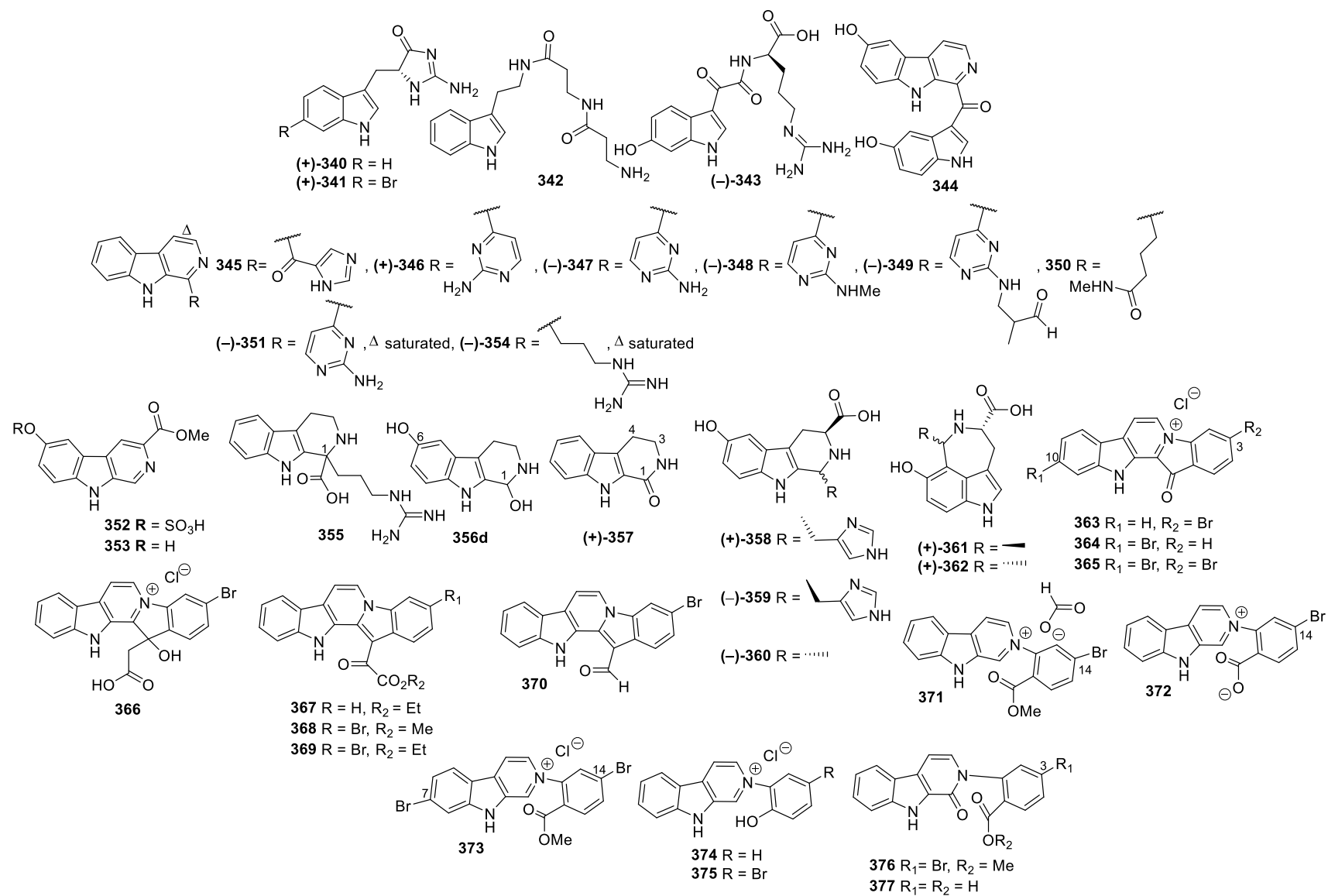


Figure S10: Structures of marine indole alkaloids from Indonesian waters found in 1970–2017.

Table S10: Marine indole alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-2-Amino-1,5-dihydro-5-(1H-indol-3-ylmethyl)-4H-imidazol-4-one 340 ^{β,η} [C ₁₂ H ₁₂ N ₄ O]	IR, MS, NMR, [α] _D , TS	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>H. aurora</i>	BLI	[277]
(+)-2-amino-5-[(6-bromo-1H-indol-3-yl)methyl]-3,5-dihydro-3-methyl-4H-imidazol-4-one 341 ^{β,η} [C ₁₃ H ₁₃ BrN ₄ O]	IR, MS, NMR, [α] _D , TS	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>H. aurora</i>	BLI	[277]
Leptoclinidamide 342 ^{β,θ} [C ₁₆ H ₂₂ N ₄ O ₂]	UV, IR, MS, NMR	Indole Alkaloid	Cytotoxic	HCT15, Jurkat	NA (30 μM)	<i>L. dubius</i>	NSW	[278]
			Antifungal	<i>M. hiemalis</i> IAM608, <i>S. cerevisiae</i> IAM 1438T	NA (250 μg/disk)			
			Antibacterial	<i>S. aureus</i> IAM 12544T, <i>E. coli</i> IAM 12119T	NA (250 μg/disk)			
(–)-Leptoclinidamine B 343 ^{β,θ} [C ₁₆ H ₁₉ N ₅ O ₅]	UV, IR, MS, NMR, [α] _D , CT	Indole Alkaloid	Cytotoxic	HCT15, Jurkat	NA (30 μM)	<i>L. dubius</i>	NSW	[278]
			Antifungal	<i>M. hiemalis</i> IAM608, <i>S. cerevisiae</i> IAM 1438T	NA (250 μg/disk)			
			Antibacterial	<i>S. aureus</i> IAM 12544T <i>E. coli</i> IAM 12119T	NA (250 μg/disk) NA (250 μg/disk)			
			Anti-inflammatory	Phospholipase A ₂ (<i>A. mellifera</i>)	IC ₅₀ > 1166 μM			
Hyrtiosulawesine 344 ^{β,η} [C ₂₀ H ₁₂ N ₃ O ₃]	UV, MS, NMR	Indole Alkaloid	Antiparasite	<i>P. falciparum</i> (FcB1) <i>P. falciparum</i> (W2 clone)	IC ₅₀ = 1.3 ± 0.2 μM NA	<i>Hyrtios</i> sp.	SSW	[279 – 281]
				<i>L. donovani</i>	IC ₅₀ = 35 μg/mL IC ₉₀ > 50 μg/mL			
			Cytotoxic	COLO-205 V79	79.2% (10 μM) NA (4.76 μg/mL)			
Des- <i>N</i> -Methylxestomanzamine A 345 ^{β,η} [C ₁₅ H ₁₀ N ₄ O]	UV, MS, NMR	Indole Alkaloid	Antiparasite	<i>P. falciparum</i> (D6 clone), <i>P. falciparum</i> (W2 clone)	NA	A sponge (<i>Petrosiidae</i>)	NSW	[252]
			Cytotoxic	<i>L. donovani</i> V79	IC ₅₀ = 35 μg/mL IC ₉₀ > 50 μg/mL NA (4.76 μg/mL)			

Table S10: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Ingenine A 346 ^{β,η} [C ₁₅ H ₁₁ N ₅]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	L5178Y	ED ₅₀ < 10 µg/mL	<i>A. ingens</i>	SSW	[282]
(-)-Ingenine B 347 ^{β,η} [C ₁₅ H ₁₁ N ₅]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	L5178Y	ED ₅₀ = 9.1 µg/mL	<i>A. ingens</i>	SSW	[282]
(-)-Ingenine C 348 ^{β,η} [C ₁₆ H ₁₃ N ₅]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	MCF7 A549 HCT116	IC ₅₀ = 4.33 µM W IC ₅₀ = 6.05 µM	<i>A. ingens</i>	UEP	[283]
(-)-Ingenine D 349 ^{β,η} [C ₁₉ H ₁₇ N ₅ O]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	MCF7, HCT116 A549	IC ₅₀ = 2.90 – 3.35 µM W	<i>A. ingens</i>	UEP	[283]
Ingenine E 350 ^{β,η} [C ₁₆ H ₁₈ N ₃ O]	UV, IR, MS, NMR	Indole Alkaloid	Cytotoxic	A549, MCF7 HCT116 L5178Y HeLa, PC12	IC ₉₀ = 2.15 – 3.50 µg/mL IC ₉₀ = 0.67 µg/mL IC ₅₀ = 4.7 µg/mL NA	<i>A. ingens</i>	UEP	[284]
(-)-Acanthomine A 351 ^{β,η} [C ₁₅ H ₁₃ N ₅]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	<i>A. salina</i> A549, MCF7 HCT116	25% (10 µg/mL, 24 h) 50% (10 µg/mL, 48 h) IC ₅₀ = 1.92 – 2.81 µM IC ₅₀ = 0.59 µM	<i>A. ingens</i>	NSW	[285]
Variabine A 352 ^{β,η} [C ₁₃ H ₁₀ O ₆ N ₂ S]	UV, MS, NMR	Indole Alkaloid	Anticancer	Chymotrypsin-like activity Ubc13 (E2)–Uev1A	IC ₅₀ = 16 µM IC ₅₀ = 20 µM	<i>L. variabilis</i>	NSW	[286]
Variabine B 353 ^{β,η} [C ₁₃ H ₁₀ O ₆ N ₂ S]	UV, MS, NMR	Indole Alkaloid	Anticancer	Chymotrypsin-like activity, Ubc13 (E2)–Uev1A, E1, P53- Hdm2 (E3)	NA (16 – 20 µM)	<i>L. variabilis</i>	NSW	[286]
Trypargimine 354 ^{β,η} [C ₁₅ H ₁₉ N ₅]	UV, MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Eudistoma</i> sp.	SSW	[287]
1-Carboxytrypargine 355 ^{β,η} [C ₁₆ H ₂₁ N ₅ O ₂]	UV, MS, NMR, ECD	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Eudistoma</i> sp.	SSW	[287]
1,6-Dihydroxy-1,2,3,4- tetrahydro- β-Carboline 356d [C ₁₁ H ₁₂ N ₂ O ₂]	Undetm.	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Hyrtios</i> sp.	SSW	[279]
(+)-1,2,3,4- Tetrahydronorharman-1-one 357 ^{β,θ} [C ₁₁ H ₁₀ N ₂ O]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone), <i>L. donovani</i>	NA	A sponge (<i>Petrosiidae</i>)	NSW	[252, 284]

Table S10: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -1,2,3,4-Tetrahydronorharman-1-one 357 ^{β,θ} [C ₁₁ H ₁₀ N ₂ O]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Cytotoxic	V79 L5178Y HeLa, PC12	NA (4.76 µg/mL) IC ₅₀ > 10 µg/mL NA	A sponge (<i>Petrosiidae</i>)	NSW	[252, 284]
				<i>A. salina</i> A549, MCF7, HCT116 A549, MGC-803, Bel-7404, NCI-H460, HepG2	20% (10 µg/mL, 24 h) 40% (10 µg/mL, 48 h) IC ₅₀ = 7.45 – 10.11 µg/mL IC ₅₀ = 12.54 ± 1.17 – 33.46 ± 0.71 µM			
(+) -Hyrtioreticulin A 358 ^{β,θ} [C ₁₆ H ₁₆ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Anticancer	E1	IC ₅₀ = 2.4 µM	<i>H. reticulatus</i>	NSW	[288]
(-) -Hyrtioreticulin B 359 ^{β,θ} [C ₁₆ H ₁₆ N ₄ O ₃]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Anticancer	E1	IC ₅₀ = 35 µM	<i>H. reticulatus</i>	NSW	[288]
(-) -Hyrtioreticulin E 360 ^{β,θ} [C ₁₃ H ₁₄ N ₂ O ₃]	UV, IR, MS, NMR, [α] _D , ECD, CT	Indole Alkaloid	Anticancer	E1	NA (100 µM)	<i>H. reticulatus</i>	NSW	[288]
(+) -Hyrtioreticulin C 361 ^{β,θ} [C ₁₃ H ₁₄ N ₂ O ₃]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Anticancer	E1	NA (100 µM)	<i>H. reticulatus</i>	NSW	[288]
(-) -Hyrtioreticulin D 362 ^{β,θ} [C ₁₃ H ₁₄ N ₂ O ₃]	UV, IR, MS, NMR, [α] _D	Indole Alkaloid	Anticancer	E1	NA (100 µM)	<i>H. reticulatus</i>	NSW	[288]
3-Bromofascaplysin 363 ^{β,ι} [C ₁₈ H ₁₀ BrN ₂ O]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210 C38–CFU-GM HT116/H125–CEM HOP-62, COLO-205, U251, SK-MEL-5	-150 z.u. (6.4 µg/disk) 0 z.u. (6.4 µg/disk) 200/150 z.u. (6.4 µg/disk) NA	<i>F. reticulata</i>	UEP	[289 – 292]
				NCI-H23, NCI-H322M, NCI-H522, HCC-2998, HCT116, SF295, M14, OVCAR-4, SNB-19, MALME-3M, UACC-62, IGROV1, OVCAR-8, UO-31, HS 578T, BT-549	IC ₅₀ = 0.49 – 0.91 µM			
				RXF-393, CAKI-1, SN12C, OVCAR-3	IC ₅₀ = 1.6 – 4.4 µM			
				HL-60	35.8% apop. (0.25 µM), IC ₅₀ = 549 nM			

Table S10: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
3-Bromofascaplysin 363 ^{β,ι} [C ₁₈ H ₁₀ BrN ₂ O]	MS, NMR	Indole Alkaloid	Cytotoxic	THP-1, MDA-MB-231, SK-MEL-28 HeLa, DLD-1, SNU-C4, J86 P ⁺ C141	IC ₅₀ = 521 – 785 nM IC ₅₀ = 238 – 337 nM	<i>F. reticulata</i>	UEP	[289 – 292]
10-Bromofascaplysin 364 ^{β,ι} [C ₁₈ H ₁₀ BrN ₂ O]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210 C38–CFU-GM HT116/H125–CEM HT116/H125–CEM HL-60 J86 P ⁺ C141, THP-1 HeLa, MDA-MB-231, DLD-1, SNU-C4 SK-MEL-28	-100 z.u. (3.4 µg/disk) 100 z.u. (3.4 µg/disk) 200/350 z.u. (3.4 µg/disk) 200/300 z.u. (0.8 µg/disk) 36.1% apop. (0.25 µM), IC ₅₀ = 142 nM IC ₅₀ = 144 – 161 nM IC ₅₀ = 86 – 173 nM IC ₅₀ > 1000 nM	<i>F. reticulata</i>	CSW	[289 – 292]
3-Bromohomofascaplysins B-1 365 ^{β,ι} [C ₁₈ H ₉ Br ₂ N ₂ O]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i> , <i>Didemnum</i> sp.	CSW	[289]
Homofascaplysate A 366 ^{β,κ} [C ₂₀ H ₁₄ BrN ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i>	CSW	[292]
Homofascaplysin B-1 367 ^{β,κ} [C ₂₂ H ₁₆ N ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i>	CSW	[292]
3-Bromohomofascaplysin B 368 ^{β,κ} [C ₂₁ H ₁₃ BrN ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i>	CSW	[292]
3-Bromohomofascaplysin B-1 369 ^{β,κ} [C ₂₂ H ₁₃ BrN ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i> , <i>Didemnum</i> sp.	CSW	[292]
3-Bromohomofascaplysin C 370 ^{β,κ} [C ₁₉ H ₁₁ BrN ₂ O]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i> , <i>Didemnum</i> sp.	CSW	[292]
14-Bromoreticulatine 371 ^{β,κ} [C ₁₉ H ₁₄ BrN ₂ O ₂]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210 C38–CFU-GM HT116/H125–CEM	50 z.u. (200 µg/disk) 150 z.u. (200 µg/disk) -/- z.u. (200 µg/disk)	<i>F. reticulata</i>	UEP	[289, 292]

Table S10: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
14-Bromoreticulatate 372 ^{β,κ} [C ₁₈ H ₁₁ BrN ₂ O ₂]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210	150 z.u. (60 µg/disk)	<i>F. reticulata</i>	UEP	[289]
7,14-Dibromoreticulatine 373 ^{β,κ} [C ₁₉ H ₁₃ Br ₂ N ₂ O ₂]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210, C38–CFU-GM HT116/H125–CEM	-100 z.u. (84 µg/disk) 0/200 z.u. (84 µg/disk)	<i>F. reticulata</i> / <i>Didemnum</i> sp.	CSW	[292]
Reticulatol 374 ^{β,κ} [C ₁₇ H ₁₃ N ₂ O]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210 C38–CFU-GM	0 z.u. (64 µg/disk) -900/-600 z.u. (64 µg/disk)	<i>F. reticulata</i>	CSW	[292]
14-Bromoreticulatol 375 ^{β,κ} [C ₁₇ H ₁₂ BrN ₂ O]	MS, NMR	Indole Alkaloid	Cytotoxic	C38–L1210, C38–CFU-GM HT116/H125–CEM	-50 z.u. (84 µg/disk) -250/-/200 z.u. (84 µg/disk)	<i>F. reticulata</i>	CSW	[292]
3-Bromosecofascaplysin A 376 ^{β,κ} [C ₁₉ H ₁₃ BrN ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i>	CSW	[292]
3-Bromosecofascaplysin B 377 ^{β,κ} [C ₁₈ H ₁₂ N ₂ O ₃]	MS, NMR	Indole Alkaloid	Undetm.	Undetm.	Undetm.	<i>F. reticulata</i>	CSW	[292]

Footnote: 1. Structure (^κmolecule isolated as salt). **Activity** (C38 murine colon adenocarcinoma, CFU-GM murine colony-forming unit-granulocyte macrophage, J86 P⁺C141 murine skin epidermal, Bel-7404 human hepatocarcinoma, BT-549 human breast cancer, CAKI-1 human renal cancer, COLO-205 human colon carcinoma, CEM human leukemia, H125 human lung cancer, HCC-2998 human colon carcinoma, HCT15 human colon carcinoma, HCT116 human colon carcinoma, HOP-62 human non-small cell lung cancer, HT116 human colon cancer, HS 578T human breast cancer, IGROV1 human ovarian cancer, M14 human melanoma cancer, MALME-3M human melanoma cancer, MGC-803 human gastric cancer, NCI-H23 human non-small cell lung cancer, NCI-H322M human non-small cell lung cancer, NCI-H522 human non-small cell lung cancer, OVCAR-4 human ovarian cancer, OVCAR-8 human ovarian cancer, RXF-393 human renal cancer, SF-295 human CNS cancer, SK-MEL-5 human melanoma cancer, SK-MEL-28 human melanoma cancer, SNB-19 human CNS cancer, SN12C human renal cancer, SNU-C4 human adenocarcinoma, THP-1 human leukemia monocyte, U251 human CNS cancer, UACC-62 human melanoma cancer, UO-31 human renal cancer, E1 ubiquitin-inhibiting enzyme, FcB1 chloroquine-resistant strain, z.u. zone unit, **apop.** apoptosis).

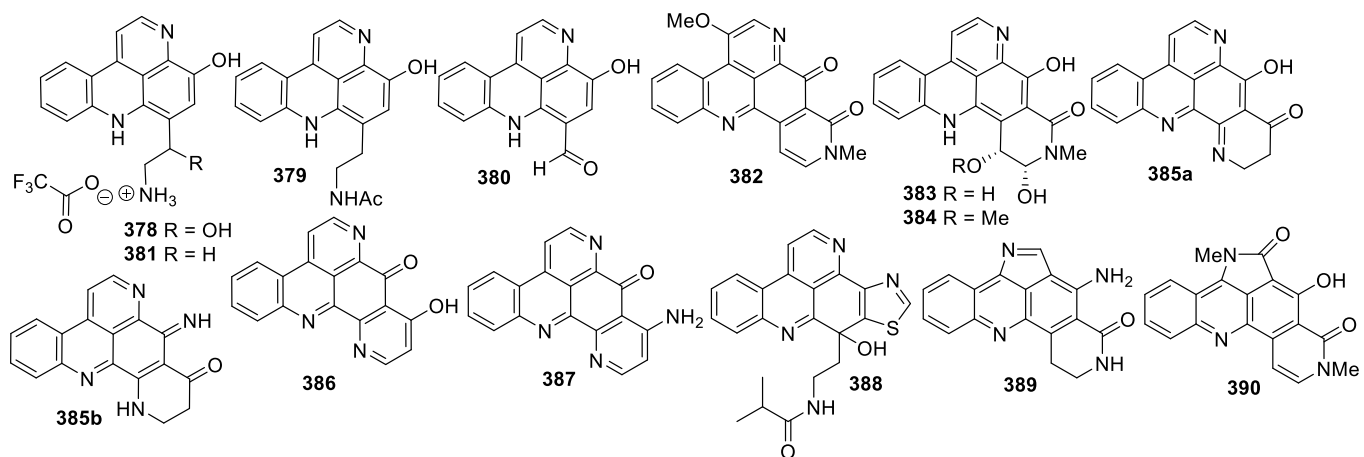


Figure S11: Structures of marine acridine alkaloids from Indonesian waters found in 1970–2017.

Table S11: Marine acridine alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Styelsamine A 378 ^{β,κ} [C ₁₇ H ₁₆ N ₃ O ₂]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic	HCT116	IC ₅₀ = 33 μM	<i>E. latericius</i>	SSW	[75 – 78, 293]
Styelsamine B 379 ^{β,κ} [C ₁₉ H ₁₇ N ₃ O ₂]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic	HCT116 Calf thymus DNA binding aff. Human tumor cells (57 cells) ^ν	IC ₅₀ = 89 μM $K_{app} = 5.33 \times 10^6 \text{ M}^{-1}$ GI ₅₀ = 3.2 μM	<i>E. latericius</i>	SSW	[75 – 78, 293]
Styelsamine C 380 ^{β,η} [C ₁₆ H ₁₀ N ₂ O ₂]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic Antibacterial	HCT116 <i>L. anguilarum</i>	IC ₅₀ = 2.6 μM MIC (micromolar level)	<i>E. latericius</i>	SSW	[75 – 78, 293]
Styelsamine D 381 ^{β,κ} [C ₁₇ H ₁₆ N ₃ O]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic	HCT116	IC ₅₀ = 1.6 μM	<i>E. latericius</i>	SSW	[75 – 78, 293]
5-Methoxyneomphimedine 382 ^{β,κ} [C ₂₀ H ₁₃ N ₃ O ₃]	MS, NMR	Acridine Alkaloid	Cytotoxic	C38–L1210 C38–CFU-GM HT116/H125–CEM	> 700 z.u. (25 μg/disk) > 800 z.u. (25 μg/disk) -250/-150 z.u. (25 μg/disk)	<i>X. cf. carbonaria</i>	UEP	[294]
Neoamphimedine Y 383 ^{β,κ} [C ₁₉ H ₁₅ N ₃ O ₄]	NMR	Acridine Alkaloid	Cytotoxic	Undetm.	Undetm.	<i>X. cf. carbonaria</i>	UEP	[294]
Neoamphimedine Z 384 ^{β,κ} [C ₂₁ H ₁₉ N ₃ O ₄]	NMR	Acridine Alkaloid	Cytotoxic	Undetm.	Undetm.	<i>X. cf. carbonaria</i>	UEP	[294]
Labuanine 385a ^{β,η} [C ₁₈ H ₁₁ N ₃ O ₂]	IR, MS, NMR, CT	Acridine Alkaloid	Anticancer	Neuro 2A	50% neuritogenesis (1 μM)	<i>B. fortis</i>	ENT	[295 – 297]
Ecionine A 385b ^{δ,κ} [C ₁₈ H ₁₂ N ₄ O]	UV, LCMS, MS, NMR	Acridine Alkaloid	Cytotoxic Anticancer Anticancer	K562 TSU-Pr1-B1, TSU-Pr1, TSU-Pr1-B2, 5637 K562 Neuro 2A	5 μg/mL IC ₅₀ = 6.48 – 6.49 μM IC ₅₀ = 3.55 – 3.66 μM 42% (25 ng/mL) 50% neuritogenesis (3 μM)	<i>Biemna</i> sp.	JPN	[296 – 297]
386 ^{β,η} [C ₁₈ H ₉ N ₃ O ₂]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic	P-388 A549, HT-29 MEL-28	IC ₅₀ = 4.18 μM IC ₅₀ = 0.03 – 0.40 μM IC ₅₀ = 0.17 μM	<i>B. fortis</i>	ENT	[295 – 297]

Table S11: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
387 ^{β,η} [C ₁₈ H ₁₀ N ₄ O]	UV, MS, NMR	Acridine Alkaloid	Anticancer	Neuro 2A	> 50% neuritogenesis (0.03 μM) ACHe and G2/M (0.03 μM, 48 h)	<i>B. fortis</i>	ENT	[295 – 297]
Sagitol C 388 ^{β,η} [C ₂₂ H ₂₀ N ₄ O ₂ S]	UV, MS, NMR	Acridine Alkaloid	Cytotoxic	L5178Y, HeLa PC12	ED ₅₀ = 0.7 – 0.9 μM ED ₅₀ = 2.3 μM	<i>Oceanapia sp.</i>	MLU	[298]
Plakinidine D 389 ^{β,η} [C ₁₇ H ₁₂ N ₄ O]	UV, MS, NMR, MS	Acridine Alkaloid	Cytotoxic	HCT116	5 μg/mL	<i>Didemnum sp.</i>	SSW	[299]
Alpkinidine 390 ^{β,λ} [C ₁₉ H ₁₃ N ₃ O ₃]	NMR, X-ray	Acridine Alkaloid*	Cytotoxic	C38–L1210, C38–CFU-GM C38–CFU-GM	300 z.u. (120 μg/disk) 300 z.u. (120 μg/disk)	X cf. <i>carbonaria</i>	UEP	[294]

Footnote: 1. **Structure** (^λmolecule isolated as HCO₂ salt); 2. Activity (^νactivity tested as freebase, **5637** human bladder carcinoma).

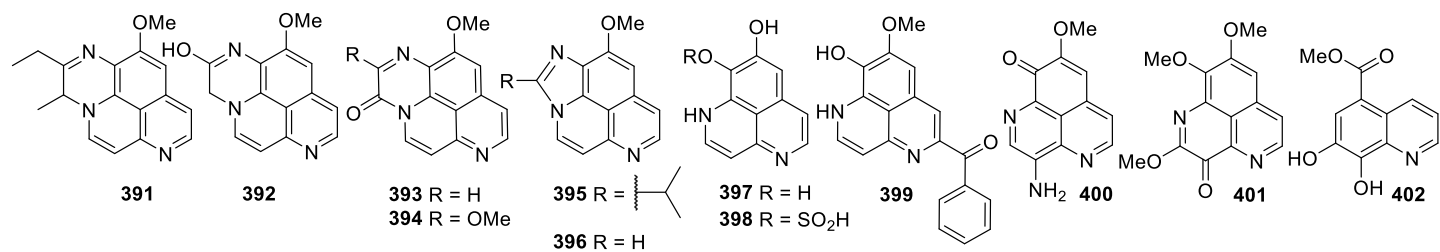


Figure S12: Structures of marine quinoline and isoquinoline alkaloids from Indonesian waters found in 1970–2017.

Table S12: Marine quinoline and isoquinoline alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
391^{β,η} [C ₁₇ H ₁₇ N ₃ O]	UV, IR, MS, NMR	Quinoline & Isoquinoline [▲]	Antibacterial	<i>S. aureus</i> , <i>E. coli</i> , <i>V. anguillarum</i>	MIC > 100 µg/mL	<i>Xestospongia</i> sp.	JSCR	[300]
			Antifungal	<i>C. tropicalis</i>	MIC > 100 µg/mL			
			Cytotoxic	KB	ID ₅₀ > 10 µg/mL			
392^{β,η} [C ₁₄ H ₁₁ N ₃ O ₂]	UV, IR, MS, NMR	Quinoline & Isoquinoline	Antibacterial	<i>S. aureus</i> , <i>E. coli</i> , <i>V. anguillarum</i>	MIC > 100 µg/mL	<i>Xestospongia</i> sp.	JSCR	[300]
			Antifungal	<i>C. tropicalis</i>	MIC > 100 µg/mL			
			Cytotoxic	KB	ID ₅₀ > 10 µg/mL			
11-Methoxy-3 <i>H</i> - [1,6]naphthyridino[6,5,4- <i>def</i>]quinoxalin-3-one 393^{β,η} [C ₁₄ H ₉ N ₃ O ₂]	UV, MS, NMR	Quinoline & Isoquinoline	Cytotoxic	L5178Y	NA (10 µg/mL)	<i>A. suberitoides</i>	MLU	[301]
2,11-Dimethoxy-3 <i>H</i> - [1,6]naphthyridino[6,5,4- <i>def</i>]- quinoxalin-3-one 394^{β,η} [C ₁₅ H ₁₁ N ₃ O ₃]	UV, MS, NMR	Quinoline & Isoquinoline	Cytotoxic	L5178Y	NA (10 µg/mL)	<i>A. suberitoides</i>	MLU	[301]
395^{β,η} [C ₁₆ H ₁₅ N ₃ O]	IR, MS, NMR	Quinoline & Isoquinoline [▲]	Antibacterial	<i>S. aureus</i> , <i>E. coli</i> , <i>V. anguillarum</i>	MIC > 100 µg/mL	<i>Xestospongia</i> sp.	JSCR	[300]
			Antifungal	<i>C. tropicalis</i>	MIC > 100 µg/mL			
			Cytotoxic	KB	ID ₅₀ > 10 µg/mL			
396^{β,η} [C ₁₃ H ₉ N ₃ O]	UV, IR, MS, NMR	Quinoline & Isoquinoline	Antibacterial	<i>S. aureus</i> , <i>E. coli</i> <i>M. smegmatis</i> (aerobic)	MIC > 100 µg/mL	<i>Xestospongia</i> sp.	JSCR	[300 – 302]
			Antifungal	<i>M. smegmatis</i> (hypoxic)	MIC = 25 µg/mL			
			Antifungal	<i>C. tropicalis</i>	MIC = 12.5 µg/mL			
			Cytotoxic	KB	MIC > 100 µg/mL			
			Cytotoxic	L5178Y	ID ₅₀ > 10 µg/mL IC ₅₀ = 13.5 µM			
Bisdemethylaaptamine 397^{ε,θ} [C ₁₁ H ₈ N ₂ O ₂]	UV, IR, NMR, MS	Quinoline & Isoquinoline	Antifungal	<i>C. neoformans</i> , <i>C. albicans</i>	MIC = 32 – 64 µg/mL	<i>Aaptos</i> sp.	NSW	[303, 304]
			Antibacterial	<i>S. aureus</i> , <i>S. pneumoniae</i> , <i>M. luteus</i>	MIC = 4 – 8 µg/mL			
			Antibacterial	<i>E. faecalis</i>	MIC = 8 – 16 µg/mL			

Table S12: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Bisdemethylaaptamine 397 ^{ζ,θ} [C ₁₁ H ₈ N ₂ O ₂]	UV, IR, NMR, MS	Quinoline & Isoquinoline	Antibacterial	<i>E. coli</i>	MIC = 16 – 64 µg/mL	<i>Aaptos</i> sp.	NSW	[302, 304]
			Antiparasite	<i>E. cloacae</i>	NA			
				<i>S. maltophilia</i>	MIC = 32 µg/mL			
				<i>N. gonorrhoeae</i>	MIC < 0.5 µg/mL			
Bisdemethylaaptamine-9-O-sulfate 398 ^{β,θ} [C ₁₁ H ₈ N ₂ O ₄ S]	UV, IR, NMR, MS	Quinoline & Isoquinoline	Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone)	NA	<i>Aaptos</i> sp.	NSW	[302, 304]
			Cytotoxic	P-388, BXPC-3, MCF7, SF268, NCI-H460, KM-20L2	EC ₅₀ = 0.12 – 0.80 µg/mL			
				DU-145	EC ₅₀ = 1.10 µg/mL			
			Undetm.	Undetm.	Undetm.			
5-Benzoyldemethyl aaptamine 399 ^{β,η} [C ₁₉ H ₁₄ N ₂ O ₃]	UV, NMR, MS	Quinoline & Isoquinoline	Cytotoxic	L5178Y	IC ₅₀ = 5.5 µM	<i>A. suberitoides</i>	MLU	[301]
			Antibacterial	<i>M. smegmatis</i> (aerobic)	MIC = 6.25 µg/mL			
				<i>M. smegmatis</i> (hypoxic)	MIC = 1.5 µg/mL			
3-Aminodemethyl (oxy)aaptamine 400 ^{β,η} [C ₁₂ H ₉ N ₃ O ₂]	UV, NMR, MS	Quinoline & Isoquinoline	Cytotoxic	L5178Y	64% (10 µg/mL)	<i>A. suberitoides</i>	MLU	[301, 302]
2-Methoxy-3-oxoaaptamine 401 ^{β,η} [C ₁₄ H ₁₂ N ₂ O ₄]	UV, NMR, MS	Quinoline & Isoquinoline	Antibacterial	<i>M. smegmatis</i> (aerobic), <i>M. smegmatis</i> (hypoxic)	MIC = 6.25 µg/mL	<i>Aaptos</i> sp.	ENT	[302]
Aaptoline 402 ^{β,θ} [C ₁₁ H ₉ NO ₄]	UV, IR, MS, NMR	Quinoline & Isoquinoline	Undetm.	Undetm.	Undetm.	<i>A. suberitoides</i>	NSW	[305]

Footnote: 1. Statistic (ID₅₀ 50% of the infective dose); 2. Activity (BXPC-3 human pancreas adenocarcinoma, DU-145 human prostate cancer, KM-20L2 human colon carcinoma).

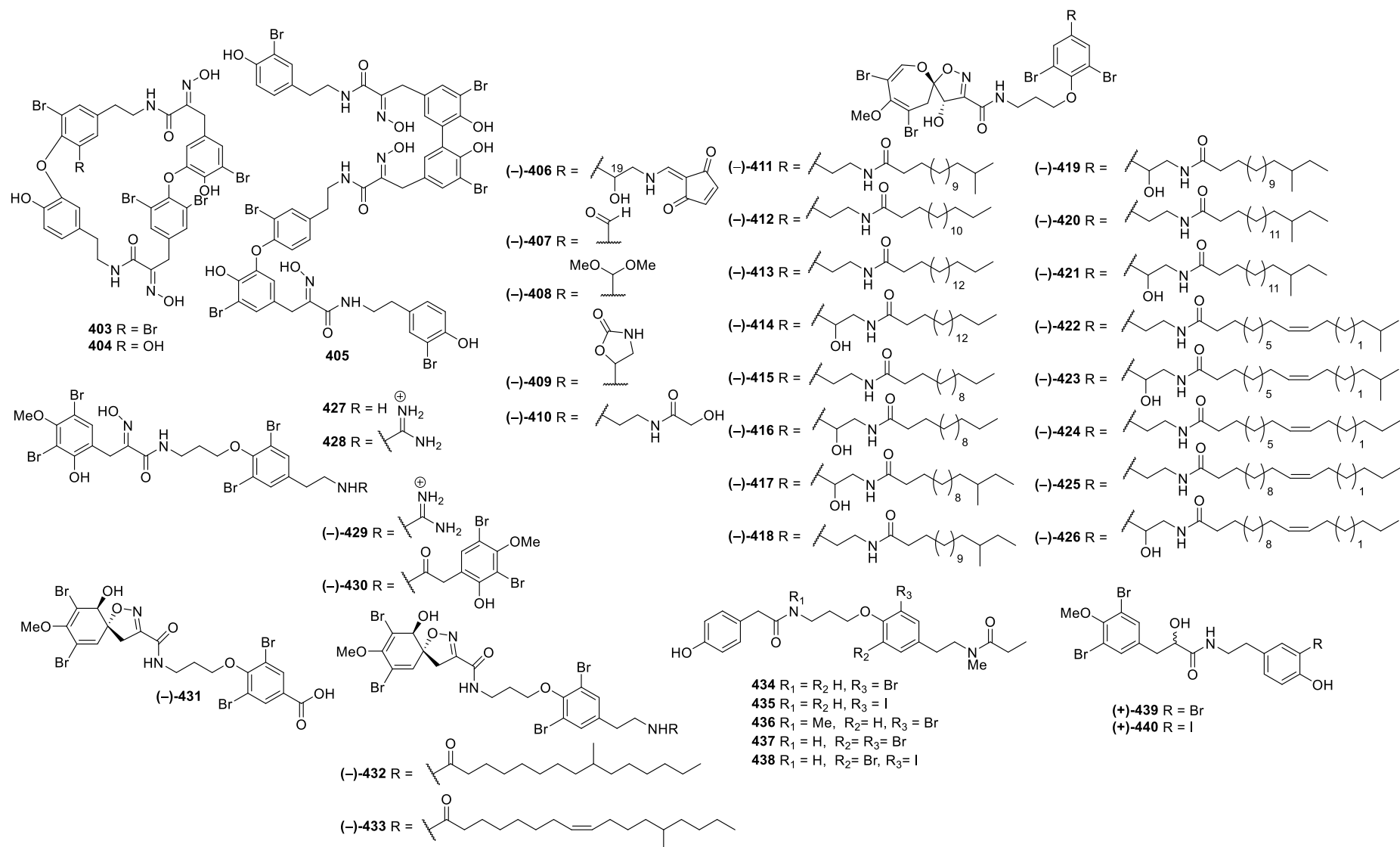


Figure S13: Structures of marine tyrosine alkaloids from Indonesian waters found in 1970–2017.

Table S13: Marine tyrosine alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Bastadin 16 403 ^{β,η} [C ₃₄ H ₂₇ Br ₅ N ₄ O ₈]	UV, IR, MS, NMR	Tyrosine Alkaloid	Cytotoxic	Human Sup-T1 cancer cell MCF7, SK-MEL-28, A549, HS683, B16F10, U373 L5178Y	IC ₅₀ = 1 × 10 ⁻¹² IC ₅₀ = 4 – 11 μM IC ₅₀ = 1.9 μM	<i>I. basta</i>	NSW	[306 – 310]
			Anticancer	Aur-A, Aur-B, EGF-R, CDK4/CycD1, ERBB2	IC ₅₀ = 1.3 – 2.9 μM IC ₅₀ = 4.0 – 4.5 μM			
			Antifouling Toxicity	<i>B. improvisus</i> <i>B. improvisus</i>	10 μM 10 μM			
			Alzheimer	Apolipoprotein E modulatory activity	NA (40 μM)			
Bastadin 17 404 ^{β,η} [C ₃₄ H ₂₈ Br ₄ N ₄ O ₉]	UV, IR, MS, NMR	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>I. basta</i>	NSW	[306]
			Cytotoxic	L5178Y ARK5, B-RAF VE, Aur-A, Aur-B, CDK4/CycD1, FLT3, INS-R, COT, ERBB2, VEGF-R3	NA IC ₅₀ = 1.3 – 2.6 μM			
Sesquibastadin 1 405 ^{β,η} [C ₅₁ H ₄₄ Br ₆ O ₁₂]	UV, IR, MS, NMR	Tyrosine Alkaloid*	Anticancer	CDK2/CycA, PLK1 EGF-R, MET, IGF1-R, SAK, SRC, TIE2, VEGF-R2 EPHB4, FAK, PDGFR-β	IC ₅₀ = 6.4 – 6.5 μM IC ₅₀ = 0.6 – 1.0 μM IC ₅₀ = 3.4 – 4.0 μM	<i>I. basta</i>	MLU	[311]
(-)-19-Hydroxy psammaplysin E 406 ^{α,β} [C ₂₇ H ₂₅ Br ₄ N ₃ O ₉]	MS, NMR, [α] _D , CT	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	IC ₅₀ = 6.4 ± 1.4 μM	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin K 407 ^β [C ₂₀ H ₁₈ Br ₄ N ₂ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin K dimethoxy acetal 408 ^β [C ₂₄ H ₂₄ Br ₄ N ₂ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin L 409 ^β [C ₂₂ H ₂₁ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin M 410 ^β [C ₂₃ H ₂₅ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]

Table S13: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Psammaplysin N 411 ^β [C ₃₇ H ₅₃ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin O 412 ^β [C ₃₇ H ₅₃ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin P 413 ^β [C ₃₉ H ₅₇ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin P 414 ^{α,β} [C ₃₉ H ₅₇ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin Q 415 ^β [C ₃₅ H ₄₉ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin Q 416 ^{α,β} [C ₃₅ H ₄₉ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin R 417 ^β [C ₃₇ H ₅₃ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin S 418 ^β [C ₃₈ H ₅₅ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin S 419 ^{α,β} [C ₃₈ H ₅₅ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin T 420 ^β [C ₄₀ H ₅₉ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin T 421 ^{α,β} [C ₄₀ H ₅₉ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin U 422 ^β [C ₃₈ H ₅₃ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin U 423 ^{α,β} [C ₃₈ H ₅₃ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-Psammaplysin V 424 ^β [C ₃₇ H ₅₁ Br ₄ N ₃ O ₇]	MS, NMR, [α] _D	Tyrosine Alkaloid	Antiparasite	<i>P. falciparum</i> (3D7)	NA (10 μM)	<i>A. strongylata</i>	BLI	[312]

Table S13: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Psammaplysin W 425 ^β [C ₄₀ H ₅₇ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
(-)-19-Hydroxy psammaplysin W 426 ^{α,β} [C ₄₀ H ₅₇ Br ₄ N ₃ O ₈]	MS, NMR, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	<i>A. strongylata</i>	BLI	[312]
Purpuramine M 427 ^β [C ₂₁ H ₂₃ Br ₄ N ₃ O ₅]	UV, IR, NMR, MS	Tyrosine Alkaloid	Alzheimer Cytotoxic	BACE1 A2780S, A2780S CP5, U251MG	36% (42 μM) IC ₅₀ = 20 – 50 μM	A sponge (Aplysi-nellidae)	EKM	[313]
Purpuramine N 428 ^{β,ι} [C ₂₂ H ₂₆ Br ₄ N ₅ O ₅]	UV, IR, NMR, MS	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	A sponge (Aplysi-nellidae)	EKM	[313]
(-)-Araplysillin VII 429 ^{β,κ} [C ₂₂ H ₂₆ Br ₄ N ₅ O ₅]	UV, IR, NMR, MS, [α] _D	Tyrosine Alkaloid	Alzheimer Cytotoxic	BACE1 A2780S, A2780S CP5, U251MG, A549, MCF7	40% (39.6 μM) Active > 50 μM	A sponge (Aplysi-nellidae)	EKM	[313]
(-)-Araplysillin VIII 430 ^β [C ₃₀ H ₂₉ Br ₆ N ₃ O ₈]	UV, IR, NMR, MS, [α] _D	Tyrosine Alkaloid	Undetm.	Undetm.	Undetm.	A sponge (Aplysi-nellidae)	EKM	[313]
(-)-Araplysillin IX 431 ^β [C ₂₀ H ₁₈ Br ₄ N ₂ O ₇]	UV, IR, NMR, MS, [α] _D	Tyrosine Alkaloid	Alzheimer Cytotoxic	BACE1 A2780S, A2780S CP5, U251MG, A549, MCF7	35% (41.9 μM) > 50 μM	A sponge (Aplysi-nellidae)	EKM	[313]
(-)-Araplysillin X 432 ^β [C ₃₇ H ₅₃ Br ₄ N ₃ O ₆]	UV, IR, NMR, MS, [α] _D	Tyrosine Alkaloid	Alzheimer Cytotoxic	BACE1 A2780S, A2780S CP5, U251MG, A549, MCF7	70% (31.4 μM) > 50 μM	A sponge (Aplysi-nellidae)	EKM	[313]
(-)-Araplysillin XI 433 ^β [C ₃₉ H ₅₆ Br ₄ N ₃ O ₆]	UV, IR, NMR, MS, [α] _D	Tyrosine Alkaloid	Alzheimer Cytotoxic	BACE1 A2780S, A2780S CP5, U251MG, A549, MCF7	60% (30.6 μM) > 50 μM	A sponge (Aplysi-nellidae)	EKM	[313]
Enisorine A 434 ^β [C ₂₃ H ₂₉ BrN ₂ O ₄]	UV, IR, MS, NMR	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	60 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]
Enisorine B 435 ^β [C ₂₃ H ₂₉ IN ₂ O ₄]	UV, IR, MS, NMR	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	120 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]
Enisorine C 436 ^β [C ₂₄ H ₃₁ BrN ₂ O ₄]	UV, IR, MS, NMR	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	30 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]
Enisorine D 437 ^β [C ₂₃ H ₂₈ Br ₂ N ₂ O ₄]	UV, IR, MS, NMR	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	120 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]

Table S13: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Enisorine E 438 ^β [C ₂₃ H ₂₈ BrIO ₄ N ₂]	UV, IR, MS, NMR	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	30 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]
(+)-1-O-Methyl hemi-bastadinol 2 439 ^β [C ₁₈ H ₁₈ Br ₃ NO ₄]	UV, IR, MS, NMR, [α] _D	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	30 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]
(+)-1-O-Methyl hemi-bastadinol 4 440 ^β [C ₁₈ H ₁₈ Br ₂ INO ₄]	UV, IR, MS, NMR, [α] _D	Tyrosine Alkaloid	Antibacterial	<i>Y. pseudotuberculosis</i>	60 μM (> 50% type III secretion system, T3SS)	<i>I. cf. iota</i>	CSW	[314]

Footnote: 1. Activity (**B16F10** murine melanoma cancer, **A2780S** human ovarian carcinoma, **CP5** human ovarian carcinoma cisplatin resistant, **HS683** human oligodendroglioma, **Sup-T1** human T cell lymphoma, **U251MG** human glioma, **U373** human astroglioma, **3D7** *Plasmodium falciparum* chloroquine-sensitive strain).

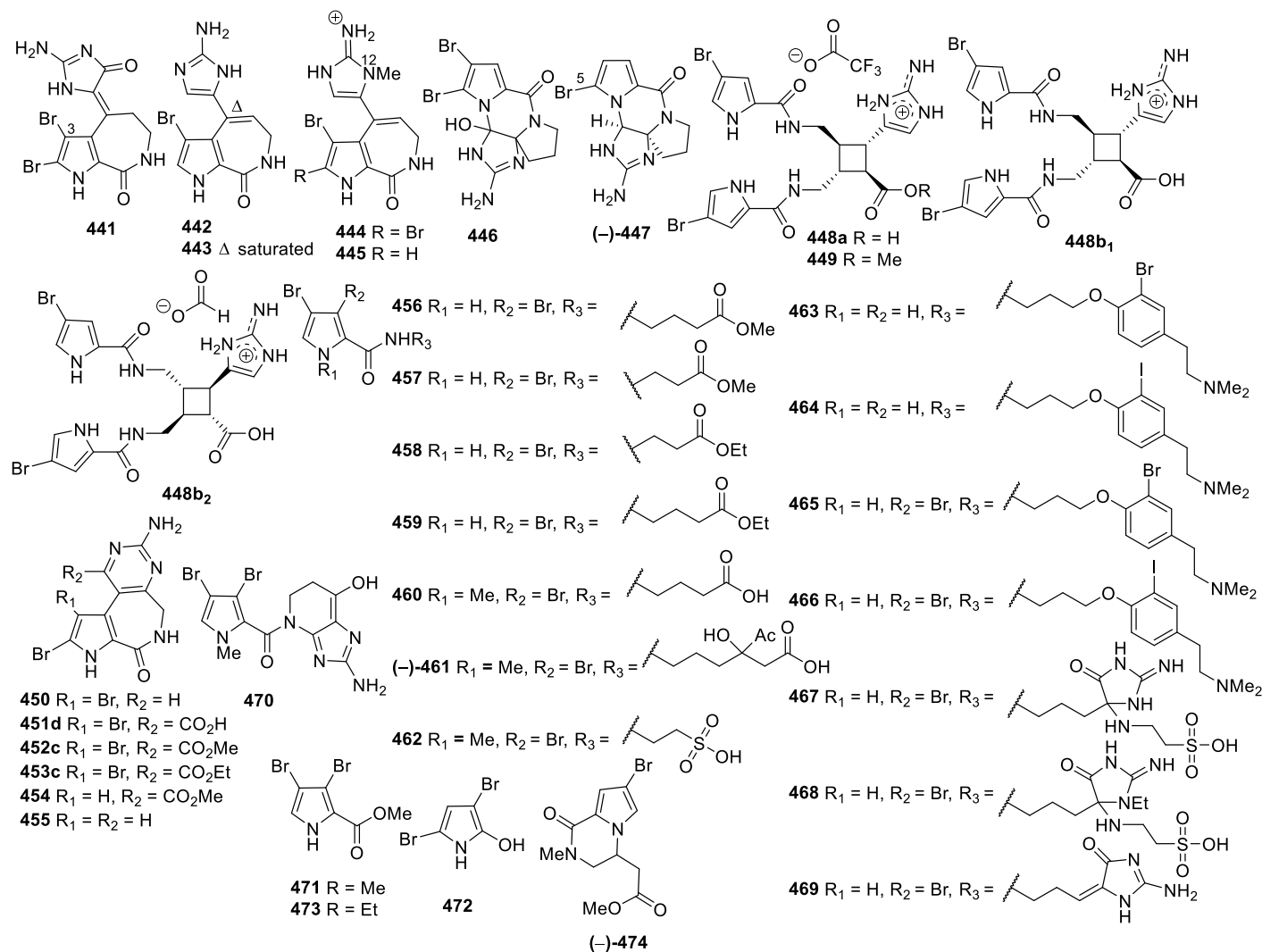


Figure S14: Structures of marine pyrrole alkaloids from Indonesian waters found in 1970–2017.

Table S14: Marine pyrrole alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(Z)-3-Bromohymenialdisine 441 ^β [C ₁₁ H ₉ Br ₂ N ₅ O ₂]	UV, NMR, MS	Pyrrole Alkaloid	Antiinsecticidal	<i>S. littoralis</i>	NA (300 µg/mL)	<i>A. carteri</i>	UEP	[315, 316]
			Cytotoxic	L5178Y	ED ₅₀ = 3.9 µg/mL 60.5% (10 µg/mL)			
			Anticancer	Aurora-A, CDK4-CycD1, FAK, VEGF-R2, SAK, PDGFR-β	20% < residual kinase activity ≤ 60%			
				Aurora-B, SRC, COT, PLK1	60% < residual kinase activity ≤ 80%			
Debromostevensine 442 ^{β,θ} [C ₁₁ H ₁₀ BrN ₅ O]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	MONO-MAC 6 L5178Y	NA 7.5% (10 µg/mL)	<i>S. carteri</i>	MLU	[317]
Debromohymenin 443 ^{β,θ} [C ₁₁ H ₁₂ BrN ₅ O]	UV, ECD, NMR, MS	Pyrrole Alkaloid	Cytotoxic	MONO-MAC 6	NA	<i>S. carteri</i>	MLU	[317]
12-N-Methyl stevensine 444 ^{β,θ} [C ₁₂ H ₁₂ Br ₂ N ₅ O]	IR, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	EC ₅₀ = 3.5 µg/mL IC ₅₀ = 8.75 µM	<i>Stylissa</i> sp.	EKM	[318]
			Anticancer	CLK-1, CDK5, CK-1 Cdk9/cyclin T, GSK-3, CDK1 Cdk2/A	IC ₅₀ = 4.1 – 6 µM IC ₅₀ = 1.3 – 3.2 µM IC ₅₀ > 10 µM			
12-N-Methyl-2-debromostevensine 445 ^{β,η} [C ₁₂ H ₁₃ BrN ₅ O]	IR, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	8.1% (10 µg/mL)	<i>Stylissa</i> sp.	EKM	[318]
(-)-Dibromohydroxy phakellin 446 ^{β,η} [C ₁₁ H ₁₁ Br ₂ N ₅ O ₂]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Cytotoxic	L5178Y	112.8% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
(-)-5-Bromophakelline 447 ^{β,θ} [C ₁₁ H ₁₂ BrN ₅ O]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Antibacterial	<i>S. epidermidis</i>	MIC > 45 µM			
(-)-Nakamuric acid 448a ^{β,θ} [C ₂₀ H ₂₁ Br ₂ N ₈ O ₃]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid*	Antibacterial	<i>B. subtilis</i> 168	9 mm (0.2 µmol/disk)	<i>A. nakamurai</i>	MLU	[80]
				<i>S. aureus</i> ATCC25923, ATCC43300	MIC = 16 µg/mL			
(-)-Nakamuric acid 448b ^{γ,Δ} [C ₂₀ H ₂₂ Br ₂ N ₇ O ₄]	TS	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.			[34]

Table S14: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Nakamuric acid 448b ^{2,λ} [C ₂₀ H ₂₂ Br ₂ N ₇ O ₄]	UV, MS, NMR, [α] _D , ECD	Pyrrole Alkaloid	Cytotoxic	PC9	IC ₅₀ > 10 µg/mL	<i>Agelas</i> sp.	PRC	[34]
(-)-Nakamuric acid methyl ester 449 ^β [C ₂₁ H ₂₄ Br ₂ N ₇ O ₄]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Antibacterial	<i>B. subtilis</i> 168	9 mm (0.2 µmol/disk)	<i>A. nakamurai</i>	MLU	[80]
Latonduine A 450 ^{β,η} [C ₁₀ H ₇ Br ₂ N ₅ O]	UV, MS, NMR, CT, TS	Pyrrole Alkaloid*	Cytotoxic	L5178Y	EC ₅₀ = 9.0 µg/mL IC ₅₀ = 26.81 µM	<i>S. carteri</i>	SSW	[45, 317, 320, 321]
			Cystic fibrosis	BHK (wild type)	45% F508del-CTFR corr. (1 µM, 24 h)			
				Salivary secretion <i>in vivo</i> assay in F508del-CFTR homo. mice	9% F508del-CTFR corr. (50 mg/kg, 2 days)			
				<i>Ex vivo</i> mouse assay (intestinal ileal epithelia from F508del-CFTR homo. mice) PARP-3	2.5% F508del-CTFR corr. (10 µM, 4 h) EC ₅₀ = 400 pM			
Latonduine B 451d ^{β,η} [C ₁₁ H ₇ Br ₂ N ₅ O ₃]	Undetm.	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>S. carteri</i>	SSW	[45]
Latonduin B methyl ester 452c ^{ε,η} [C ₁₂ H ₉ Br ₂ N ₅ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	1.7% (10 µg/mL)	<i>S. carteri</i>	SSW	[45, 317, 320, 321]
			Cystic fibrosis	BHK (wild type)	15% F508del-CTFR corr. (1 µM, 24 h)			
			Cytotoxic	L5178Y	1.7% (10 µg/mL)			
Latonduin B ethyl ester 453c ^{ε,η} [C ₁₃ H ₁₁ Br ₂ N ₅ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Anticancer	CLK-1, CDK5, GSK-3, DYRK1A, CK-1, CDK1, Cdk9/cyclin T	IC ₅₀ > 10 µM	<i>S. carteri</i>	SSW	[45, 317, 320, 321]
			Cystic fibrosis	BHK (wild type)	30% F508del-CTFR corr. (1 µM, 24 h)			
3-Debromolatonduine B methyl ester 454 ^{β,η} [C ₁₂ H ₁₀ BrN ₅ O ₃]	IR, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	10.2% (10 µg/mL)	<i>Stylissa</i> sp.	EKM	[317, 321]

Table S14: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
3-Debromo latonduine A 455 ^{β,η} [C ₁₀ H ₈ BrN ₅ O]	IR, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	6.6% (10 µg/mL)	<i>Stylissa</i> sp.	EKM	[317, 321]
			Anticancer	CLK-1, DYRK1A GSK-3, CK-1 CDK1, CDK2/A, CDK5, CDK9/cyclin T	IC ₅₀ = 1.7 – 2 µM IC ₅₀ = 0.21 – 0.78 µM IC ₅₀ > 10 µM			
Acanthamide A 456 ^{β,λ} [C ₁₀ H ₁₂ Br ₂ N ₂ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
Acanthamide B 457 ^{β,λ} [C ₉ H ₁₀ Br ₂ N ₂ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
Acanthamide C 458 ^{β,λ} [C ₁₀ H ₁₂ Br ₂ N ₂ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
Acanthamide D 459 ^{β,λ} [C ₁₁ H ₁₄ Br ₂ N ₂ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
4-(4,5-dibromo-1-methyl pyrrole-2-carboxamido)-butanoic acid 460 ^{β,η} [C ₁₀ H ₁₂ Br ₂ N ₂ O ₃]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	107.5% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 54 µM			
(-)-Agelatin B 461 ^{β,η} [C ₁₄ H ₁₈ Br ₂ N ₂ O ₅]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Cytotoxic	L5178Y	92.8% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 44 µM			
Mauritamide D 462 ^{β,η} [C ₈ H ₁₀ Br ₂ N ₂ O ₄ S]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	117.6% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 52 µM			
Agelanesin A 463 ^{β,η} [C ₁₈ H ₂₃ Br ₂ N ₃ O ₂]	UV, NMR, MS	Pyrrole Alkaloid	Cytotoxic	L5178Y	IC ₅₀ = 9.55 µM	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 42 µM			
Agelanesin B 464 ^{β,η} [C ₁₈ H ₂₃ BrIN ₃ O ₂]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	IC ₅₀ = 9.25 µM	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 38 µM			
Agelanesin C 465 ^{β,η} [C ₁₈ H ₂₂ Br ₃ N ₃ O ₂]	UV, NMR, MS	Pyrrole Alkaloid	Cytotoxic	L5178Y	IC ₅₀ = 16.76 µM	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 36 µM			
Agelanesin D 466 ^{β,η} [C ₁₈ H ₂₂ Br ₂ IN ₃ O ₂]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	IC ₅₀ = 13.06 µM	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 33 µM			
(-)-Mauritamide B 467 ^{β,η} [C ₁₃ H ₁₈ Br ₂ N ₆ O ₅ S]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Cytotoxic	L5178Y	92.5% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 37 µM			
(-)-Mauritamide C 468 ^{β,η} [C ₁₅ H ₂₂ Br ₂ N ₆ O ₅ S]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Cytotoxic	L5178Y	98.6% (23 mM)	<i>A. linnaei</i>	JSCR	[125]
			Antibacterial	<i>S. epidermidis</i>	MIC > 35 µM			

Table 14. Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Dispacamide E 469 ^{β,η} [C ₁₁ H ₁₁ Br ₂ N ₅ O ₂]	MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	27.2% (10 µg/mL)	<i>A. linnaei</i>	PUA	[321]
Agelatin A 470 ^{β,η} [C ₁₂ H ₁₁ Br ₂ N ₅ O ₂]	UV, MS, NMR	Pyrrole Alkaloid	Cytotoxic Antibacterial	L5178Y <i>S. epidermidis</i>	98.9% (23 mM) MIC > 48 µM	<i>A. linnaei</i>	JSCR	[125]
Methyl 3,4-dibromo-1H-pyrrole-2-carboxylate 471 ^β [C ₆ H ₅ Br ₂ NO ₂]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
3,5-Dibromo-1H-pyrrole-2-carboxylic acid 472 ^{β,η} [C ₄ H ₃ Br ₂ NO]	UV, MS, NMR	Pyrrole Alkaloid	Undetm.	Undetm.	Undetm.	<i>Acanthostylotella</i> sp.	BLI	[322]
Ethyl 3,4-dibromo-1H-pyrrole-2-carboxylate 473 ^{β,η} [C ₇ H ₇ Br ₂ NO ₂]	MS, NMR	Pyrrole Alkaloid	Cytotoxic	L5178Y	77.2% (10 µg/mL)	<i>S. massa</i>	PUA	[321]
(-)-Longamide C 474 ^{β,η} [C ₁₁ H ₁₃ BrN ₂ O ₃]	UV, MS, NMR, [α] _D	Pyrrole Alkaloid	Cytotoxic	Undetm.	Undetm.	<i>A. nakamurai</i>	JSCR	[125]

Footnote: 1. Activity (MONO-MAC 6 human monocytic leukemia cells, PC9 human lung cancer, BHK baby hamster kidney cells, **Aurora-A** serine/threonine-protein kinase, **Aurora-B** serine/threonine-protein kinase, **CDK2** cyclin-dependent kinase 2, **CDK4** cyclin-dependent kinase 4, **CDK5** cyclin-dependent kinase 5, **CDK9** cyclin-dependent kinase 9, **CK-1** creatine kinase 1, **COT** cancer Osaka thyroid oncogene, **DYRK1A** dual specificity tyrosine-phosphorylation regulated kinase 1A, **GSK-3** glycogen synthase kinase 3, **SRC** non-receptor tyrosine kinase, **PARP-3** poly(ADP-ribose) polymerase 3, **PDGFR** platelet-derived growth factor receptor, **SAK** serine/threonine-protein kinase, **CTFR** cystic fibrosis transmembrane conductance regulator, **corr.** correction).

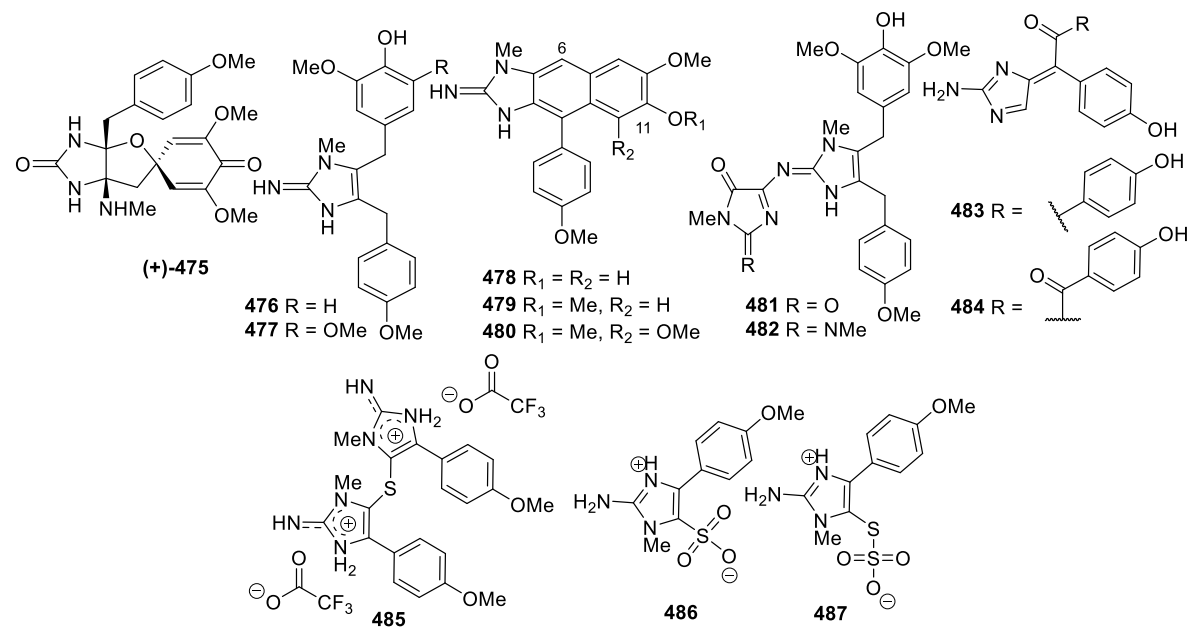


Figure S15: Structures of marine imidazole alkaloids from Indonesian waters found in 1970–2017.

Table S15: Marine imidazole alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Spironaamidine 475 ^{β,η} [C ₂₁ H ₂₅ N ₃ O ₆]	MS, NMR, [α] _D	Imidazole Alkaloid*	Antibacterial	<i>B. cereus</i>	12 mm (10 mg/disk)	<i>L. microraphis</i>	NSW	[323]
Naamine F 476 ^{β,θ} [C ₂₀ H ₂₃ N ₃ O ₃]	UV, MS, NMR	Imidazole Alkaloid	Undetm.	Undetm.	Undetm.	<i>L. chagosensis</i>	SSW	[324]
Naamine G 477 ^{β,θ} [C ₂₁ H ₂₅ N ₃ O ₄]	UV, MS, NMR	Imidazole Alkaloid	Cytotoxic	L5178Y, HeLa PC12	29 – 46% (10 µg/mL) NA	<i>L. chagosensis</i>	SSW	[324, 325]
Kealiinine A 478 ^{β,η} [C ₂₀ H ₁₉ N ₃ O ₃]	UV, MS, NMR	Imidazole Alkaloid	Antifungal Cytotoxic Antifungal	<i>A. salina</i> <i>C. herbarum</i> <i>A. salina</i> <i>C. herbarum</i>	10% (10 µg/mL) 20 mm (20 µg/disk) 50% (10 µg/mL) NA (20 µg)	<i>L. chagosensis</i>	SSW	[324, 326]
Kealiinine B 479 ^{β,η} [C ₂₁ H ₂₁ N ₃ O ₃]	UV, MS, NMR	Imidazole Alkaloid	Cytotoxic	MCF10A, MCF7, T47D, MDA-MB-231	IC ₅₀ = 10.0 – 13.3 µM	<i>L. chagosensis</i>	SSW	[324, 326, 327]
Kealiinine C 480 ^{β,η} [C ₂₂ H ₂₃ N ₃ O ₄]	UV, MS, NMR	Imidazole Alkaloid	Cytotoxic	MCF10A, MCF7, T47D, MDA-MB-231	IC ₅₀ > 50 µM	<i>L. chagosensis</i>	SSW	[324, 326, 327]
Naamidine H 481 ^{β,η} [C ₂₅ H ₂₇ N ₅ O ₆]	UV, IR, MS, NMR	Imidazole Alkaloid	Cytotoxic Antibacterial	HeLa <i>E. coli</i>	IC ₅₀ = 5.6 µg/mL NA (50 µg/disk)	<i>L. chagosensis</i>	NSW	[328]
Naamidine I 482 ^{β,η} [C ₂₆ H ₃₀ N ₆ O ₅]	UV, IR, MS, NMR	Imidazole Alkaloid	Cytotoxic Antibacterial	<i>B. cereus</i> HeLa <i>E. coli</i>	8 mm (10 mg/disk) IC ₅₀ = 15 µg/mL NA (50 µg/disk)	<i>L. chagosensis</i>	NSW	[328]
Lissodendrin A 483 ^{β,η} [C ₁₇ H ₁₃ N ₃ O ₃]	UV, MS, NMR	Imidazole Alkaloid*	Cytotoxic	L5178Y	NA (10 µg/mL)	<i>Lisso-dendoryx (Acantho-doryx) fibrosa</i>	MLU	[81]
Lissodendrin B 484 ^{β,η} [C ₁₇ H ₁₃ N ₃ O ₄]	UV, MS, NMR	Imidazole Alkaloid*	Cytotoxic	L5178Y	NA (10 µg/mL)	<i>Lisso-dendoryx (Acantho-doryx) fibrosa</i>	MLU	[81]

Table S15: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Polycarpaurine A 485 ^{B,θ} [C ₂₂ H ₂₆ N ₆ O ₂ S ₂]	UV, IR, MS, NMR	Imidazole Alkaloid [•]	Cytotoxic	V79	EC ₅₀ = 6.8 μM	<i>P. aurata</i>	NSW	[329, 330]
			Antiviral ^u	TMV	NA (100 μg/mL)			
			<i>in vitro</i>		14 ± 2% (500 μg/mL)			
					28 ± 2% inactivation (500 μg/mL)			
			Antiviral ^u	TMV	33 ± 2% cur. (500 μg/mL)			
Polycarpaurine B 486 ^{B,θ} [C ₂₂ H ₂₆ N ₆ O ₂ S ₂]	UV, IR, MS, NMR	Imidazole Alkaloid	<i>in vivo</i>		24 ± 2% protection (500 μg/mL)			
				<i>F. oxysporium f.sp.cucumeris</i> , <i>C. arachidicola</i> Hori, <i>P. piricola</i> , <i>A. solani</i> , <i>F. graminearum</i> , <i>F. moniliforme</i> , <i>S. sclerotiorum</i> , <i>P. capsici</i> , <i>R. cerealis</i> , <i>B. maydis</i> , <i>W. Anthracnose</i> , <i>P. infestans</i> , <i>R. solani</i> , <i>B. cinerea</i>	6 ± 2 – 48 ± 1% (50 mg/kg)			
			Antifungal					
			<i>in vitro</i> ^u					
				<i>P. capsici</i> , <i>R. cerealis</i> , <i>B. graminis f. sp. tritici</i> , <i>S. sclerotiorum</i> , <i>R. solani</i> , <i>B. cinerea</i> , <i>C. cassiicola</i>	8 ± 2 – 30 ± 1% (200 mg/kg)			
Polycarpaurine C 487 ^{B,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid [•]	Antifungal			<i>P. aurata</i>	NSW	[329, 330]
			<i>in vivo</i> ^u					
Polycarpaurine C 487 ^{B,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid [•]	Cytotoxic	V79	EC ₅₀ > 10 μM	<i>P. aurata</i>	NSW	[329, 330]
			Cytotoxic	V79	EC ₅₀ = 8.6 μM			
Polycarpaurine C 487 ^{B,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid [•]			NA (100 μg/mL)	<i>P. aurata</i>	NSW	[329, 330]
			Antiviral	TMV	23 ± 2% (500 μg/mL)			
Polycarpaurine C 487 ^{B,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid [•]	<i>in vitro</i>		30 ± 2% inactivation (500 μg/mL)	<i>P. aurata</i>	NSW	[329, 330]
					15 ± 1% cur. (500 μg/mL)			
Polycarpaurine C 487 ^{B,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid [•]			34 ± 2% protection (500 μg/mL) (<i>in vivo</i>)			

Table S15: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Polycarpaurine C 487 ^{b,θ} [C ₁₁ H ₁₃ N ₃ O ₄ S ₂]	UV, IR, MS, NMR	Imidazole-Alkaloid*	Antifungal <i>in vitro</i>	<i>F. oxysporium f.sp.cucumeris</i> , <i>C. arachidicola</i> Hori, <i>P. piricola</i> , <i>A. solani</i> , <i>F. graminearum</i> , <i>F. moniliforme</i> , <i>S. sclerotiorum</i> , <i>P. capsici</i> , <i>R. cerealis</i> , <i>R. solani</i> , <i>B. maydis</i> , <i>W. Anthracnose</i> , <i>P. infestans</i> , <i>B. cinerea</i>	2 ± 2 – 40 ± 2% (50 mg/kg)	<i>P. aurata</i>	NSW	[329, 330]
			Antifungal <i>in vivo</i>	<i>P. capsici</i> , <i>R. cerealis</i> , <i>B. graminis f. sp. tritici</i> , <i>S. sclerotiorum</i> , <i>R. solani</i> , <i>B. cinerea</i> , <i>C. cassiicola</i>	0 – 30 ± 2% (200 mg/kg)			
				TMV	34 ± 2% protection (500 µg/mL) (<i>in vivo</i>)			

Footnote: 1. Activity (^aActivity tested chloride salt, **MCF10A** human normal breast cell, **T47D** human breast adenocarcinoma, **TMV** tobacco mosaic virus, **cur.** curative).

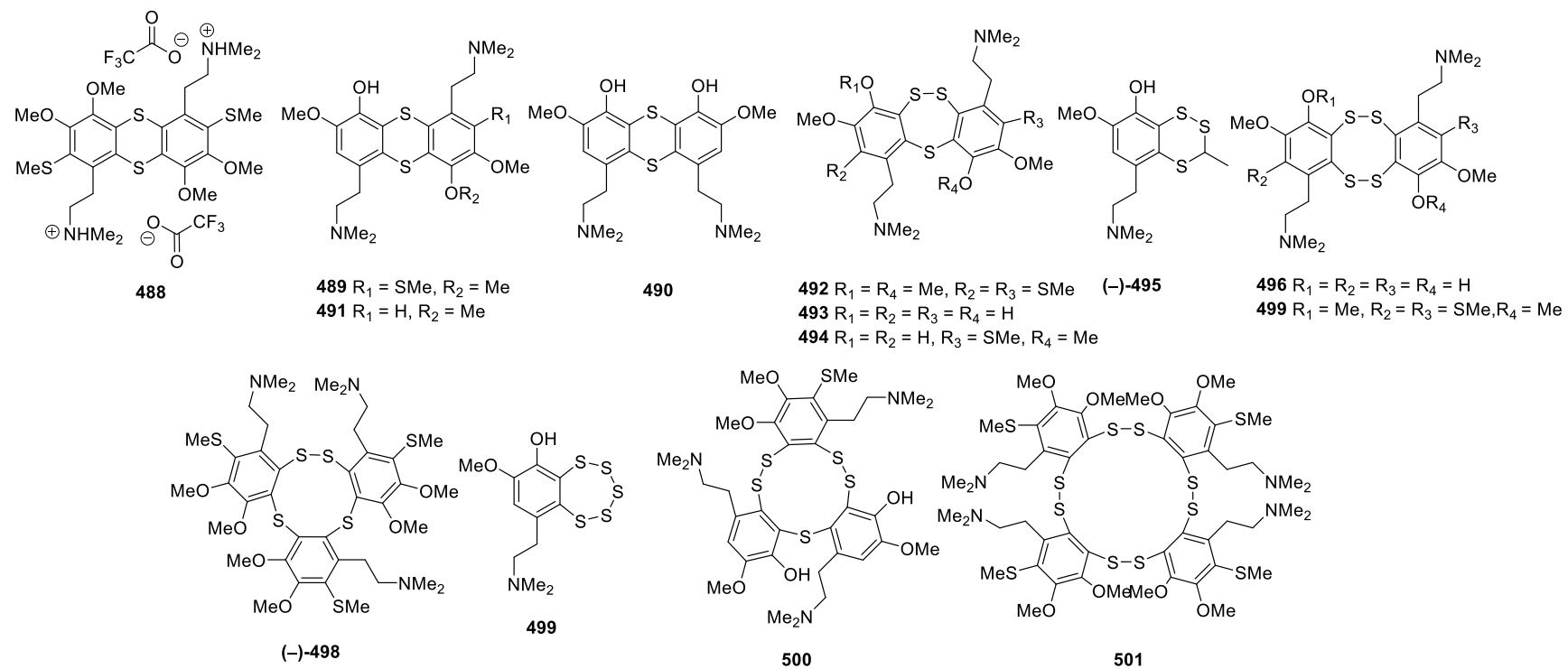


Figure S16: Structures of marine polysulfur aromatic alkaloids from Indonesian waters found in 1970–2017.

Table S16: Marine polysulfur aromatic alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Lissoclibadin 3 488 ^{B,0} [C ₂₆ H ₃₈ N ₂ O ₄ S ₅]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Antifungal	<i>M. hiemalis</i> IAM 6088	NA (20 – 50 µg/disk)	<i>L. cf. badium</i>	NSW	[331, 332, 334]
				<i>S. cerevisiae</i> IAM 1438T	7.8 mm (50 µg/disk)			
					NA (20 µg/disk)			
			Antibacterial	<i>S. aureus</i> IAM 12544T	10 – 13.8 mm (20 – 50 µg/disk)			
				<i>E. coli</i> IAM 12119T	10.8 mm (50 µg/disk)			
				V79	NA (20 µg/disk)			
Lissoclibadin 6 489 ^{B,0} [C ₂₄ H ₃₄ N ₂ O ₄ S ₃]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid		HCT-15, HeLa-S3	IC ₅₀ = 0.34 µM	<i>L. cf. badium</i>	NSW	[335]
					IC ₅₀ = 13.2 ± 1.4 – 16.0 ± 1.8 µM			
			Cytotoxic	T-47D, MDA-MB-231, NCI-H460, ACHN, UO31, DLD-1, HCT116, MALME-3M	IC ₅₀ = 1.22 – 3.59 µM			
				PC3, HL-60	IC ₅₀ = 5.50 – 7.02 µM			
			Antifungal	<i>M. hiemalis</i> IAM 6088	NA (20 – 50 µg/disk)			
				<i>S. cerevisiae</i> IAM 1438T	7.8 mm (50 µg/disk)			
Lissoclibadin 11 490 ^{B,0} [C ₂₄ H ₃₄ N ₂ O ₄ S ₃]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Antibacterial	<i>S. aureus</i> IAM 12544T	10 – 13.8 mm (20 – 50 µg/disk)	<i>L. cf. badium</i>	NSW	[336]
				<i>E. coli</i> IAM 12119T	10.8 (50 µg/disk)			
				V79	NA (20 µg/disk)			
			Cytotoxic		EC ₅₀ = 0.06 µM			
				V79, L1210	IC ₅₀ > 20 µM			
			Cytotoxic		IC ₅₀ = 7.90 µM			
Lissoclibadin 12 491 ^{B,0} [C ₂₂ H ₃₀ N ₂ O ₄ S ₂]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid		V79	IC ₅₀ = 7.90 µM	<i>L. cf. badium</i>	NSW	[336]
			Cytotoxic	L1210	IC ₅₀ > 20 µM			
Lissoclibadin 2 492 ^{B,0} [C ₂₆ H ₃₈ N ₂ O ₄ S ₅]	UV, IR, MS, NMR, Mol. Mod.	Polysulfur Aromatic Alkaloid	Antifungal	<i>M. hiemalis</i> IAM 6088	13.8 mm (50 µg/disk)	<i>L. cf. badium</i>	NSW	[331, 332, 334]

Table S16: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Lissoclibadin 2 492 ^{B,θ} [C ₂₆ H ₃₈ N ₂ O ₄ S ₅]	UV, IR, MS, NMR, Mol. Mod.	Polysulfur Aromatic Alkaloid	Antibacterial	<i>R. atlantica</i> TUF-D	12.2 – 28.2 mm (5 – 50 µg/disk)	<i>L. cf. badium</i>	NSW	[331, 332, 334]
				<i>S. cerevisiae</i> IAM 1438T, <i>E. coli</i> IAM 12119T	NA (20, 50 µg/disk)			
				<i>S. aureus</i> IAM 12544T	NA (50 µg/disk)			
			Cytotoxic	T-47D, MDA-MB-231, NCI-H460, PC3, ACHN, UO31, DLD-1, HCT116, MALME-3M	IC ₅₀ = 0.10 – 0.77 µM			
				HL-60	20 – 90% (1 – 10 µM) IC ₅₀ = 0.21 µM			
				V79	IC ₅₀ = 0.08 µM			
Lissoclibadin 4 493 ^{B,θ} [C ₂₂ H ₃₀ N ₂ O ₄ S ₃]	UV, IR, MS, NMR, Mol. Mod.	Polysulfur Aromatic Alkaloid	Anti-inflammatory	IL-8 (PMA, HL-60)	3 – 10 µM	<i>L. cf. badium</i>	NSW	[333, 335]
			Antifungal	<i>M. hiemalis</i> IAM 6088	NA (20, 50 µg/disk)			
			Antibacterial	<i>S. cerevisiae</i> IAM 1438T	NA (20, 50 µg/disk)			
				<i>S. aureus</i> IAM 12544T	10.8 mm (20 µg/disk)			
			Cytotoxic	<i>E. coli</i> IAM 12119T	15.7 mm (50 µg/disk) NA (20 µg/disk)			
				V79	EC ₅₀ = 0.71 µM			
Lissoclibadin 5 494 ^{B,θ} [C ₂₄ H ₃₄ N ₂ O ₄ S ₄]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Antifungal	HCT-15, HeLa-S3	IC ₅₀ = 17.2 ± 5.2 – 17.8 ± 5.3 µM	<i>L. cf. badium</i>	NSW	[335]
				<i>M. hiemalis</i> IAM 6088	NA (20, 50 µg/disk)			
			Antibacterial	<i>S. cerevisiae</i> IAM 1438T	NA (20, 50 µg/disk)			
				<i>S. aureus</i> IAM 12544T	10.8 – 13.1 mm (20 – 50 µg/disk)			
				<i>E. coli</i> IAM 12119T	15.7 mm (50 µg/disk)			
				<i>E. coli</i> IAM 12119T	NA (20 µg/disk) EC ₅₀ = 0.71 µM			
(–)-Lissoclibadin 13 495 ^{B,θ} [C ₁₃ H ₁₉ NO ₂ S ₃]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Cytotoxic	V79, L1210	IC ₅₀ = 0.44 – 2.20 µM	<i>L. cf. badium</i>	NSW	[336]

Table S16: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Lissoclibadin 7 496 ^{B,0} [C ₂₂ H ₃₀ N ₂ O ₄ S ₄]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Antifungal	<i>M. hiemalis</i> IAM 6088	NA (20 – 50 µg/disk)	<i>L. cf. badium</i>	NSW	[333, 335]
				<i>S. cerevisiae</i> IAM 1438T	16.8 mm (50 µg/disk)			
			Antibacterial	<i>S. aureus</i> IAM 12544T	NA (20 µg/disk)			
				<i>E. coli</i> IAM 12119T	9.4 – 13.1 mm (20 – 50 µg/disk)			
			Cytotoxic	V79 HCT-15, HeLa-S3	9.9 mm (50 µg/disk) NA (20 µg/disk) EC ₅₀ = 0.17 µM IC ₅₀ = 14.2 ± 1.3 – 15.7 ± 1.2 µM			
Lissoclibadin 10 497 ^{B,0} [C ₂₆ H ₃₈ N ₂ O ₄ S ₆]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid	Undetm.	Undetm.	Undetm.	<i>L. cf. badium</i>	NSW	[336]
			Antifungal	<i>M. hiemalis</i> IAM 6088	NA (20 – 50 µg/disk)			
				<i>R. atlantica</i> TUF-D	15.2 – 28.2 mm (20 – 50 µg/disk)			
			Antibacterial	<i>S. cerevisiae</i> IAM 1438T <i>S. aureus</i> IAM 12544T <i>E. coli</i> IAM 12119T	NA (5 µg/disk) NA (20 – 50 µg/disk) NA (50 µg/disk)			
				HCT-15, HeLa-S3, MCF7, NCI-H28	NA (20 – 50 µg/disk) IC ₅₀ = 4.0 ± 1.8 – 7.6 ± 4.0 µM			
(–)-Lissoclibadin 1 498 ^{B,0} [C ₃₉ H ₅₇ N ₃ O ₆ S ₇]	UV, IR, MS, NMR, Mol. Mod	Polysulfur Aromatic Alkaloid*		V79, HL-60, T-47D, MDA- MB-231, NCI-H460, PC3, ACHN, UO31, DLD-1, HCT116, MALME-3M	IC ₅₀ = 0.13 – 0.82 µM	<i>L. cf. badium</i>	NSW	[330 – 333]
			Cytotoxic	HCT-15	Apop. 5 µM (24 h) caspa- se-dependent pathway			
				Nude mice (HCT)	60% (25 mg/kg per day wo. signt. secon. adv. eff)			
				V79, L1210	IC ₅₀ = 0.70 – 1.80 µM			
				HCT-15, HeLa-S3, MCF7, NCI-H28	IC ₅₀ = 4.2 ± 2.4 – 6.4 ± 2.7 µM			
Lissoclibadin 14 499 ^{B,0} [C ₁₁ H ₁₅ NO ₂ S ₅]	UV, IR, MS, NMR	Polysulfur Aromatic Alkaloid*	Cytotoxic			<i>L. cf. badium</i>	NSW	[336]

Table S16: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Lissoclibadin 9 500 ^{B,θ} [C ₃₅ H ₄₉ N ₃ O ₆ S ₆]	UV, IR, MS, NMR, Mol. Mod.	Polysulfur Aromatic Alkaloid	Cytotoxic	V79, L1210	IC ₅₀ = 0.38 – 0.63 μM	<i>L. cf. badium</i>	NSW	[336]
Lissoclibadin 8 501 ^{B,θ} [C ₅₂ H ₇₆ N ₄ O ₈ S ₁₂]	UV, IR, MS, NMR, Mol. Mod.	Polysulfur Aromatic Alkaloid*	Cytotoxic	V79, L1210 HCT-15, HeLa-S3, MCF7, NCI-H28	IC ₅₀ = 0.14 – 2.00 μM IC ₅₀ = 4.9 ± 1.9 – 11.8 ± 3.1 μM	<i>L. cf. badium</i>	NSW	[336]

Footnote: 1. Activity (ACHN human renal carcinoma, DLD-1 human colorectal adenocarcinoma, HeLa-S3 human cervical adenocarcinoma, MALME-3M human melanoma cancer, NCI-H28 human non-small cell lung cancer, IL-8 interleukin 8, PMA phorbol myristate acetate).

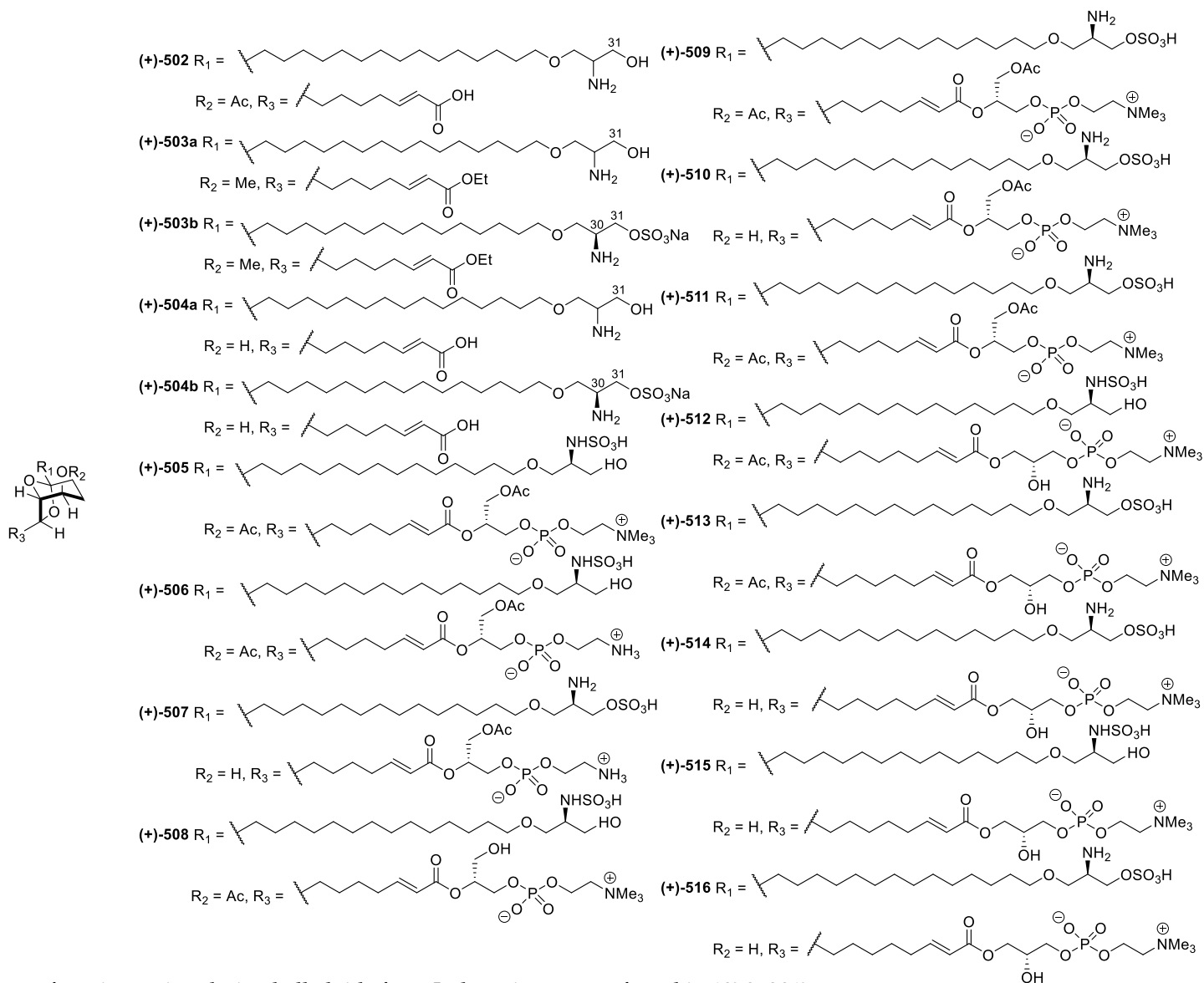


Figure S17: Structures of marine serine-derived alkaloids from Indonesian waters found in 1970–2017.

Table S17: Marine serine-derived alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Didemniserinolipid A 502^β [C ₃₃ H ₅₉ NO ₈]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid*	Cytotoxic	P-388, A549, HT-29	IC ₅₀ > 20 μg/mL	<i>Didemnum</i> sp.	NMU	[337]
(+)-Didemniserinolipid B 503a^β [C ₃₄ H ₆₃ NO ₇]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Cytotoxic	P-388, A549, HT-29	IC ₅₀ > 20 μg/mL	<i>Didemnum</i> sp.	NMU	[337]
(+)-Didemniserinolipid B 503b^γ [C ₃₄ H ₆₂ NNaO ₁₀ S]	TS	Serine Alkaloid	Undetm.	Undetm.	Undetm.			[338, 339]
(+)-Didemniserinolipid C 504a^β [C ₃₁ H ₅₇ NO ₇]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Cytotoxic	P-388, A549, HT-29	IC ₅₀ > 20 μg/mL	<i>Didemnum</i> sp.	NMU	[337]
(+)-Didemniserinolipid C 504b^γ [C ₃₁ H ₅₆ NNaO ₁₀ S]	TS	Serine Alkaloid	Undetm.	Undetm.	Undetm.			[339]
(+)-Siladenoserinol A 505^β [C ₄₃ H ₇₉ N ₂ O ₁₇ PS]	UV, IR, MS, NMR, ECD, [α] _D , CT, TS	Serine Alkaloid*•	Anticancer	p53-Hdm	IC ₅₀ = 2.0 – 17 μM (natural) IC ₅₀ = 17 μM (synthetic)	A tunicate (Didemnidae)	NSW	[340, 341]
(+)-Siladenoserinol B 506^β [C ₄₀ H ₇₃ N ₂ O ₁₇ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 2.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol C 507^β [C ₃₈ H ₇₁ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 4.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol D 508^β [C ₄₁ H ₇₇ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 7.7 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol E 509^β [C ₄₁ H ₇₇ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 18.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol F 510^β [C ₄₁ H ₇₇ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 29.0 μM	A tunicate (Didemnidae)	NSW	[340]

Table S17: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Siladenoserinol G 511 ^β [C ₄₃ H ₇₉ N ₂ O ₁₇ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 53.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol H 512 ^β [C ₄₁ H ₇₇ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 2.5 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol I 513 ^β [C ₄₁ H ₇₇ N ₂ O ₁₆ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 9.3 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol J 514 ^β [C ₃₆ H ₆₉ N ₂ O ₁₅ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 11.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol K 515 ^β [C ₃₉ H ₇₅ N ₂ O ₁₅ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 13.0 μM	A tunicate (Didemnidae)	NSW	[340]
(+)-Siladenoserinol L 516 ^β [C ₃₉ H ₇₅ N ₂ O ₁₅ PS]	UV, IR, MS, NMR, [α] _D	Serine Alkaloid	Anticancer	p53-Hdm	IC ₅₀ = 55.0 μM	A tunicate (Didemnidae)	NSW	[340]

Footnote: 1. Activity (p53 tumor protein).

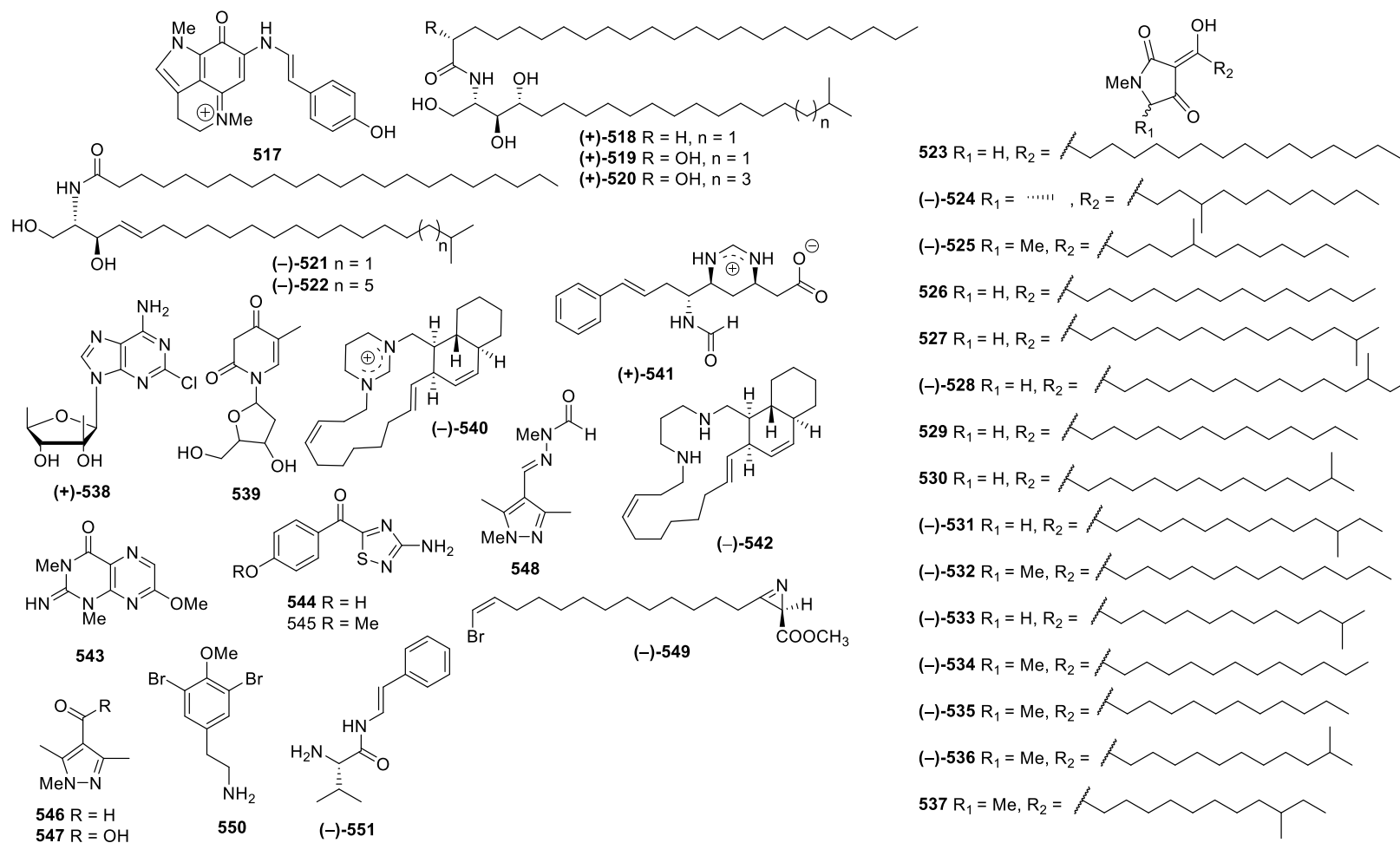


Figure S18: Structures of other marine alkaloids from Indonesian waters found in 1970–2017.

Table S18: Other marine alkaloids from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Makavulamine G 517 ^{β,κ} [C ₂₀ H ₂₀ N ₃ O ₂]	UV, IR, MS, NMR	Pyrroloimino-quinone Alkaloid	Cytotoxic	P-388, A549, HT-29, MCF7, KB	IC ₅₀ = 0.35 – 0.5 µg/mL	<i>Histodermella</i> sp.	NSW	[342 – 344]
			Antifungal	<i>A. oryzae</i> , <i>C. albicans</i> , <i>P. notatum</i> , <i>S. cerevisiae</i> , <i>T. mentagrophytes</i>	NA			
			Antiviral	Herpes simplex 1, Herpes simplex 2, Polio virus	NA			
			Immuno modulatory	MLR	IC ₅₀ = 0.28 µg/mL			
			Anticancer	Murine resting lymphocyte DNA topoisomerase-I	IC ₅₀ = 24.4 µg/mL			
				DNA topoisomerase-II	IC ₅₀ = 3.0 µM			
			Neurodisease	RNA, DNA protein synthesis	NA			
				Affinity to <i>L. stagnalis</i> AchBP (radioligand assay)	IC ₅₀ = 15 µM			
				Affinity to <i>T. californica</i> nAChR (radioligand assay)	IC ₅₀ = 21 µM			
				Affinity to human nAChR (radioligand assay)	<i>K_i</i> = 0.55 ± 0.01 µM			
(+)–Strepsiamide A 518 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	Murine muscle nAChR	<i>K_i</i> = 1.60 ± 0.17 µM	<i>S. lendenfeldi</i>	UEP	[345]
				Human nAChR expressed in <i>X. laevis</i>	<i>K_i</i> = 18 ± 2 µM			
(+)–Strepsiamide B 519 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	Electro-physiological activity	IC ₅₀ = 3.3 ± 0.3 µM	<i>S. lendenfeldi</i>	UEP	[345]
				Stimulator at initial stage of development of agricultural plant	40% (10 µM)			
(+)–Strepsiamide A 518 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	<i>H. vulgare</i> (root)	Active	<i>S. lendenfeldi</i>	UEP	[345]
				<i>F. esculentum</i> Moench (root)	Active			
(+)–Strepsiamide B 519 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	L5178Y	ED ₅₀ = 7.6 µg/mL	<i>S. lendenfeldi</i>	UEP	[345]
				HeLa	4% (10 µg/mL)			
(+)–Strepsiamide A 518 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	PC12	3% (3 µg/mL)	<i>S. lendenfeldi</i>	UEP	[345]
				L5178Y	NA			
(+)–Strepsiamide B 519 ^β [C ₄₃ H ₈₇ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	HeLa, PC12	ED ₅₀ > 10 µg/mL	<i>S. lendenfeldi</i>	UEP	[345]
					NA			

Table S18: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Strepsiamide C 520 ^β [C ₄₄ H ₈₉ NO ₄]	IR, MS, NMR, [α] _D , CT	Ceramide	Cytotoxic	L5178Y HeLa, PC12	68% (10 µg/mL) NA	<i>S. lendenfeldi</i>	UEP	[345]
(-)-Iotrochotamide I 521 ^β [C ₄₃ H ₈₅ NO ₃]	IR, MS, NMR, [α] _D , CT	Ceramide	Undetm.	Undetm.	Undetm.	<i>I. purpurea</i>	SSW	[346]
(-)-Iotrochotamide II 522 ^β [C ₄₇ H ₉₃ NO ₃]	IR, MS, NMR, [α] _D , CT	Ceramide	Undetm.	Undetm.	Undetm.	<i>I. purpurea</i>	SSW	[346]
Melophlin A 523 ^β [C ₂₁ H ₃₇ NO ₃]	UV, IR, MS, NMR, CT	Tetramic acid Alkaloid	Cytotoxic	HL-60 L1210, KB-3-1, A498, U-937, L-929	0.2 – 0.4 µg/mL IC ₅₀ = 8.55 – 26 µM	<i>M. sarassinorum</i>	SSW	[84 – 87, 347]
				Revers. the morp. H- <i>ras</i> transf. NIH3T3 fibroblast to normal	5 µg/mL			
				Modulator signal transduction (Ras) Arrest. NIH3T3 (G1)	Dynamins II and I- like protein 1 µg/mL			
				Mg and Zn complex	Cytotoxic in various cancer cells			
			Antibacterial	<i>M. smegmatis</i> (Glucose), <i>M.</i> <i>bovis</i> BCG (Glucose), <i>M.</i> <i>bovis</i> BCG (Palmitate)	MIC = 25 µg/mL			
				<i>M. smegmatis</i> (Propionate), <i>M. bovis</i> BCG (Propionate), <i>M. smegmatis</i> (Palmitate)	MIC = 0.8 – 3.0 µg/mL			
			Anti-inflammatory	<i>M. smegmatis</i> (Palmitate) V79	MIC = 12.5 µg/mL ED ₅₀ = 27.2 µM			
				IL-8 (PMA, HL-60) HL-60	NA 0.2 – 0.4 µg/mL			
(-)-Melophlin B 524 ^β [C ₁₉ H ₃₃ NO ₃]	UV, IR, MS, NMR, ECD, [α] _D , CT	Tetramic acid Alkaloid	Cytotoxic	Revers. the morp. H- <i>ras</i> transf. NIH3T3 fibroblast to normal Arrest. NIH3T3 at G1 phase	5 µg/mL 1 µg/mL	<i>M. sarassinorum</i>	SSW	[347]

Table S18: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(–)-Melophlin C 525 ^β [C ₁₉ H ₃₃ NO ₃]	UV, IR, MS, NMR, ECD, [α] _D , CT	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
			Antibacterial	<i>A. salina</i> <i>S. aureus</i> <i>S. aureus</i> , <i>B. subtilis</i>	IC ₅₀ = 36.6 µg/mL 18 mm (5 µg/disk) 11 – 16 mm (5 – 10 µg/disk)			
			Antifungal	<i>C. albicans</i>	15 mm (10 µg/disk)			
			Antiinsecticidal	<i>S. littoralis</i>	M			
Melophlin D 526 ^β [C ₂₀ H ₃₅ NO ₃]	UV, IR, MS, NMR	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
Melophlin E 527 ^β [C ₂₁ H ₃₇ NO ₃]	UV, IR, MS, NMR	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
(–)-Melophlin F 528 ^β [C ₂₁ H ₃₇ NO ₃]	UV, IR, MS, NMR, [α] _D	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
Melophlin G 529 ^β [C ₁₉ H ₃₃ NO ₃]	UV, IR, MS, NMR	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
			Antibacterial	<i>A. salina</i> <i>S. aureus</i> <i>B. subtilis</i>	IC ₅₀ = 48.8 µg/mL 9 – 10 mm (5 – 10 µg/disk) 15 mm (10 µg/disk)			
			Antiinsecticidal	<i>S. littoralis</i>	Moderate			
Melophlin H 530 ^β [C ₂₀ H ₃₅ NO ₃]	UV, IR, MS, NMR	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
(–)-Melophlin I 531 ^β [C ₂₀ H ₃₅ NO ₃]	UV, IR, MS, NMR, [α] _D	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
			Antibacterial	<i>A. salina</i> <i>S. aureus</i> , <i>B. subtilis</i> <i>S. littoralis</i>	IC ₅₀ = 52.6 µg/mL 9 mm (5 – 10 µg/disk) M			
(–)-Melophlin J 532 ^β [C ₂₀ H ₃₅ NO ₃]	UV, IR, MS, NMR, [α] _D , CT	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
Melophlin K 533 ^β [C ₁₉ H ₃₄ NO ₃]	UV, IR, MS, NMR	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
(–)-Melophlin L 534 ^β [C ₁₉ H ₃₄ NO ₃]	UV, IR, MS, NMR, [α] _D , CT	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
(–)-Melophlin M 535 ^β [C ₁₈ H ₃₁ NO ₃]	UV, IR, MS, NMR, [α] _D , CT	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]
Melophlin N 536 ^β [C ₁₉ H ₃₃ NO ₃]	UV, IR, MS, NMR, CT	Tetramic acid Alkaloid	Cytotoxic	HL-60, HeLa, TF-1	NA	<i>M. sarassinorum</i>	SSW	[348]

Table S18: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Melophlin O 537 ^β [C ₁₉ H ₃₃ NO ₃]	UV, IR, MS, NMR, CT	Tetramic acid Alkaloid	Undetm.	Undetm.	Undetm.	<i>M. sarassinorum</i>	SSW	[348]
(+)-Kumusine 538 ^β [C ₁₁ H ₁₄ ClN ₅ O ₃]	UV, IR, MS, NMR, ECD, [α] _D	Nucleoside Alkaloid*	Imuuno suppressive activity	MLR LcV	IC ₅₀ = 0.195 µg/mL IC ₅₀ = 5.0 µg/mL, potency > 256	<i>Theonella</i> sp.	NSW	[349, 350]
1-(Tetrahydro-4-hydroxy-5-(hydroxymethyl)furan-2-yl)-5-methyl pyrimidine-2,4(1H,3H)-dione 539 ^β [C ₁₁ H ₁₅ NO ₅]	IR, MS, NMR	Nucleoside Alkaloid	Cytotoxic	P-388, HT-29 A549, <i>C. aethiops</i> kidney cells Myeloma cells	IC ₅₀ = 5.0 µg/mL IC ₅₀ = 2.5 µg/mL IC ₅₀ = 0.18 µg/mL	<i>Kaliopsis</i> sp.	BLI	[351]
(-)-Neopetrocyclamine A 540 ^β [C ₂₆ H ₄₁ N ₂]	IR, MS, NMR, [α] _D , Mol. Mod.	Formamido Alkaloid*	Cytotoxic	UO-31, A498, SF295	GI ₅₀ > 20 µg/mL	<i>N. exigua</i>	EKM	[352]
(+)-Lanesoic acid 541 ^β [C ₁₇ H ₂₁ N ₃ O ₃]	UV, IR, MS, NMR, [α] _D , QCC	Formamido Alkaloid*	Cytotoxic	A549 (ATCC CCL-185), HT-29 (ATCC HTB-38), MDA-MB-231 (ATCC HTB-26) PSN-1 (ATCC CRM-CRL-3211)	NA IC ₅₀ = 8.9 µg/mL	<i>Theonella</i> sp.	CSW	[353]
(-)-Neopetrocyclamine B 542 ^β [C ₂₅ H ₄₂ N ₂]	IR, MS, NMR, [α] _D	Polycyclic diamine Alkaloid	Cytotoxic	UO-31, A498, SF295	GI ₅₀ > 20 µg/mL	<i>N. exigua</i>	EKM	[352]
1,3, O ⁷ -Trimethyl isoxanthopterin 543 ^{β,θ} [C ₉ H ₁₁ N ₅ O ₂]	UV, IR, MS, NMR	Pterin Alkaloid	Undetm.	Undetm.	Undetm.	<i>Eudistoma</i> sp.	SSW	[354]
Polycarpathiamine A 544 ^β [C ₉ H ₇ N ₃ O ₂ S]	UV, IR, MS, NMR, CT	Thiadiazole Alkaloid*	Cytotoxic	L5178Y	IC ₅₀ = 0.41 µM	<i>P. aurata</i>	MLU	[46]
Polycarpathiamine B 545 ^β [C ₁₀ H ₉ N ₃ O ₂ S]	UV, IR, MS, NMR	Thiadiazole Alkaloid	Cytotoxic	L5178Y	NA	<i>P. aurata</i>	MLU	[46]
Cinachyrazole A 546 ^ε [C ₇ H ₁₀ N ₂ O]	UV, MS, NMR	Pyrazole Alkaloid	Cytotoxic	L5178Y	NA (IC ₅₀ > 10 µM)	<i>Cinachyrella</i> sp.	MLU	[355]
Cinachyrazole B 547 ^ε [C ₇ H ₁₀ N ₂ O ₂]	UV, MS, NMR	Pyrazole Alkaloid	Cytotoxic	L5178Y	NA (IC ₅₀ > 10 µM)	<i>Cinachyrella</i> sp.	MLU	[355]

Table 18. Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Cinachyrazole C 548 ^β [C ₉ H ₁₄ N ₄ O]	UV, MS, NMR	Pyrazole Alkaloid*	Cytotoxic	L5178Y	NA (IC ₅₀ > 10 μM)	<i>Cinachyrella</i> sp.	MLU	[355]
(-)-Debromoantazirine 549 ^β [C ₁₇ H ₂₈ BrNO ₂]	IR, MS, NMR, [α] _D	Azirine Alkaloid*	Cytotoxic	NBT-T2	IC ₅₀ = 4.7 μg/mL	<i>Dysidea</i> sp.	PUA	[355]
3',5'-dibromo-4'-methoxyphenethylamine 550 ^{β,θ} [C ₉ H ₁₁ Br ₂ NO]	UV, MS, NMR, CT	Simple Amine/Amide Alkaloid	Undetm.	Undetm.	Undetm.	<i>Eudistoma</i> sp.	SSW	[287]
(-)-(E)-2-Amino-3-methyl-N-styrylbutanamide 551 ^β [C ₁₃ H ₁₈ N ₂ O]	UV, MS, NMR, [α] _D	Simple Amine/Amide Alkaloid*	Antibacterial	<i>E. faecium</i> BM4147-1, <i>S. aureus</i> ATCC29213 <i>K. pneumoniae</i> ATCC 12657, <i>E. aerogenes</i> ATCC 13048	MIC > 50 MIC > 100	<i>C. basilana</i>	MLU	[357]

Footnote: 1. **Activity** (NIH3T3 murine fibroblast, TF-1 human erythroleukemia, U-937 human leukemia, **Raji cells** human Burkitt's lymphoma, **PSN-1** human pancreas carcinoma, **A498** human kidney carcinoma, **Revers.** reversing, **Arres.** arresting).

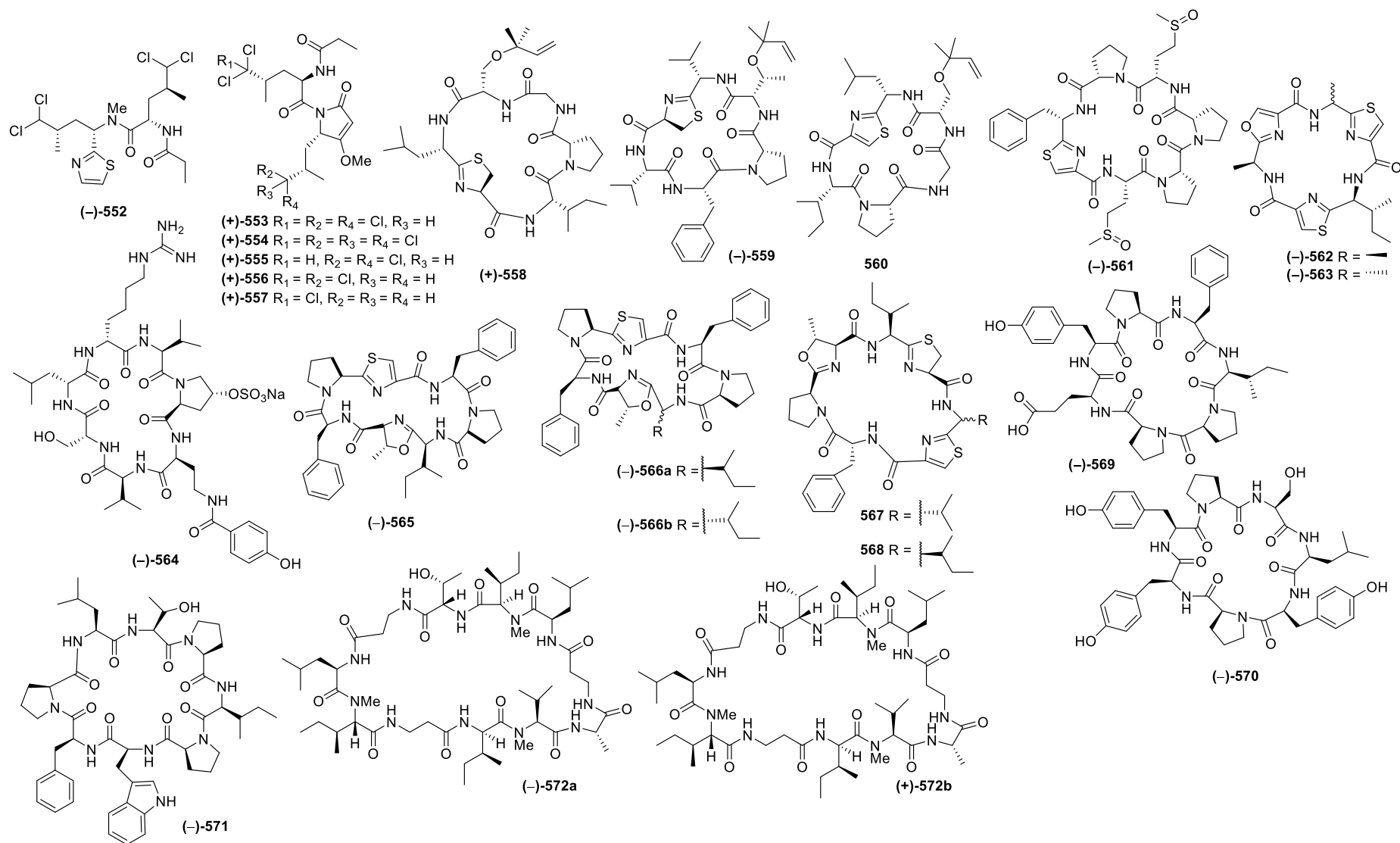


Figure S19: *Cont.*

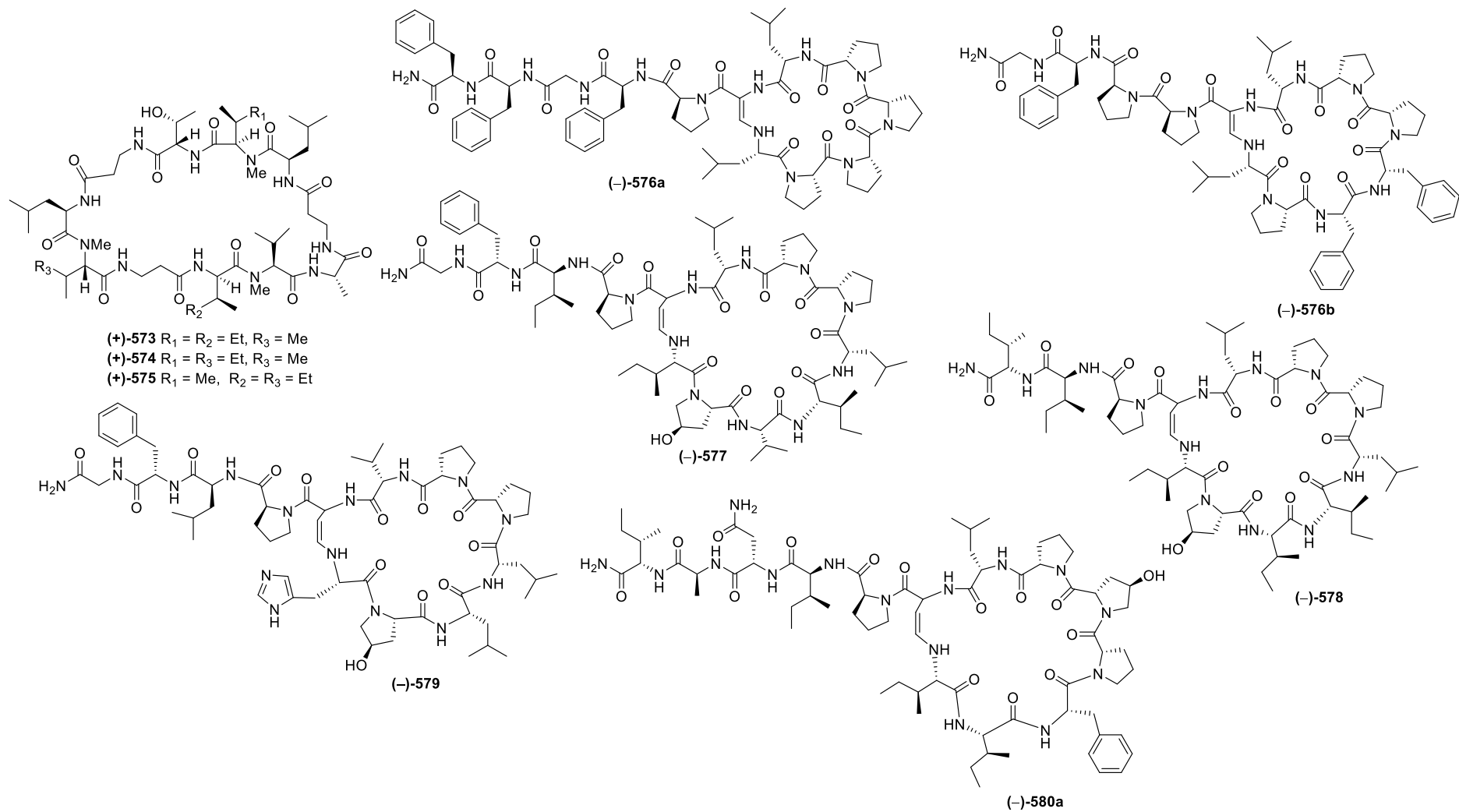


Figure S19: *Cont.*

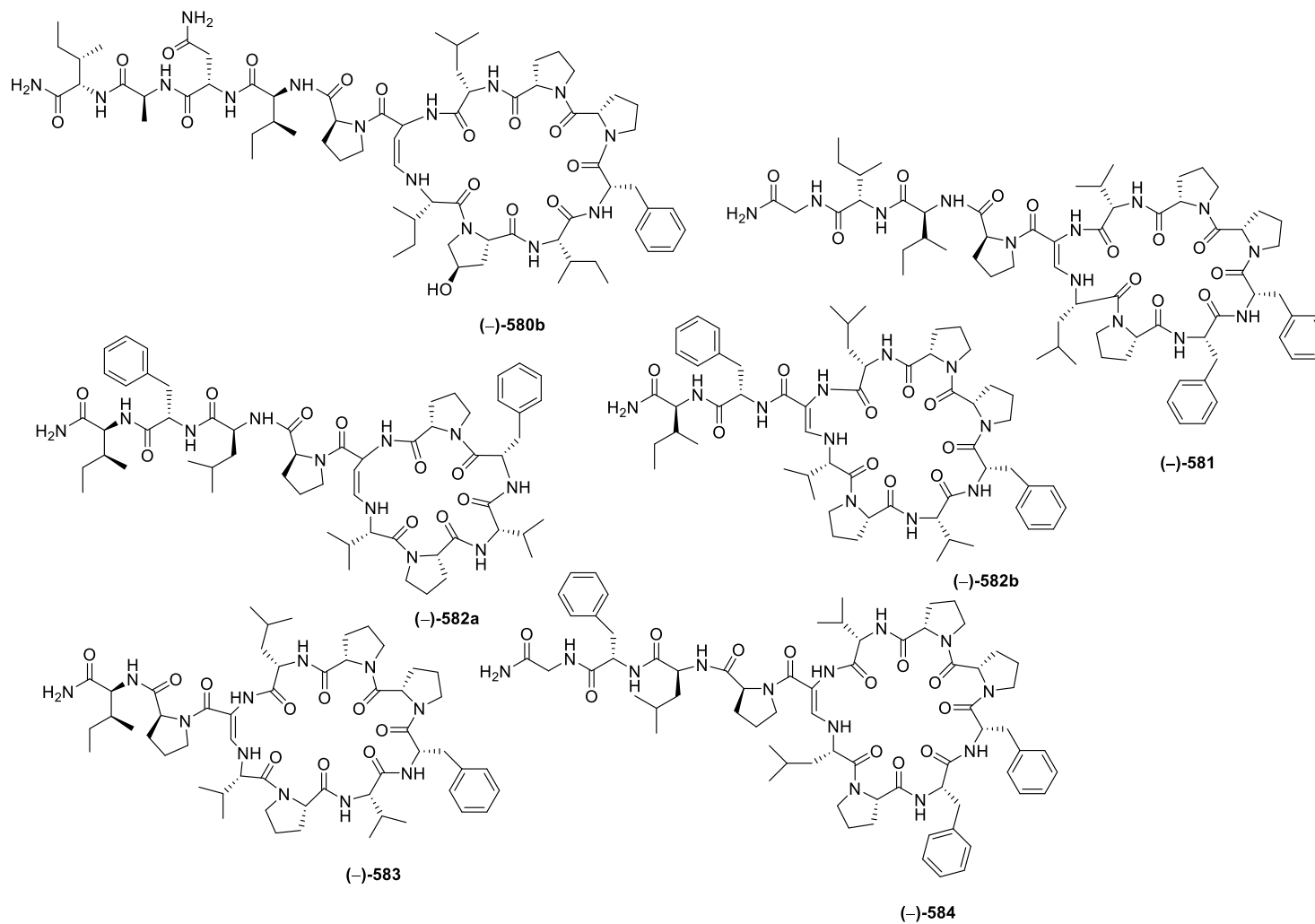


Figure S19: *Cont.*

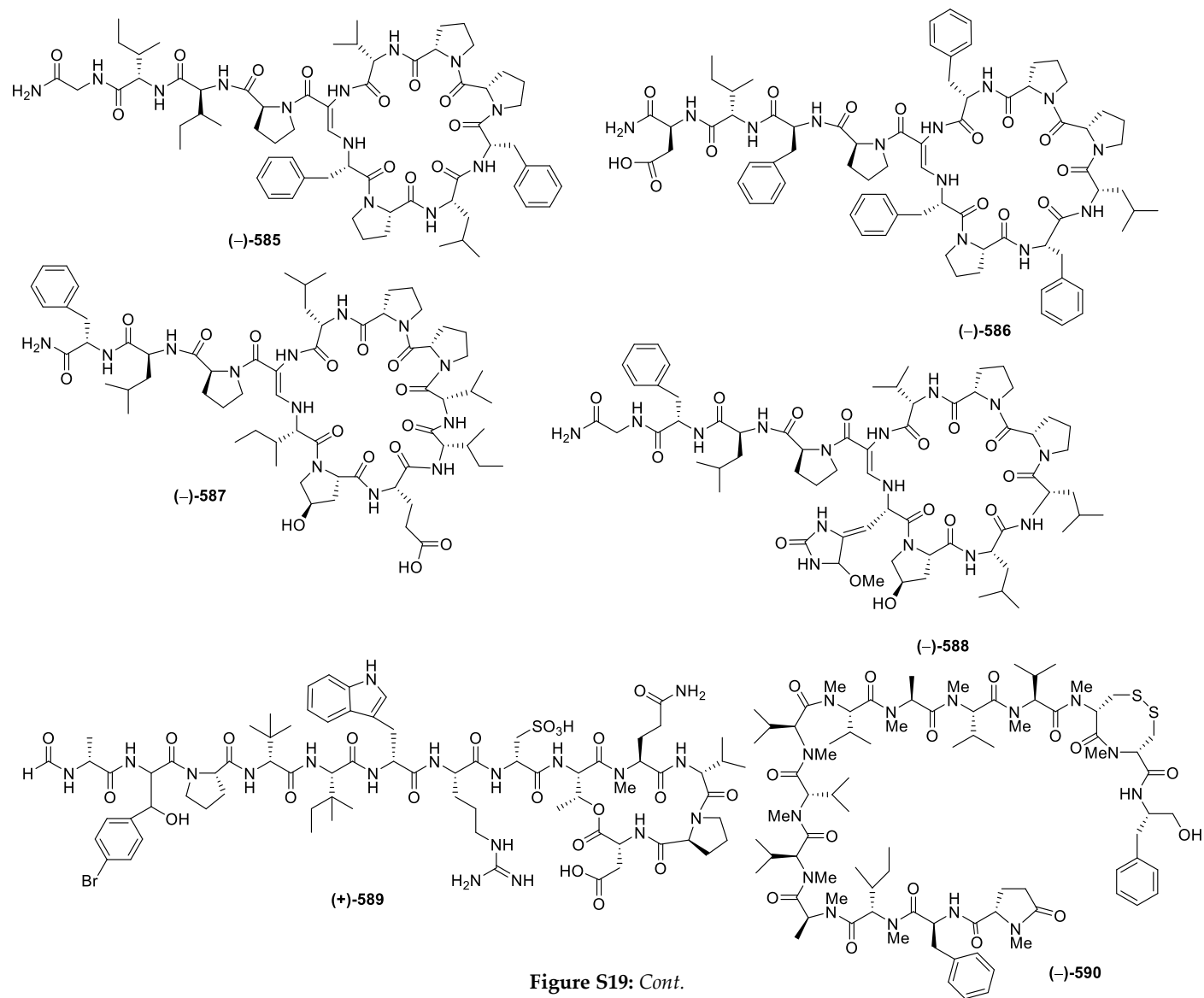


Figure S19: *Cont.*

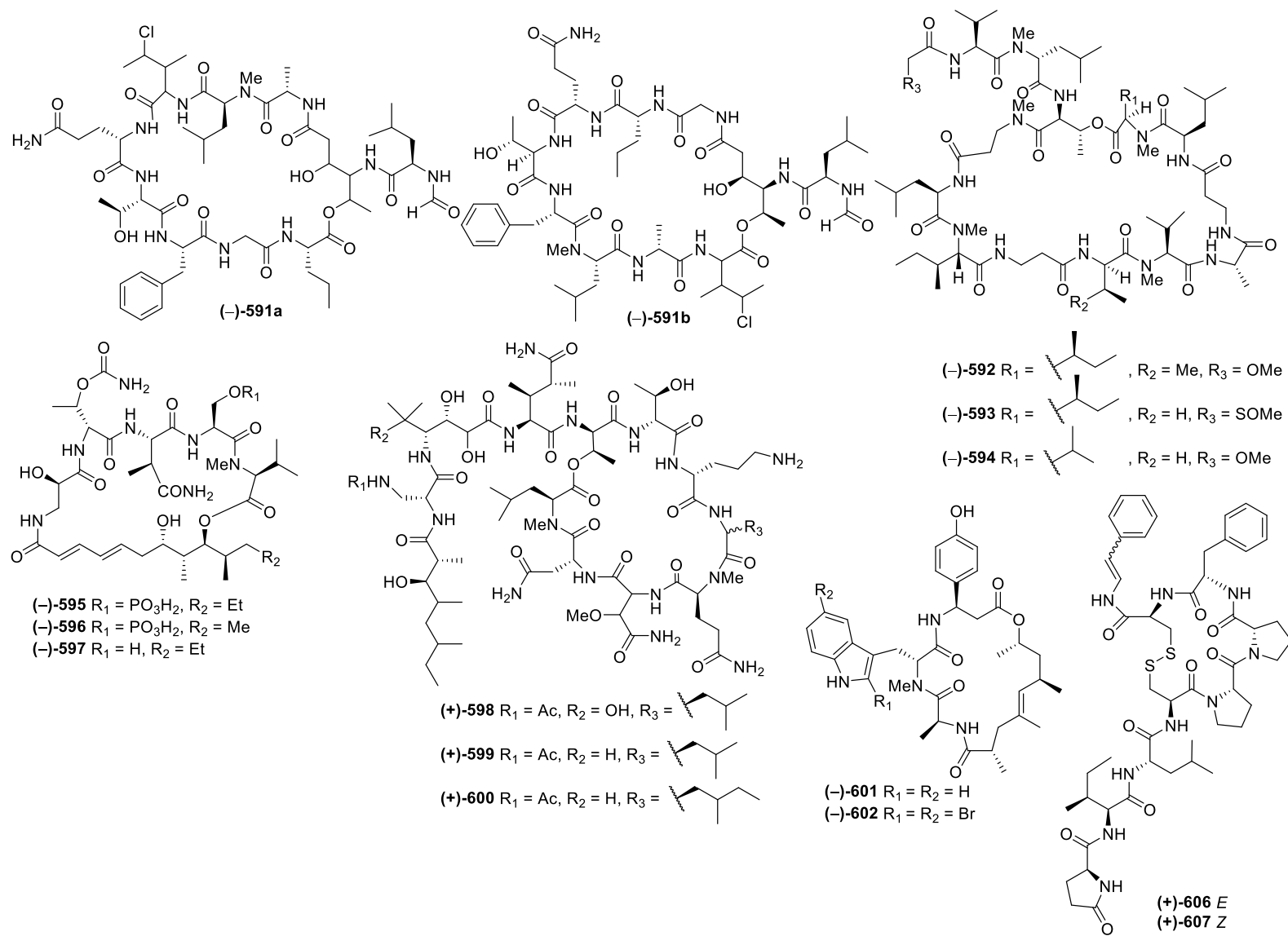


Figure S19: *Cont.*

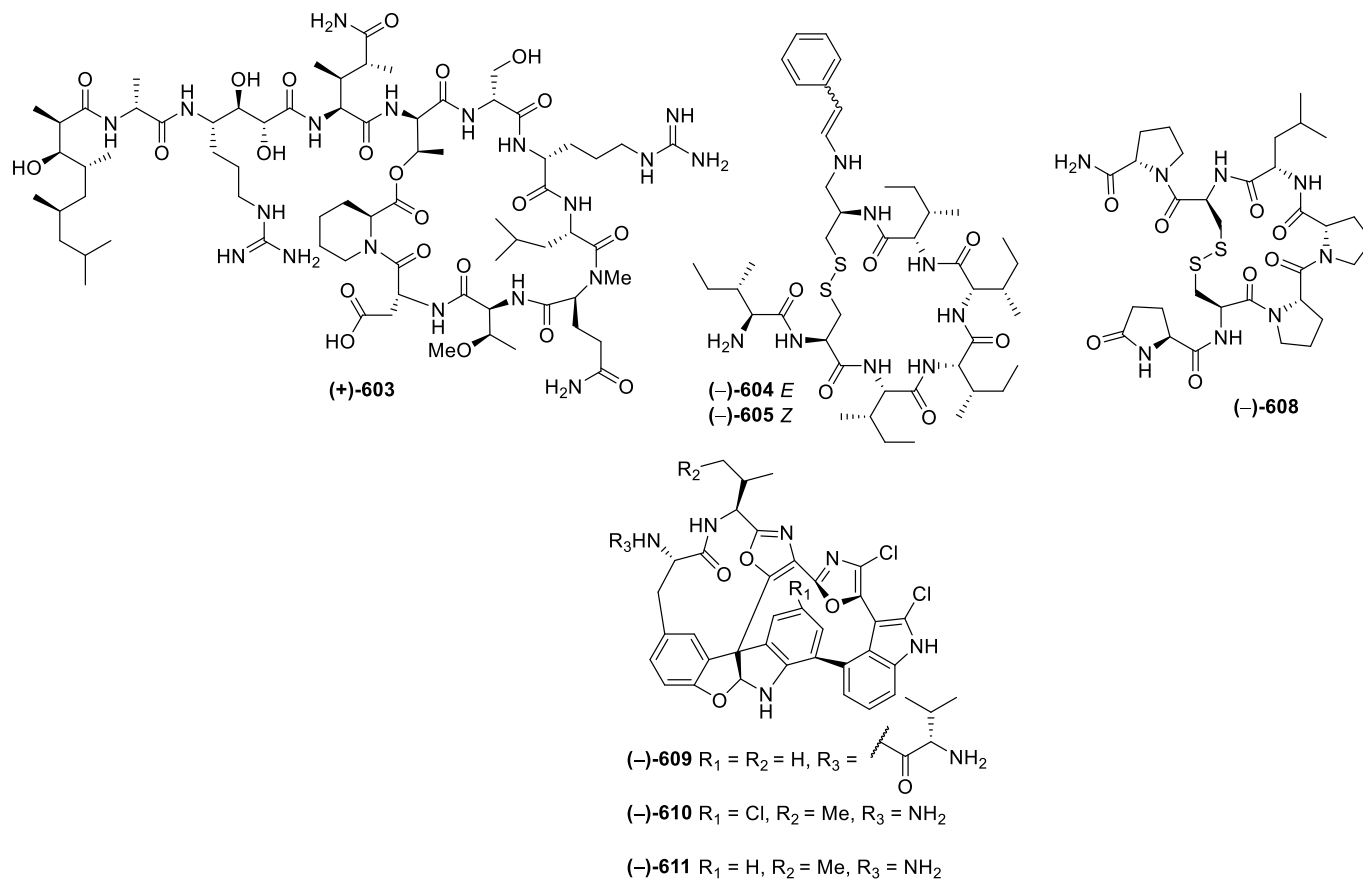


Figure S19: Structures of marine peptides from Indonesian waters found in 1970–2017.

Table S19: Marine peptides from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Dysithiazolamide 552 ^β [C ₁₈ H ₂₇ Cl ₄ N ₃ O ₂ S]	UV, IR, MS, NMR, QCC, TS	Linear Dipeptide	Undetm.	Undetm.	Undetm.	<i>Dysidea</i> sp.	NMU	[358, 359]
(+)-Sintokamide A 553 ^β [C ₁₈ H ₂₅ Cl ₅ N ₂ O ₄]	UV, MS, NMR, [α] _D , X-ray, TS	Linear Dipeptide [▲]	Anticancer	AR NTD in LNCaP AR-positive Cytotoxic	5 µg/mL Active NA (10 µg/mL)	<i>Dysidea</i> sp.	CJV	[360 – 362]
(+)-Sintokamide B 554 ^β [C ₁₈ H ₂₄ Cl ₆ N ₂ O ₄]	UV, MS, NMR, [α] _D , TS	Linear Dipeptide	Undetm.	Undetm.	Undetm.	<i>Dysidea</i> sp.	CJV	[360, 361]
(+)-Sintokamide C 555 ^β [C ₁₈ H ₂₆ Cl ₄ N ₂ O ₄]	UV, MS, NMR, [α] _D , TS	Linear Dipeptide	Anticancer	Androgen receptor-expressing LNCaP	IC ₅₀ = 35 µM	<i>Dysidea</i> sp.	CJV	[360, 363]
(+)-Sintokamide D 556 ^β [C ₁₈ H ₂₆ Cl ₄ N ₂ O ₄]	UV, MS, NMR, [α] _D	Linear Dipeptide	Anticancer	Undetm.	Undetm.	<i>Dysidea</i> sp.	CJV	[360]
(+)-Sintokamide E 557 ^β [C ₁₈ H ₂₇ Cl ₃ N ₂ O ₄]	UV, MS, NMR, [α] _D , TS	Linear Dipeptide	Undetm.	Undetm.	Undetm.	<i>Dysidea</i> sp.	CJV	[360, 361]
(+)–Keenamamide 558 ^β [C ₃₀ H ₄₈ N ₆ O ₆ S]	UV, MS, NMR, [α] _D	Cyclo hexapeptide	Cytotoxic	P-388, A549, MEL-28 HT-29	IC ₅₀ = 2.5 µg/mL IC ₅₀ = 5.0 µg/mL	<i>P. forskalii</i>	NSW	[364, 365]
			Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone)	NA			
			Antibacterial	MRSA, <i>M. intracellulaire</i>	NA			
			Antifungal	<i>C. albicans</i> , <i>C. glabrata</i> , <i>C. krusei</i> , <i>C. neoformans</i>	NA			
			Anti-inflammatory	COX-2 (rat neonatal microglia)	NA			
			Antibacterial	MRSA, <i>M. intracellulaire</i>	NA			
			Antifungal	<i>C. albicans</i> , <i>C. glabrata</i> , <i>C. krusei</i> , <i>C. neoformans</i>	NA			
			Anti-inflammatory	COX-2 (rat neonatal microglia)	NA			
			Antiparasite	<i>P. falciparum</i> (D6 clone), (W2 clone)	IC ₅₀ = 2.0 – 2.1 µg/mL			
(-)-Mollamide B 559 ^β [C ₃₆ H ₅₂ N ₆ O ₆ S]	UV, MS, NMR, [α] _D , Mol. Mod.	Cyclo hexapeptide	Antiparasite	<i>L. donovani</i>	IC ₅₀ = 18 – 35 µg/mL	<i>D. molle</i>	NSW	[364, 365]
			Antiviral	HIV-1 (human PBM cells)	EC ₅₀ = 48.7 µM			
			Cytotoxic	H460 MCF7, SF268	29% (100 µM) 42-44% (100 µM)			

Table S19: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Mollamide C 560 ^β [C ₃₃ H ₄₆ N ₆ O ₆ S]	UV, MS, NMR, [α] _D , CT	Cyclo hexapeptide	Anti-inflammatory	COX-2 (rat neonatal microglia)	NA	<i>D. molle</i>	NSW	[365]
			Cytotoxic	L1210	100 z.u. (CFU-GM)			
				CCRF-CEM, C38	NA			
				HCT116	250 z.u. (CFU-GM)			
(-)-Waiakeamide 561 ^β [C ₃₇ H ₄₉ N ₇ O ₈ S ₃]	UV, MS, NMR, [α] _D , CT	Cyclo hexapeptide [▲]	Cytotoxic	H125, MCF7, LNCap	NA	<i>I. dendroides</i>	NSW	[366]
(-)-Bistramide M 562 ^β [C ₂₁ H ₂₄ N ₆ O ₄ S ₂]	UV, MS, NMR, [α] _D , CT	Cyclo hexapeptide	Cytotoxic	MDA-MB-231, HT-29, A549	GI ₅₀ = 9.1 – 18 μM TGI > 20.0 μM LC ₅₀ > 20.0 μM	<i>L. bistratum</i>	SRWP	[367]
(-)-Bistramide N 563 ^β [C ₂₁ H ₂₄ N ₆ O ₄ S ₂]	UV, MS, NMR, [α] _D , CT	Cyclo hexapeptide	Cytotoxic	MDA-MB-231	GI ₅₀ > 20.0 μM TGI > 20.0 μM LC ₅₀ > 20.0 μM	<i>L. bistratum</i>	SRWP	[367]
(-)-Cupolamide A 564 ^β [C ₄₂ H ₆₆ N ₁₁ NaO ₁₄ S]	UV, MS, NMR, [α] _D , CT	Cyclo heptapeptide [▲]	Cytotoxic	HT-29, A549, PSN-1	GI ₅₀ = 11.0 – 15.0 μM TGI > 20.0 μM LC ₅₀ > 20.0 μM	<i>T. cupola</i>	NSW	[368]
(-)- <i>cis, cis</i> -Ceratospongamide 565 ^β [C ₄₁ H ₄₉ N ₇ O ₆ S]	UV, MS, NMR, [α] _D , Mol. Mod., CT, X-ray, TS	Cyclo heptapeptide	Anti-inflammatory	P-388	IC ₅₀ = 7.5 μg/mL			
			Cytotoxic	Phospholipase A ₂ (HEPG2/ IL-1β)	NA	<i>C. spongiosum</i>	NSW	[369 – 373]
(-)- <i>trans, trans</i> -Ceratospongamide 566a ^β [C ₄₁ H ₄₉ N ₇ O ₆ S]	UV, MS, NMR, [α] _D , Mol. Mod., CT, TS	Cyclo heptapeptide	Anti-inflammatory	<i>A. salina</i>	LD ₅₀ = 13–19 μM (<i>ca</i>)			
(-)- <i>trans, trans</i> -[D- <i>allo</i> -ile] Ceratospongamide 566b ^γ [C ₄₁ H ₄₉ N ₇ O ₆ S]	UV, MS, NMR (¹ H, GCOSY, TOCSY, (ROESY), [α] _D , Mol. Mod., X-ray, TS	Cyclo heptapeptide	Anti-inflammatory	Phospholipase A ₂ (HEPG2/ IL-1β)	ED ₅₀ = 32 nM	<i>C. spongiosum</i>	NSW	[369 – 373]
			Cytotoxic	Phospholipase A ₂ (HEPG2/ IL-1β)	50% red. (reporter) 90% red. (plasmid)			
				<i>A. salina</i>	LD ₅₀ = 13 – 19 μM (<i>ca</i>)			
Lissoclinamide 9 567 ^β [C ₃₅ H ₄₅ N ₇ O ₅ S ₂]	UV, IR, MS, NMR, ECD, Mol. Mod.	Cyclo heptapeptide	Metal binding selectivity	Undetm.	Undetm.	<i>L. patella</i>	NMU	[374, 375]

Table S19: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Lissoclinamide 10 568 ^β [C ₃₆ H ₄₉ N ₇ O ₅ S ₂]	UV, IR, MS, NMR, ECD, Mol. Mod.	Cyclo heptapeptide	Metal binding selectivity	Selective (Cu ²⁺ , excess Zn ²⁺)	High selectivity	<i>L. patella</i>	NMU	[374, 375]
(-)-Carteritin A 569 ^β [C ₄₄ H ₅₇ N ₇ O ₁₀]	UV, IR, MS, NMR, [α] _D , CT	Cyclo heptapeptide	Cytotoxic	HeLa, HCT116, RAW264	IC ₅₀ = 0.70 – 1.50 μM	<i>S. carteri</i>	NSW	[376]
(-)-Carteritin B 570 ^β [C ₄₆ H ₅₇ N ₇ O ₁₁]	UV, IR, MS, NMR, [α] _D , CT	Cyclo heptapeptide	Cytotoxic	HeLa, HCT116, RAW264	IC ₅₀ > 50 μM	<i>S. carteri</i>	NSW	[376]
(-)-Styllissamide X 571 ^β [C ₅₁ H ₆₉ N ₉ O ₉]	UV, IR, MS, NMR, [α] _D , CT	Cyclo octapeptide	Anticancer	HeLa (migration)	0.1-10 μM (wound-healing)	<i>Styllissa</i> sp.	PUA	[377]
				HeLa (via.)	75% (10 μM) (chemotaxicell chamber)			
				HeLa-EGF induced	0.1 – 10 μM			
(-)-Barangamide A 572a ^β [C ₅₄ H ₉₇ N ₁₁ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclo undecapeptide	Cytotoxic	L1210	NA (10 μg/mL)	<i>T. swinhoei</i>	SSW	[378]
(+)-Barangamide A 572b ^γ [C ₅₄ H ₉₇ N ₁₁ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclo undecapeptide	Cytotoxic	MLR	NA (100 μg/mL)	<i>T. swinhoei</i>	SSW	[379]
(+)-Barangamide B 573 ^β [C ₅₃ H ₉₅ N ₁₁ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclo undecapeptide	Undetm.	Undetm.	Undetm.	<i>T. swinhoei</i>	SSW	[379]
(+)-Barangamide C 574 ^β [C ₅₃ H ₉₅ N ₁₁ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclo undecapeptide	Undetm.	Undetm.	Undetm.	<i>T. swinhoei</i>	SSW	[379]
(+)-Barangamide D 575 ^β [C ₅₃ H ₉₅ N ₁₁ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclo undecapeptide	Undetm.	Undetm.	Undetm.	<i>T. swinhoei</i>	SSW	[379]
(-)-Callyaerin G 576a ^β [C ₆₉ H ₉₁ N ₁₃ O ₁₂]	UV, IR, MS, NMR, [α] _D , CT	Cyclohexapeptide (penta peptide sc.)	Cytotoxic	L5178Y	ED ₅₀ = 0.41 – 0.53 μg/mL	<i>C. aerizusa</i>	MLU	[380, 381]
(-)-Callyaerin G 576b ^γ [C ₆₉ H ₉₁ N ₁₃ O ₁₂]	NMR (COSY, TOCSY, ROESY)	Cyclohexapeptide (penta peptide sc.)	Cytotoxic	HeLa, PC12	ED ₅₀ = 3.8 – 4.43 μM			
			Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 μM			
(-)-Callyaerin A 577 ^β [C ₇₀ H ₁₁₀ N ₁₄ O ₁₄]	UV, MS, NMR, [α] _D , CT, TS	Cyclooctapeptide (tetrapeptide sc.)	Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 μM	<i>C. aerizusa</i>	MLU	[381 – 383]
				L5178Y	ED ₅₀ = 3.61 μM			
				<i>C. albicans</i>	25 – 30 mm (5 – 10 μL)			
				<i>S. aureus</i> , <i>B. subtilis</i>	0 – 9 mm (5 – 10 μL)			
				<i>E. coli</i>	10 – 15 mm (5 – 10 μL)			
				<i>M. tuberculosis</i>	MIC ₉₀ = 2 μM			
					MIC ₁₀₀ = 6 μM			

Table S19: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Callyaerin A 577 ^β [C ₇₀ H ₁₁₀ N ₁₄ O ₁₄]	UV, MS, NMR, [α] _D , CT, TS	Cyclooctapeptide (tetrapeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ = 20 – 50 μM IC ₉₀ = 40 – 50 μM	<i>C. aerizusa</i>	MLU	[381 – 383]
			Cytotoxic	<i>A. salina</i> L5178Y HeLa, PC12	15% (20 μg/mL) 35% (50 μg/mL) ED ₅₀ = 4.14 μM ED ₅₀ > 8 μM			
(-)-Callyaerin B 578 ^β [C ₆₆ H ₁₁₁ N ₁₃ O ₁₃]	UV, MS, NMR, [α] _D , CT	Cyclooctapeptide (tripeptide sc.)	Antifungal	THP-1, MRC-5	IC ₅₀ = 2 – 5 μM IC ₉₀ = 6 – 30 μM	<i>C. aerizusa</i>	MLU	[381, 382]
			Antibacterial	<i>C. albicans</i> <i>S. aureus</i> <i>B. subtilis</i> <i>E. coli</i>	15 mm (5 – 10 μL) 7 – 10 mm (5 – 10 μL) 0 mm (5 – 10 μL) 11 mm (5 – 10 μL)			
				<i>M. tuberculosis</i>	MIC ₉₀ = 5 μM MIC ₁₀₀ = 10 μM			
			Cytotoxic	L5178Y THP-1, MRC-5	ED ₅₀ = 2.92 μM IC ₅₀ > 100 μM, IC ₉₀ > 100 μM			
(-)-Callyaerin C 579 ^β [C ₆₄ H ₉₅ N ₁₅ O ₁₃]	UV, MS, NMR, [α] _D , CT	Cycloheptapeptide (tetrapeptide sc.)	Antifungal	<i>C. albicans</i> <i>S. aureus</i>	0 mm (5 – 10 μL) 7 – 10 mm (5 μL)	<i>C. aerizusa</i>	MLU	[381, 382]
			Antibacterial	<i>B. subtilis</i> , <i>E. coli</i> <i>M. tuberculosis</i>	0 mm (5 – 10 μL) MIC ₉₀ = 40 μM MIC ₁₀₀ = 100 μM			
(-)-Callyaerin D 580a ^β [C ₇₀ H ₁₀₉ N ₁₅ O ₁₅]	UV, MS, NMR, [α] _D , CT	Cycloheptapeptide (pentapeptide sc.)	Cytotoxic	L5178Y THP-1, MRC-5	ED ₅₀ = 3.03 μM IC ₅₀ > 10 μM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	<i>S. aureus</i> , <i>E. coli</i> <i>B. subtilis</i> <i>M. tuberculosis</i>	0 mm (5 – 10 μL) 12 mm (5 – 10 μL) MIC ₉₀ > 100 μM			
			Antifungal	<i>C. albicans</i>	0 – 7 mm (5 – 10 μL)			
(-)-Callyaerin D 580b ^γ [C ₇₀ H ₁₀₉ N ₁₅ O ₁₅]	HMBC, ROESY, ESIMS	Cycloheptapeptide (pentapeptide sc.)	Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 μM	<i>C. aerizusa</i>	MLU	[381]

Table S19: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Callyaerin E 581 ^β [C ₆₆ H ₉₅ N ₁₃ O ₁₂]	UV, MS, NMR, [α] _D , CT	Cyclohepta-peptide (tetra peptide sc.)	Cytotoxic	<i>A. salina</i> L5178Y	45 – 70% (20 – 50 µg/mL) ED ₅₀ = 0.39 µM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	HeLa, PC12	ED ₅₀ = 3.4 – 3.8 µM			
				<i>S. aureus</i>	9 – 10 mm (5 – 10 µL)			
			Antifungal	<i>B. subtilis</i> <i>E. coli</i> <i>M. tuberculosis</i> <i>C. albicans</i>	15 – 17 mm (5 – 10 µL) 9 – 11 mm (5 – 10 µL) MIC ₉₀ = 100 µM 20 mm (5 – 10 µL)			
(-)-Callyaerin F 582a ^β [C ₅₉ H ₈₅ N ₁₁ O ₁₀]	UV, MS, NMR, [α] _D , chemical transformation	Cyclo pentapeptide (tetrapeptide sc.)	Cytotoxic	L5178Y, HeLa, PC12	ED ₅₀ > 9 µM	<i>C. aerizusa</i>	MLU	[381]
			Antibacterial	THP-1, MRC-5	IC ₅₀ > 10 µM			
			Antifungal	<i>S. aureus</i> , <i>B. subtilis</i> <i>E. coli</i>	0 – 9 mm (5 – 10 µL) 0 mm (5 – 10 µL)			
			Antibacterial	<i>C. albicans</i> <i>M. tuberculosis</i>	0 mm (5 – 10 µL) MIC ₉₀ = 50 µM			
(-)-Callyaerin F 582b ^γ [C ₅₈ H ₈₃ N ₁₁ O ₁₀]	MS, NMR (ROESY)	Cyclo pentapeptide (dipeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 µM	<i>C. aerizusa</i>	SSW	[382]
(-)-Callyaerin H 583 ^β [C ₅₄ H ₈₁ N ₁₁ O ₁₀]	UV, MS, NMR, [α] _D , CT	Cyclo heptapeptide (dipeptide sc.)	Cytotoxic	<i>A. salina</i> L5178Y	30 – 55% (20 – 50 µg/mL) ED ₅₀ = 0.48 µM	<i>C. aerizusa</i>	MLU	[381]
			Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 µM			
(-)-Callyaerin I 584 ^β [C ₆₉ H ₉₃ N ₁₃ O ₁₂]	UV, MS, NMR, [α] _D , CT	Cyclo heptapeptide (tetrapeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 µM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 µM			
(-)-Callyaerin J 585 ^β [C ₆₆ H ₉₅ N ₁₃ O ₁₂]	UV, MS, NMR, [α] _D , CT	Cyclo heptapeptide (tetrapeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 µM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 µM			
(-)-Callyaerin K 586 ^β [C ₇₅ H ₉₅ N ₁₃ O ₁₂]	UV, MS, NMR, [α] _D , CT	Cyclo heptapeptide (tetrapeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 µM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 µM			
(-)-Callyaerin L 587 ^β [C ₆₆ H ₁₀₁ N ₁₃ O ₁₅]	UV, MS, NMR, [α] _D , CT	Cyclo octapeptide (tripeptide sc.)	Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 µM	<i>C. aerizusa</i>	MLU	[382]
			Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 µM			

Table S19: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Callyaerin M 588 ^β [C ₆₄ H ₉₅ N ₁₅ O ₁₅]	UV, MS, NMR, [α] _D , CT	Cyclo octapeptide (tetrapeptide sc.)	Antibacterial	<i>M. tuberculosis</i>	MIC ₉₀ > 100 μM	<i>C. aerizusa</i>	MLU	[382]
			Cytotoxic	THP-1, MRC-5	IC ₅₀ > 10 μM			
(+)–Microspinosamide 589 ^β [C ₇₅ H ₁₀₉ BrN ₁₈ O ₂₂ S]	UV, IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide (octapeptide sc.) [★]	Antiviral	CEM-SS infected HIV-1	EC ₅₀ = 0.2 μg/mL	<i>S. microspinosus</i>	NSW	[384]
			Cytotoxic	CEMS-SS	EC ₅₀ ~ 3.0 μg/mL			
(-)-Kendarimide 590 ^β [C ₈₃ H ₁₃₄ N ₁₄ O ₁₅ S ₂]	UV, IR, MS, NMR, [α] _D , CT	Linear trideca-peptide [★]	Cytotoxic	KB-C2, P-gp type	87% (KB-C2, 0.1 μg/mL colchine) (6 μM)	<i>Haliclona</i> sp.	SES	[385, 386]
				KB-3-1	NA in KB-3-1 (6 μM)			
(-)-Cyclolithistide A 591a ^β [C ₅₄ H ₈₆ ClN ₁₁ O ₁₅]	UV, IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide [★]	Antifungal	<i>C. albicans</i> (ATCC 24433)	90% (20 μg/disk)	<i>T. swinhoei</i>	NSW	[387]
			Antibacterial	<i>E. coli</i> , <i>B. subtilis</i>	NA			
(-)-Cyclolithistide A 591b ^γ [C ₅₄ H ₈₆ ClN ₁₁ O ₁₅]	UV, MS, GC-MS, NMR, HMBC, [α] _D , CT	Cyclo depsipeptide	Undetm.	Undetm.	Undetm.	<i>D. japonica</i>	JPN	[388]
			Cytotoxic	L1210	NA			
(-)–Theonellapeptolide Ile 592 ^β [C ₇₀ H ₁₂₅ N ₁₃ O ₁₆]	IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide	Cytotoxic			<i>T. swinhoei</i>	SSW	[379]
(-)–Sulfinyltheonellapeptolide 593 ^β [C ₆₉ H ₁₂₃ N ₁₃ O ₁₆ S]	MS, NMR, [α] _D , CT	Cyclo depsipeptide [★]	Cytostatic	HepG2	IC ₅₀ = 3 μM	<i>T. swinhoei</i>	NSW	[389]
(-)–Theonellapeptolide If 594 ^β [C ₆₉ H ₁₂₁ N ₁₃ O ₁₆]	MS, NMR, [α] _D , CT	Cyclo depsipeptide	Cytostatic	HepG2	IC ₅₀ = 3 μM	<i>T. swinhoei</i>	NSW	[389]
(-)-Celebeside A 595 ^β [C ₃₇ H ₆₂ N ₇ O ₁₆ P]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT	Cyclo depsipeptide [★]	Antiviral	HIV-1 SF162 envelope	IC ₅₀ = 1.9 ± 0.4 μg/mL	<i>S. mirabilis</i>	UEP	[390]
			Cytotoxic	HCT-116	IC ₅₀ = 8.8 ± 3.0 μg/mL			
(-)-Celebeside B 596 ^β [C ₃₆ H ₆₀ N ₇ O ₁₆ P]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT	Cyclo depsipeptide	Antibacterial	<i>B. subtilis</i> , <i>E. coli</i> , <i>P. aeruginosa</i> , <i>S. aureus</i>	NA (50 μg/disk)	<i>S. mirabilis</i>	UEP	[390]
			Antifungal	<i>C. albicans</i>	NA (50 μg/disk)			
(-)-Celebeside B 596 ^β [C ₃₆ H ₆₀ N ₇ O ₁₆ P]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT	Cyclo depsipeptide	Antibacterial	<i>B. subtilis</i> , <i>E. coli</i> , <i>P. aeruginosa</i> , <i>S. aureus</i>	NA (50 μg/disk)	<i>S. mirabilis</i>	UEP	[390]
			Antifungal	<i>C. albicans</i>	NA (50 μg/disk)			

Table S19: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Celebeside C 597 ^β [C ₃₇ H ₆₁ N ₇ O ₁₃]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT	Cyclo depsipeptide	Antiviral	HIV-1 SF162 envelope	IC ₅₀ > 50 µg/mL	<i>S. mirabilis</i>	UEP	[390]
			Cytotoxic	HCT-116	IC ₅₀ > 25 µg/mL			
			Antibacterial	<i>B. subtilis</i> , <i>E. coli</i> , <i>P. aeruginosa</i> , <i>S. aureus</i>	NA (50 µg/disk)			
			Antifungal	<i>C. albicans</i>	NA (50 µg/disk)			
(+) -Theopapuamide B 598 ^β [C ₇₁ H ₁₂₅ N ₁₇ O ₂₄]	UV, IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide	Antiviral	HIV-1 SF162 envelope	IC ₅₀ = 0.8 ± 0.3 µg/mL	<i>S. mirabilis</i>	UEP	[390]
			Cytotoxic	HCT-116	IC ₅₀ = 2.1 ± 0.7 µg/mL			
			Antifungal	<i>C. albicans</i> , <i>C. albicans</i> (amphotericin B-resistant)	10 mm (5 µg/disk)			
(+) -Theopapuamide C 599 ^β [C ₇₁ H ₁₂₅ N ₁₇ O ₂₃]	UV, IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide	Cytotoxic	HCT-116	IC ₅₀ = 4.0 ± 1.7 µg/mL	<i>S. mirabilis</i>	UEP	[390]
			Antifungal	<i>C. albicans</i> , <i>C. albicans</i> (amphotericin B-resistant)	10 mm (5 µg/disk)			
(+) -Theopapuamide D 600 ^β [C ₇₂ H ₁₂₇ N ₁₇ O ₂₃]	UV, IR, MS, NMR, [α] _D , CT	Cyclo depsipeptide	Cytotoxic	HCT-116	IC ₅₀ = 2.1 ± 0.9 µg/mL	<i>S. mirabilis</i>	UEP	[390]
(-) -Jaspamide Q 601 ^β [C ₃₆ H ₄₆ N ₄ O ₆]	UV, MS, NMR, [α] _D	Cyclo depsipeptide	Cytotoxic	L5178Y	IC ₅₀ < 0.1 µg/mL	<i>J. splendens</i>	EKM	[391]
(-) -Jaspamide R 602 ^β [C ₃₆ H ₄₄ Br ₂ N ₄ O ₆]	UV, MS, NMR, [α] _D	Cyclo depsipeptide	Cytotoxic	L5178Y	IC ₅₀ < 0.1 µg/mL	<i>J. splendens</i>	EKM	[391]
(+) -Daedophamide 603 ^β [C ₇₁ H ₁₂₅ N ₁₉ O ₂₂]	UV, IR, MS, NMR, [α] _D	Cyclo depsipeptide	Cytotoxic	MDA-MB-231, HT-29, A549, PSN-1	GI ₅₀ = 0.2 – 0.6 µM TGI = 0.3 – 0.8 µM LC ₅₀ = 0.6 – 1.3 µM	<i>Daedalopelta</i> sp.	ENT	[392]
			Cytotoxic	A2780, Jurkat, Ramos, Nomo-1, HL-60	IC ₅₀ = 0.45 – 1.90 µM			
				Apop. (caspase activation) Starvation-ind. autophagy	1 µM 10 µM			
(-) -Microcionamide C 604 ^β [C ₄₄ H ₇₄ N ₈ O ₆ S ₂]	UV, MS, NMR, [α] _D	Cyclo pentapeptide ^Δ	Antibacterial	<i>E. faecium</i> BM4147-1 <i>S. aureus</i> ATCC29213 <i>M. tuberculosis</i> H37Rv, <i>K. pneumoniae</i> ATCC 12657, <i>E. aerogenes</i> ATCC 13048, <i>E. coli</i> ATCC 25922, <i>P. aeruginosa</i> ATCC 27853, <i>A. baumannii</i> 09987	MIC = 12.5 µM MIC = 6.3 µM	<i>C. basilana</i>	MLU	[357]
					MIC > 100			

Table S19: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Microcionamide D 605 ^β [C ₄₄ H ₇₄ N ₈ O ₆ S ₂]	UV, MS, NMR, [α] _D	Cyclo pentapeptide	Cytotoxic	A2780, Ramos, Jurkat, Nomo-1, HL-60	IC ₅₀ = 0.53 – 2.50 μM	<i>C. basilana</i>	MLU	[357]
			Cytotoxic	Apop. (caspase activation) starvation-ind. autophagy	1 μM 10 μM			
(+)–Gombamide B 606 ^β [C ₅₀ H ₆₇ N ₉ O ₉ S ₂]	UV, MS, NMR, [α] _D	Cyclo tetrapeptide (tripeptide sc)	Antibacterial	A2780, Ramos, Jurkat, Nomo-1, HL-60 <i>E. faecium</i> BM4147-1, <i>S. aureus</i> ATCC29213 <i>M. tuberculosis</i> H37Rv, <i>K. pneumoniae</i> ATCC 12657, <i>E. aerogenes</i> ATCC 13048, <i>A. baumannii</i> 09987, <i>P. aeruginosa</i> ATCC 27853	MIC > 50 MIC > 100	<i>C. basilana</i>	MLU	[357]
			Cytotoxic	Ramos, Jurkat, Nomo-1, HL-60	NA			
(+)–Gombamide C 607 ^β [C ₅₀ H ₆₇ N ₉ O ₉ S ₂]	UV, MS, NMR, [α] _D	Cyclo tetrapeptide (tripeptide sc)	Antibacterial	<i>E. faecium</i> BM4147-1, <i>S. aureus</i> ATCC29213 <i>M. tuberculosis</i> H37Rv, <i>K. pneumoniae</i> ATCC 12657, <i>E. aerogenes</i> ATCC 13048, <i>A. baumannii</i> 09987, <i>P. aeruginosa</i> ATCC 27853	MIC > 50 MIC > 100	<i>C. basilana</i>	MLU	[357]
			Cytotoxic	Ramos, Jurkat, Nomo-1, HL-60	NA			
(-)-Gombamide D 608 ^β [C ₃₂ H ₄₈ N ₈ O ₈ S ₂]	UV, MS, NMR, [α] _D	Cyclo tetrapeptide	Antibacterial	<i>E. faecium</i> BM4147-1, <i>S. aureus</i> ATCC29213 <i>K. pneumoniae</i> ATCC 12657, <i>E. aerogenes</i> ATCC 13048, <i>P. aeruginosa</i> ATCC 13048	MIC > 50 MIC > 100	<i>C. basilana</i>	MLU	[357]
(-)–Diazonamide C 609 ^β [C ₄₀ H ₃₅ Cl ₂ N ₇ O ₅]	MS, NMR, [α] _D	Macrocyclic peptide	Cytotoxic	A549, MDA-MB-231, HT-29	GI ₅₀ = 1.8 – 2.2 μM	<i>Diazona</i> sp.	PUA	[393]
(-)–Diazonamide D 610 ^β [C ₃₆ H ₂₇ Cl ₃ N ₆ O ₄]	MS, NMR, [α] _D	Macrocyclic peptide	Cytotoxic	A549, HT-29, MDA-MB-231	GI ₅₀ = 2.9 – 3.1 μM	<i>Diazona</i> sp.	PUA	[393]

Table S19: *Cont.*

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Diazonamide E 611 ^β [C ₃₆ H ₂₇ BrCl ₂ N ₆ O ₄]	MS, NMR, [α] _D	Macrocyclic peptide	Cytotoxic	A549, HT-29, MDA-MB-231	GI ₅₀ = 1.8 – 2.2 μM	<i>Diazona</i> sp.	PUA	[393]

Footnote: 1. **Statistic** (TGI total growth inhibition); 2. **Activity** (**HepG2** human hepatocarcinoma, **LNCaP** in human prostate carcinoma, **MRC-5** human fetal lung fibroblast, **Nomo-1** human adult monocyclic leukemia, **Ramos** human Burkitt lymphoma, **RAW 264** murine macrophage, **AR NTD** androgen receptor N-terminus domain, **HIV-1** human immunodeficiency virus 1).

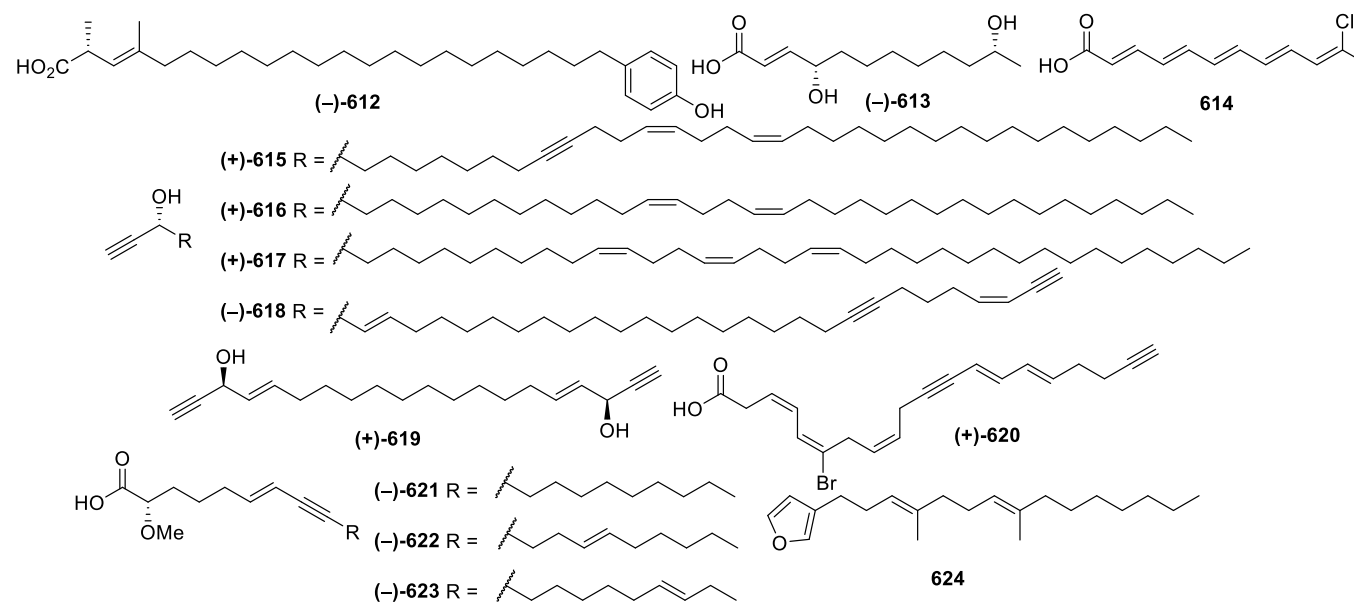


Figure S20: Structures of marine fatty acids and linear molecules from Indonesian waters found in 1970–2017.

Table S20: Marine fatty acids and linear molecules from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Elenic acid 612 ^β [C ₃₀ H ₅₀ O ₃]	UV, IR MS, NMR, [α] _D , CT, TS	Fatty acid [▲]	Cytotoxic	P-388, A549, MEL-28	IC ₅₀ = 5 µg/mL	<i>Plakinastrella</i> sp.	NSW	[394]
			Cytotoxic	Topoisomerase II	0.1 µg/mL			– [396]
(-)- <i>seco</i> -Patulolide C 613 ^β [C ₁₂ H ₂₂ O ₄]	UV, MS, NMR, [α] _D , CT, TS	Fatty acid	Antihyperlipidemic	A549, HeLa, SMMC-7721	IC ₅₀ > 50 µM	A fungus (symbiont) a sponge (host)	SSW	[397]
			Antibacterial	HepG2	IC ₅₀ = 13.1 µM			– [399]
Aurantoic acid 614 ^β [C ₁₂ H ₁₃ ClO ₂]	UV, MS, NMR	Fatty acid	Cytotoxic	Gram-positive, Gram-negative bacteria	NA (250 µg/disk)	<i>T. swinhoei</i>	NSW	[212]
			Cytotoxic	C6, HeLa, H9c2	IC ₅₀ > 70 µM			
(+)–Lembehyne A 615 ^β [C ₃₆ H ₆₂ O]	IR, MS, NMR, [α] _D , CT, TS	Poly acetylenene	Neuro disease	Neurite outgrowth (PC12)	2 µg/mL	<i>Haliclona</i> sp.	NSW	[400]
			Neuro disease	Neuritogenesis (Neuro2A)	<i>ca.</i> 60% inducer (3.0 µg/mL)			– [403]
(+)–Lembehyne B 616 ^β [C ₃₆ H ₆₆ O]	IR, MS, NMR, [α] _D , CT, TS	Poly acetylenene	Neuro disease	Enhancer cyclic dependent kinase inhibitor (p21/WAF1)	G1 arres.	<i>Haliclona</i> sp.	NSW	[401]
			Cytotoxic	Neuritogenesis (Neuro2A)	<i>ca.</i> 60% inducer (3.0 µg/mL)			– [407]
(+)–Lembehyne C 617 ^β [C ₃₆ H ₆₆ O]	IR, MS, NMR, [α] _D , CT, TS	Poly acetylenene	Neuro disease	Jurkat	IC ₅₀ = 2.0 µM, 72.6% apop. (2.0 µM)	<i>Haliclona</i> sp.	NSW	[404]
			Neuro disease	HL-60 K562	IC ₅₀ = 2.2 µM IC ₅₀ = 3.0 µM			
(-)– 618 ^β [C ₃₁ H ₄₈ O]	IR, MS, NMR, [α] _D , CT	Poly acetylenene	Cytotoxic	Neuritogenesis (Neuro2A)	<i>ca.</i> 60% inducer (3.0 µg/mL)	<i>Haliclona</i> sp.	NSW	[404]
(+)–(3 <i>S</i> ,18 <i>S</i> ,4 <i>E</i> ,16 <i>E</i>)-eicosa-1,19-diyne-3,18-diol-4,16-diene 619 ^β [C ₂₀ H ₃₀ O ₂]	IR, MS, NMR, [α] _D , CT	Poly acetylenene	Cytotoxic	NBT-T2	5 – 10 µg/mL	<i>Callyspongia</i> sp.	ENT	[408]
620 ^β [C ₂₀ H ₂₁ BrO ₂]	IR, MS, NMR, [α] _D	Poly acetylenene [▲]	Cytotoxic	<i>A. salina</i>	LD ₅₀ = 2.0 µg/mL	<i>C. pseudo-reticulata</i>	SSW	[409]
				NBT-T2	IC ₅₀ = 36 µg/mL	<i>Haliclona</i> sp.	ENT	[410]

Table S20: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Cinachylenic acid B 621 ^β [C ₁₉ H ₃₂ O ₃]	UV, IR, MS, NMR, [α] _D	Poly acetylenene	Cytotoxic	L5178Y	IC ₅₀ = 0.3 μM	<i>Cinachyrella</i> sp.	MLU	[355]
(-)-Cinachylenic acid C 622 ^β [C ₁₉ H ₃₀ O ₃]	UV, IR, MS, NMR, [α] _D	Poly acetylenene	Cytotoxic	L5178Y	IC ₅₀ = 0.3 μM	<i>Cinachyrella</i> sp.	MLU	[355]
(-)-Cinachylenic acid D 623 ^β [C ₁₉ H ₃₀ O ₃]	UV, IR, MS, NMR, [α] _D	Poly acetylenene	Cytotoxic	L5178Y	IC ₅₀ = 0.3 μM	<i>Cinachyrella</i> sp.	MLU	[355]
3-[(3 <i>E</i> ,7 <i>E</i>)-4,8-dimethylpentadeca-3,7dienyl]furan 624 ^β [C ₂₁ H ₃₄ O]	UV, IR, MS, NMR	Furanolipid	Antitumor	HIF-1 (T47D)	NA (10 μM)	<i>Lendenfeldia</i> sp.	UEP	[411]

Footnote: : 1. Activity (HIF-1 hypoxia-inducible factor-1).

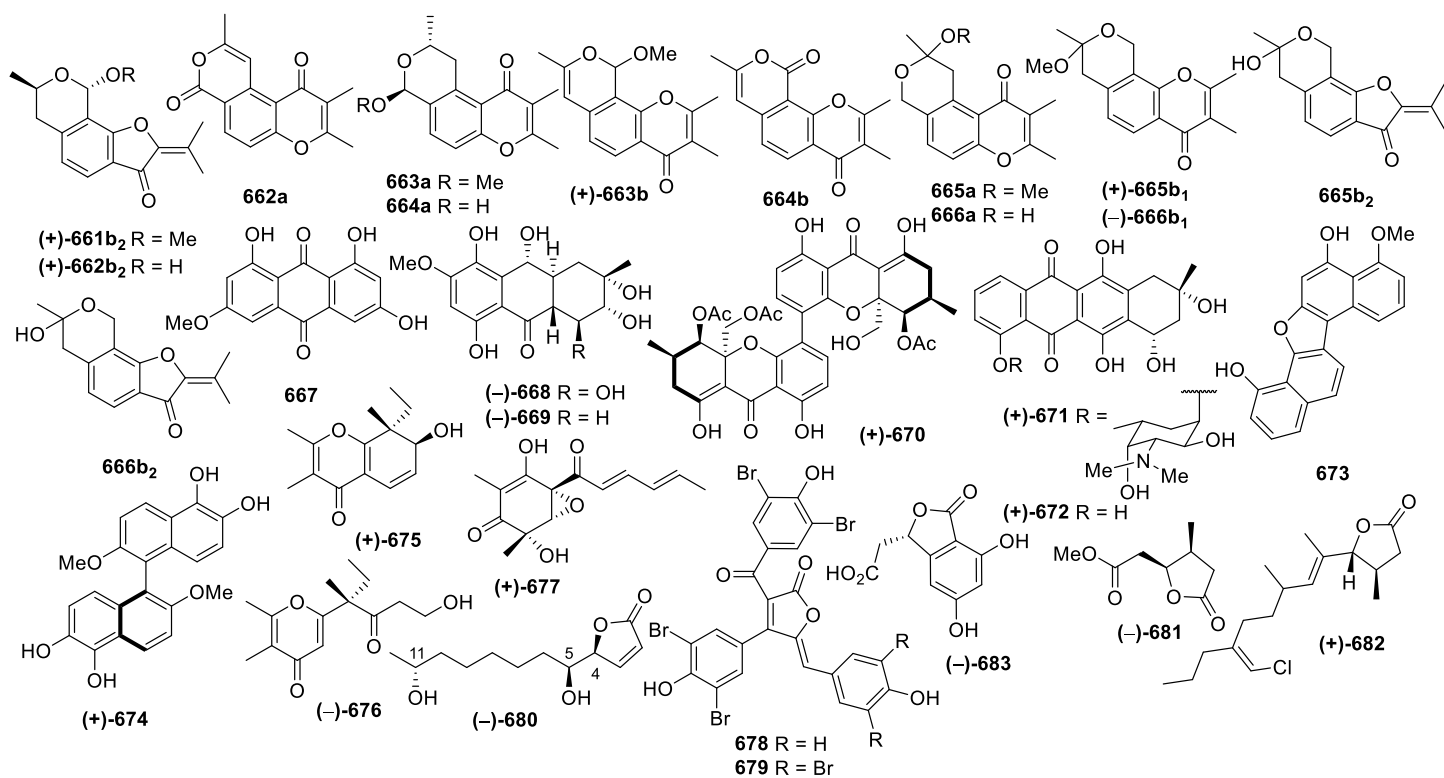


Figure S21: *Cont.*

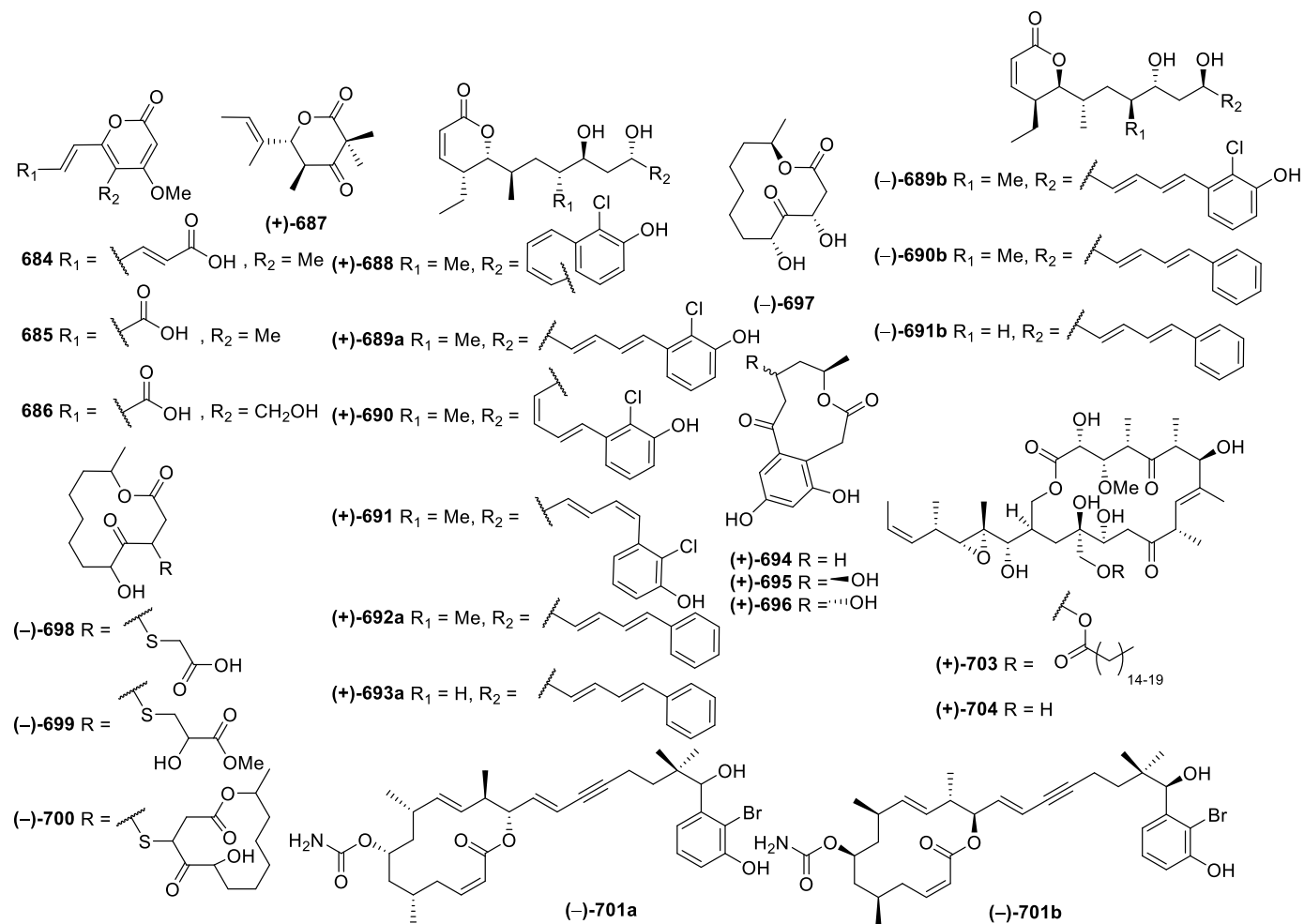


Figure S21: Cont.

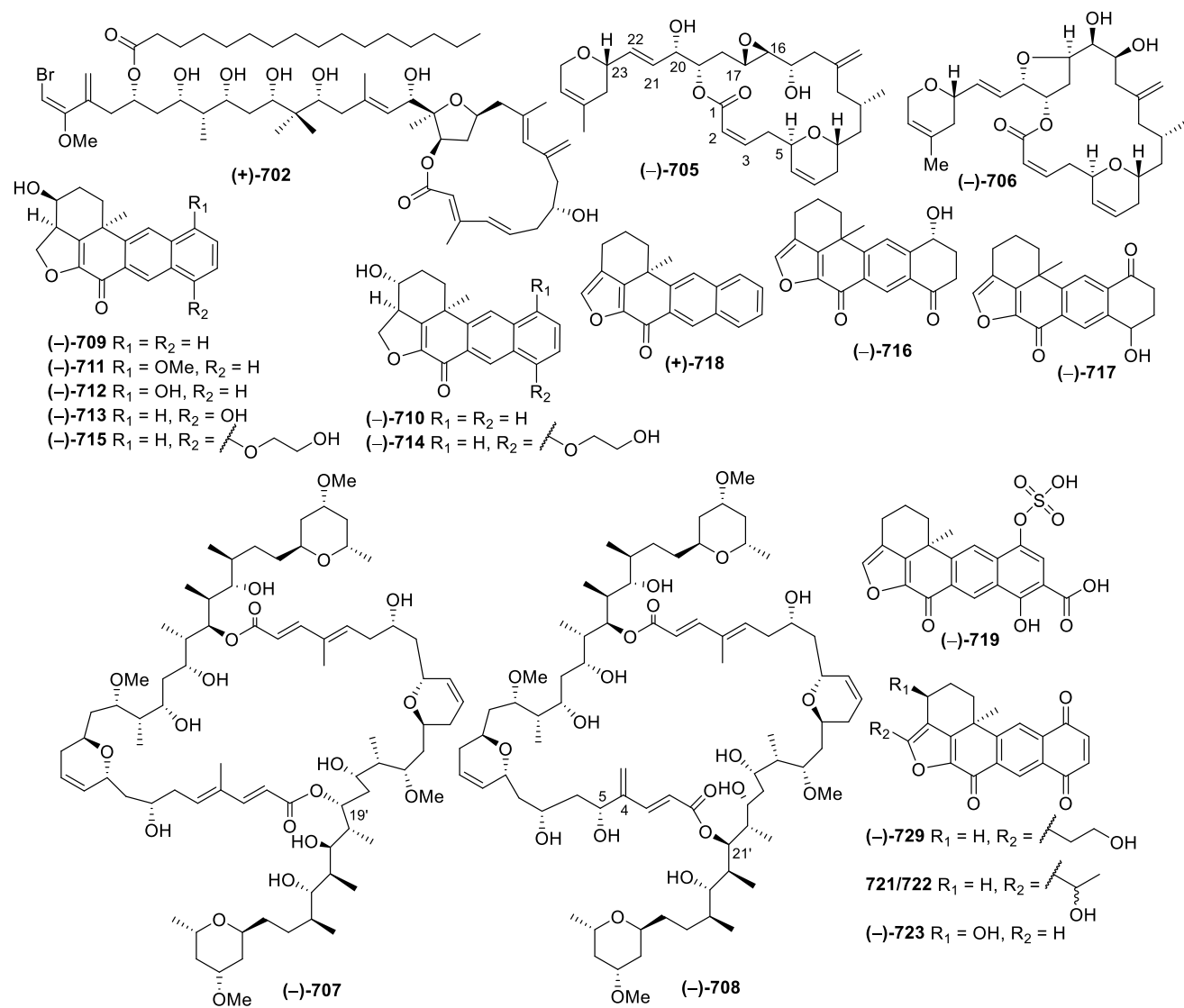


Figure S21: Cont.

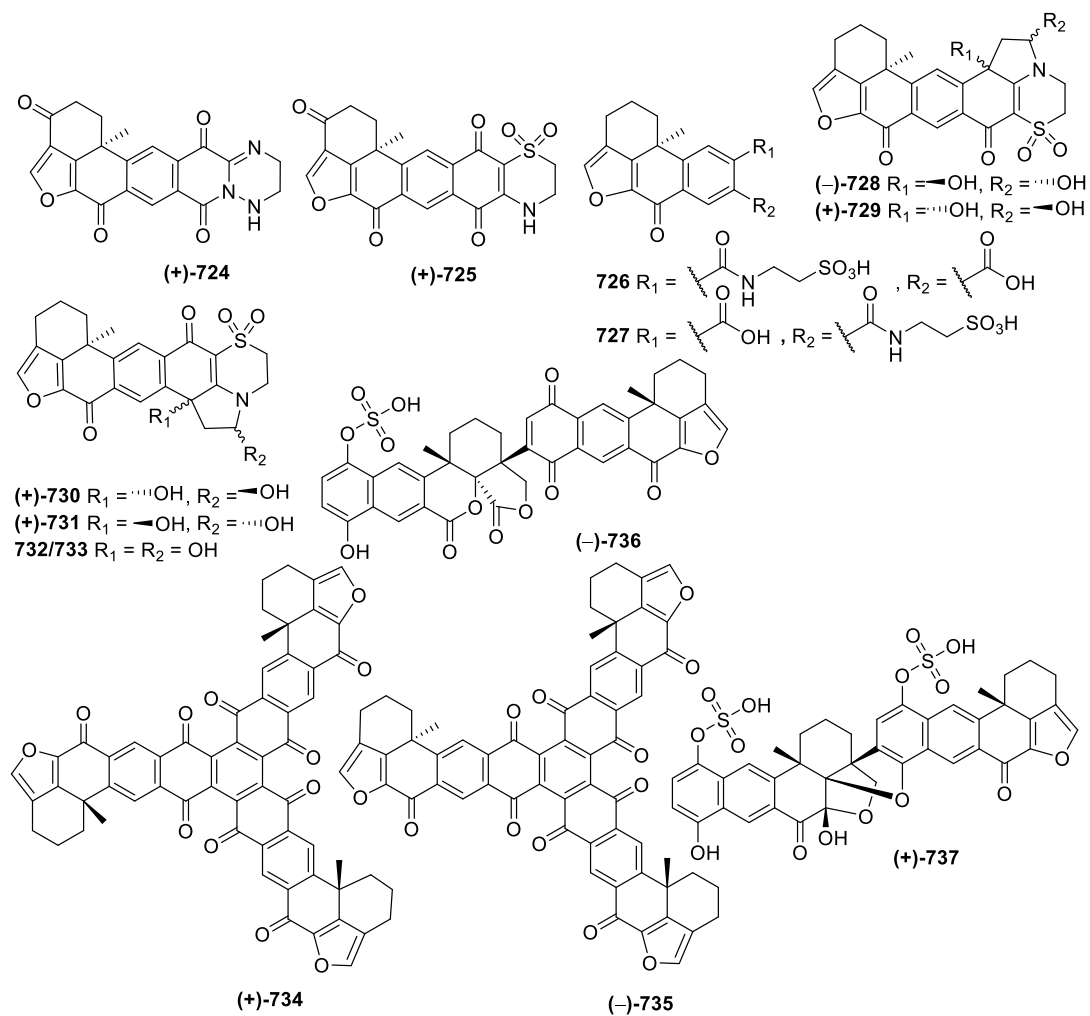


Figure S21: *Cont.*

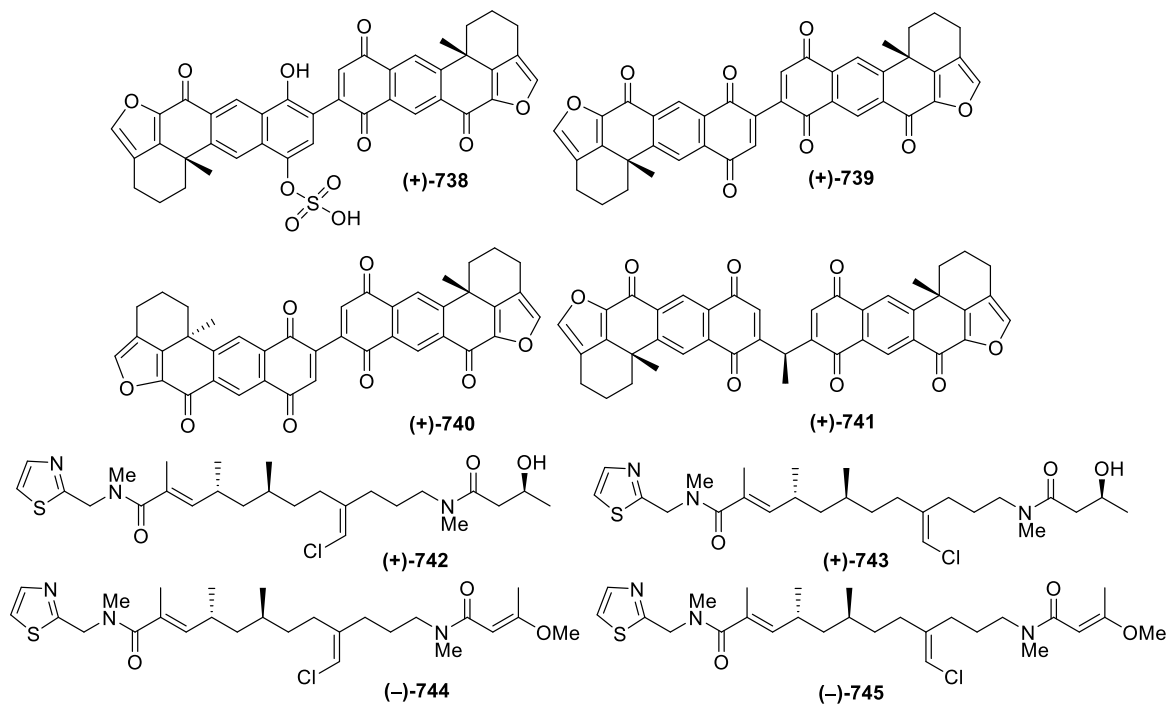


Figure S21: Structures of polyketides from Indonesian waters found in 1970–2017.

Table S21: Marine polyketides from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Karatungiol A 625 ^β [C ₇₃ H ₁₃₂ O ₂₈]	UV, IR, MS, NMR, [α] _D , CT	Polyol	Antifungal	<i>A. niger</i> NBRC4407 <i>T. foetus</i>	12 µg/disk 1 µg/mL	<i>Amphidinium</i> sp.	NSW	[44]
(+)-Karatungiol B 626 ^β [C ₇₃ H ₁₃₀ O ₂₇]	UV, IR, MS, NMR, [α] _D	Polyol	Undetm.	Undetm.	Undetm.	<i>Amphidinium</i> sp.	NSW	[44]
(+)-Manadic acid A 627 ^β [C ₁₇ H ₂₈ O ₅]	UV, IR, MS, NMR, [α] _D	Cyclic peroxide	Cytotoxic	P-388, A459, HT-29, LcV, MEL-28 MLR	IC ₅₀ = 0.5 – 5 µM IC ₅₀ = 0.015 µM	<i>Plakortis</i> sp.	NSW	[412]
(+)-Manadic acid B 628 ^β [C ₁₈ H ₃₀ O ₅]	UV, IR, MS, NMR, [α] _D , CT	Cyclic peroxide	Cytotoxic	P-388, A549, HT-29, MEL-28 MLR, LcV	IC ₅₀ = 0.5 – 5 µM NA	<i>Plakortis</i> sp.	NSW	[412]
(-)-(9S, 10S)-Plakorstatin 1 629 /(-)-(9R, 10R)-Plakorstatin 1 630 ^{α,β} [C ₁₉ H ₃₂ O ₅]	MS, NMR, [α] _D	Cyclic peroxide	Cytotoxic	P-388 BXPC-3, MCF7, NCI-H460, KM-20L2, DU-145 SF268	ED ₅₀ = 1.1 µg/mL GI ₅₀ > 10 µg/mL GI ₅₀ > 1.8 µg/mL	<i>P. nigra</i>	NSW	[413]
(-)-(9R, 10R)-Plakorstatin 1 631 /(-)-(9S, 10S)-Plakorstatin 1 632 ^{α,β} [C ₁₉ H ₃₂ O ₅]	MS, NMR, [α] _D	Cyclic peroxide	Cytotoxic	P-388 BXPC-3, KM-20L2 MCF7 SF268, DU-145 NCI-H460	ED ₅₀ = 0.91 µg/mL GI ₅₀ = 6.4 – 6.7 µg/mL GI ₅₀ = 3.8 µg/mL GI ₅₀ = 1.6 – 1.7 µg/mL GI ₅₀ > 10 µg/mL	<i>P. nigra</i>	NSW	[413]
(-)-Manadoperoxide A 633 ^β [C ₁₈ H ₃₀ O ₅]	UV, IR, MS, NMR, [α] _D , Mol. Mod., CT	Cyclic peroxide	Antiparasite	<i>P. falciparum</i> (D10), (W2)	IC ₅₀ = 3.74 ± 0.92 – 6.88 ± 0.37 µM	<i>P. cfr. simplex</i>	NSW	[414]
(-)-Manadoperoxide B 634 ^β [C ₁₉ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D , Mol. Mod.	Cyclic peroxide	Antiparasite	<i>P. falciparum</i> (D10), (W2) <i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 3.69 ± 0.88 – 6.76 ± 0.32 µM IC ₅₀ = 0.003 µg/mL IC ₅₀ = 0.589 µg/mL	<i>P. cfr. simplex</i>	NSW	[414, 415]
(-)-Manadoperoxide C 635 ^β [C ₁₆ H ₂₆ O ₆]	UV, IR, MS, NMR, [α] _D , Mol. Mod	Cyclic peroxide	Antiparasite	<i>P. falciparum</i> (D10), (W2) <i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 2.33 ± 0.48 – 4.54 ± 0.66 µM IC ₅₀ = 0.678 µg/mL IC ₅₀ = 3.24 µg/mL	<i>P. cfr. simplex</i>	NSW	[414, 415]
(-)-Manadoperoxide D 636 ^β [C ₁₈ H ₃₂ O ₇]	UV, IR, MS, NMR, [α] _D , Mol. Mod	Cyclic peroxide	Antiparasite	<i>P. falciparum</i> (D10), (W2) <i>T. brucei rhodesiense</i>	IC ₅₀ = 7.93 ± 0.68 – 10.38 ± 0.76 µM IC ₅₀ = 19.2 – 36.7 µg/mL	<i>P. cfr. simplex</i>	NSW	[414, 415]
(-)-Manadoperoxide E 637 ^β [C ₁₉ H ₃₄ O ₇]	MS, NMR, [α] _D , CT	Cyclic peroxide	Undetm.	Undetm.	Undetm.	<i>P. cfr. simplex</i>	NSW	[415]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Manadoperoxide F 638 ^β [C ₁₉ H ₃₄ O ₇]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 0.792 µg/mL IC ₅₀ = 5.73 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Manadoperoxide G 639 ^β [C ₁₈ H ₃₀ O ₈]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 1.84 µg/mL IC ₅₀ = 3.22 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Manadoperoxide H 640 ^β [C ₁₉ H ₃₄ O ₆]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 0.375 µg/mL IC ₅₀ = 2.44 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Manadoperoxide I 641 ^β [C ₁₈ H ₃₀ O ₇]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 0.062 µg/mL IC ₅₀ = 0.633 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Manadoperoxide J 642 ^β [C ₁₉ H ₃₁ ClO ₇]	MS, NMR, [α] _D	Cyclic peroxide	Undetm.	Undetm.	Undetm.	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Manadoperoxide K 643 ^β [C ₂₀ H ₃₅ ClO ₇]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 0.087 µg/mL IC ₅₀ = 1.89 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
(-)-Peroxyplakoric ester C 644 ^β [C ₁₆ H ₂₆ O ₆]	MS, NMR, [α] _D	Cyclic peroxide	Antiparasite	<i>T. brucei rhodesiense</i> <i>L. donovani</i>	IC ₅₀ = 30.9 µg/mL IC ₅₀ = 43.4 µg/mL	<i>P. cfr.</i> <i>simplex</i>	NSW	[415]
14,15- <i>seco</i> -Curvularin 645 ^β [C ₁₆ H ₂₂ O ₅]	MS, NMR	Aromatic Polyketide [▲]	Antibacterial	<i>B. subtilis</i>	20% (200 µg/disk)	A fungus (symbiont) <i>S. vagabunda</i> (host)	CSW	[416]
(-)-Vidalenolone 646 ^β [C ₁₃ H ₁₄ O ₄]	UV, IR, MS, NMR, [α] _D , ECD, QCC	Aromatic Polyketide	Anticancer	Fyn-SH2	NA	<i>Vidalia</i> sp.	UEP	[417, 418]
Cosmochlorin A 647 ^β [C ₁₈ H ₁₈ Cl ₂ O ₄]	UV, IR, MS, NMR	Aromatic Polyketide	Antibacterial	<i>S. aureus</i> NRBC 13276 <i>C. albicans</i> ATCC 2019 <i>A. clavatus</i> F318a,	MIC = 15.6 µg/mL MIC = 125 µg/mL	<i>C. vilior</i> IM2-155 (symbiont) <i>S. alba</i> (host)	WJV	[419]
			Antifungal	<i>T. harzianum</i> NBRC 33016 <i>V. dahliae</i> Klebahn NBRC 9470	MIC = 15.6 – 62.5 µg/mL MIC > 125 µg/mL			
			Cytotoxic	HL-60	IC ₅₀ = 73.7 µM			
			Growth restore activity	<i>S. cerevisiae</i> YNS17 (0.3 M CaCl ₂) GSK-3β	5 µg IC ₅₀ = 62.5 µM			
			Cytotoxic	HL-60	IC ₅₀ = 53.6 µM			
Cosmochlorin B 648 ^β [C ₁₈ H ₁₈ Cl ₂ O ₄]	UV, IR, MS, NMR	Aromatic Polyketide	Growth restore activity	<i>S. cerevisiae</i> YNS17 (0.3 M CaCl ₂) GSK-3β	1.25-5 µg IC ₅₀ = 60.6 µM	<i>C. vilior</i> IM2-155 (symbiont) <i>S. alba</i> (host)	WJV	[419]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Cosmochlorin C 649 ^β [C ₁₈ H ₁₈ Cl ₂ O ₄]	UV, IR, MS, NMR	Aromatic Polyketide	Antibacterial	<i>S. aureus</i> NRBC 13276	MIC = 15.6 µg/mL	<i>C. vilior</i> IM2-155 (symbiont) <i>S. alba</i> (host)	WJV	[419]
			Antifungal	<i>C. albicans</i> ATCC 2019	MIC = 125 µg/mL			
				<i>A. clavatus</i> F318a	MIC = 62.5 µg/mL			
				<i>T. harzianum</i> NBRC 33016	MIC = 15.6 µg/mL			
				<i>V. dahliae</i> Klebahn NBRC 9470	MIC > 125 µg/mL			
Acetyl sumiki's acid 650 ^β [C ₈ H ₈ O ₅]	MS, NMR	Aromatic Polyketide	Antibacterial	<i>B. subtilis</i> , <i>S. aureus</i>	7 mm (5 µg)	<i>C. herbarum</i> (symbiont) <i>C. aerizusa</i> (host)	BLI	[420]
				<i>E. coli</i> , <i>C. albicans</i>	NA			
2-Carboxy-8-methoxy-naphthalene-1-ol 651 ^β [C ₁₂ H ₁₀ O ₄]	MS, NMR	Aromatic Polyketide	Antibacterial	<i>S. aureus</i> , <i>P. aeruginosa</i> , <i>E. coli</i>	NA (100 µg/disk)	<i>M. sterillum</i>	UEP	[421]
			Antifungal	<i>C. maltosa</i>	NA (200 µg/disk)			
			Cytotoxic	5637 (ATCC HTB-9)	NA (IC ₅₀ = 0.34 mM)			
3,4,5-Tribromo-2-(2'-bromophenoxy)phenol 652 ^β [C ₁₂ H ₆ Br ₄ O ₂]	UV, MS, NMR	Aromatic polyketide	Antibacterial	<i>B. subtilis</i>	MIC = 6.25 µg/mL	<i>D. herbacea</i>	WST	[422]
			Antifungal	<i>C. cucumerinum</i>	8.0 – 13.0 mm (25 – 50 mmol)			
			Cytotoxic	<i>A. salina</i>	LC ₅₀ = 3.30 ± 0.51 µg/mL			
3,5,6-Tribromo-2-(2'-bromophenoxy)phenol 653 / 3,4,6-Tribromo-2-(2'-bromophenoxy)phenol 654 ^{α,β} [C ₁₂ H ₆ Br ₄ O ₂]	UV, MS, NMR	Aromatic polyketide	Antibacterial	<i>B. subtilis</i>	MIC = 1.56 µg/mL	<i>D. herbacea</i>	WST	[422]
			Antifungal	<i>C. cucumerinum</i>	8.0 – 13.0 mm (25 – 50 mmol)			
			Cytotoxic	<i>A. salina</i>	LC ₅₀ = 3.30 ± 0.35 µg/mL			
3,5,6-Tribromo-1-(2'-bromophenoxy)-2-benzene methyl ether 655 ^β [C ₁₃ H ₈ Br ₄ O ₂]	UV, MS, NMR	Aromatic polyketide	Antibacterial	<i>B. subtilis</i>	MIC = 104.00 µg/mL	<i>D. herbacea</i>	WST	[422]
			Cytotoxic	<i>A. salina</i>	LC ₅₀ = 26.25 ± 0.42 µg/mL			
656 ^β [C ₁₃ H ₇ Br ₅ O ₃]	UV, MS, NMR, CT	Aromatic polyketide	Antibacterial	<i>B. subtilis</i>	6 – 14 mm (0.1 – 10 µg/disk)	<i>L. herbacea</i>	BTN	[423]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
657^β [C ₁₂ H ₆ Br ₄ O ₃]	UV, MS, NMR, CT	Aromatic Polyketide*	Antibacterial	<i>B. subtilis</i> NBT-T2	7-20 mm (0.1 – 10 µg/disk) IC ₅₀ >15 µg/mL	<i>L. herbacea</i>	BTN	[423]
658^β [C ₁₃ H ₇ Br ₅ O ₃]	UV, MS, NMR, chemical transformation	Aromatic Polyketide	Antibacterial Cytotoxic	<i>B. subtilis</i> NBT-T2	7 – 17 mm (0.1 – 10 µg/disk) IC ₅₀ >15 µg/mL	<i>L. herbacea</i>	BTN	[423]
659a^β [C ₁₃ H ₇ Br ₅ O ₃]	UV, MS, NMR	Aromatic Polyketide	Antibacterial	<i>B. subtilis</i>	6 – 13 mm (0.1 – 10 µg/disk)	<i>L. herbacea</i>	BTN	[423]
659b^δ [C ₁₃ H ₇ Br ₅ O ₃]	X-ray	Aromatic Polyketide	Anticancer	Mc1-1	IC ₅₀ >10 µg/mL	<i>L. herbacea</i>	PNG	[423, 424]
			Antibacterial	<i>B. subtilis</i>	6 – 13 mm (0.1 – 10 µg/disk)			
2-hydroxy-6-(2'-hydroxy-3'-hydroxymethyl-5-methylphenoxy)-benzoic acid 660^β [C ₁₅ H ₁₄ O ₆]	UV, MS, NMR	Aromatic Polyketide*	Antidiabetic	PTP1B, TCPTP, VHR CD45	IC ₅₀ >35 µg/mL IC ₅₀ >43 µg/mL	<i>P. albobiverticillium</i> (symbiont) a tunicate (host)	NSW	[425]
Aspergione A 661a^β [C ₁₆ H ₁₆ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]
(+)-Aspergione A 661b¹⁷ [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[272]
(+)-Aspergione A = (+)-Ustusorane C 661b^{2δ} [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR, [α] _D , TS	Aromatic Polyketide	Cytotoxic	A549, HL-60	IC ₅₀ > 100 µM	<i>A. ustus</i> 094102	PRC	[426, 427]
Aspergione B 662a^β [C ₁₅ H ₁₂ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Aspergione B 662b ¹⁷ [C ₁₅ H ₁₆ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[272]
(+)-Aspergione B = (+)-pseudodeflectusin 662b ²⁶ [C ₁₅ H ₁₆ O ₄]	TS	Aromatic Polyketide	Undetm.	Undetm.	Undetm.			[427, 428]
Aspergione C 663a ^β [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]
(+)–Aspergione C 663b ⁷ [C ₁₆ H ₁₆ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Antibacterial	<i>B. subtilis</i> , <i>E. coli</i>	NA	<i>A. versicolor</i> (symbiont)	BLI	[272]
			Antifungal	<i>S. cerevisiae</i>	NA	<i>X. exigua</i> (host)		
Aspergione D 664a ^β [C ₁₅ H ₁₆ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]
Aspergione D 664b ⁷ [C ₁₅ H ₁₂ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[272]
Aspergione E 665a ^β [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]
(+)–Aspergione E 665b ¹⁷ [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Antibacterial	<i>B. subtilis</i> , <i>E. coli</i>	NA	<i>A. versicolor</i> (symbiont)	BLI	[272]
			Antifungal	<i>S. cerevisiae</i>	NA	<i>X. exigua</i> (host)		

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Aspergione E = pergillin 665b ^δ [C ₁₅ H ₁₆ O ₄]	TS	Aromatic Polyketide	Undetm.	Undetm.	Undetm.			[429]
Aspergione F 666a ^β [C ₁₅ H ₁₆ O ₄]	UV, MS, NMR	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[271]
(-)-Aspergione F 666b ¹⁷ [C ₁₆ H ₁₈ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>A. versicolor</i> (symbiont) <i>X. exigua</i> (host)	BLI	[272]
Aspergione F = pergillin 666b ₂ = 666b ^δ [C ₁₅ H ₁₆ O ₄]	TS	Aromatic Polyketide	Undetm.	Undetm.	Undetm.			[429]
Lunatin 667 ^β [C ₁₅ H ₁₀ O ₆]	MS, NMR	Aromatic Polyketide	Antibacterial	<i>S. aureus</i> <i>E. coli</i> <i>E. coli</i> HBI-101 <i>B. subtilis</i> <i>C. albicans</i>	8.5 – 10.0 mm (5 – 10 µg) 9.0 – 11.0 mm (5 – 10 µg) 8.0 – 10.5 mm (5 – 10 µg) 7.5 – 9.0 mm (5 – 10 µg) NA	<i>C. lunata</i> (symbiont) <i>N. olemda</i> (host)	BLI	[430]
(-)-Tetrahydrobostrycin 668 ^β [C ₁₆ H ₂₀ O ₈]	UV, IR, MS, NMR, [α] _D	Aromatic Polyketide	Antifungal					
			Antibacterial	<i>S. aureus</i> , <i>E. coli</i>	9.2 – 15 mm (100 µg/disk)	<i>Aspergillus</i> sp.	NSW	[431]
			Antibacterial	<i>S. aureus</i>	12 mm (100 µg/disk)	<i>Aspergillus</i> sp.		
(-)-1-Deoxytetrahydrobostrycin 669 ^β [C ₁₆ H ₂₀ O ₇]	UV, IR, MS, NMR, [α] _D	Aromatic Polyketide	Antifungal	<i>S. cerevisiae</i> , <i>M. hiemalis</i>	NA (100 µg/disk)	(symbiont) an alga (host)	NSW	[431]
(+)-12-O-Deacetyl-phomoxanthone A 670 ^β [C ₃₆ H ₃₆ O ₁₅]	UV, IR, MS, NMR, ECD, [α] _D , CT	Aromatic Polyketide	Antifungal	<i>T. harzianum</i> NBRC 33016, <i>V. dahliae</i> Klebahn NBRC 9470, <i>D. medusae</i> Nitschke NBRC 30895 <i>S. sclerotiorum</i> de Bary NBRC 103652, <i>B. cinerea</i> Persoon NBRC 100717	NA 11–12 mm (30 µg)	<i>Phomopsis</i> sp. (symbiont) <i>R. mucronata</i> (host)	JSCR	[432 – 433]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+) -12-O-Deacetyl-phomoxanthone A 670 ^β [C ₃₆ H ₃₆ O ₁₅]	UV, IR, MS, NMR, ECD, [α] _D , CT	Aromatic Polyketide	Antibacterial	<i>S. aureus</i> NBRC 13276 <i>P. aeruginosa</i> ATCC 15442	9 mm (30 μg) NA	<i>Phomopsis</i> sp. (symbiont)	JSCR	[432 – 433]
			Cytotoxic	L5178Y	IC ₅₀ = 2.8 μM	<i>R. mucronata</i> (host)		
(+) -Komodoquinone A 671 ^β [C ₂₈ H ₃₃ NO ₉]	UV, IR, MS, NMR, [α] _D , CT	Aromatic Polyketide [•]	Anticancer	Neuro 2A	1 μg/mL (morp. cng. mltplr. proc.) 1 μg/mL (G1 phase)	<i>Streptomyces</i> sp. KS3	ENT	[434, 435]
(+) -Komodoquinone B 672 ^β [C ₁₉ H ₁₆ O ₇]	UV, IR, MS, NMR, [α] _D , CT	Aromatic Polyketide	Anticancer	Neuro 2A	NA (3 μg/mL)	<i>Streptomyces</i> sp. KS3 A fungus (<i>Xylariales</i> , symbiont) a sponge (host)		
Xylarianaphthol-1 673 ^β [C ₂₁ H ₁₄ O ₄]	IR, MS, NMR, TS	Aromatic Polyketide [•]	Anticancer	Transfected MG63	Activator p21 promoter (0.3 μM)		UEP	[436]
(+) - (S)-2,2'-dimethoxy-1,1'-binaphthyl-5,5',6,6'-tetraol 674 ^β [C ₂₂ H ₁₈ O ₆]	UV, IR, MS, NMR, [α] _D , ECD	Aromatic Polyketide	Antitumor	HIF-1 (T47D)	IC ₅₀ = 4.3 μM	<i>Lendenfeldia</i> sp.	UEP	[411]
			Cytotoxic	MDA-MB-231 – T47D (hypoxic)	IC ₅₀ = 7.0 – 8.3 μM			
(+) -Spiciferol A 675 ^β [C ₁₄ H ₁₈ O ₃]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>D. hawaiiensis</i> (symbiont) <i>C. aerizusa</i> (host)	BLI	[437]
(-) -Butoxyl spiciferin A 676 ^β [C ₁₄ H ₂₀ O ₄]	UV, MS, NMR, [α] _D	Aromatic Polyketide	Undetm.	Undetm.	Undetm.	<i>D. hawaiiensis</i> (symbiont) <i>C. aerizusa</i> (host)	BLI	[437]
(+) -Epoxysorbicillinol 677 ^β [C ₁₄ H ₁₆ O ₅]	UV, IR, MS, NMR, [α] _D , ECD	Vertinoid Polyketide [•]	Undetm.	Undetm.	Undetm.	<i>T. longibrachiatums</i> (symbiont) <i>Halichlona</i> (host)	UEP	[438]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Cadiolide A 678 ^β [C ₂₄ H ₁₂ Br ₄ O ₆]	UV, IR, MS, NMR	γ-Lactone*	Cytotoxic	HCT116	NA	<i>Botryllus</i> sp.	SSW	[439]
Cadiolide B 679 ^β [C ₂₄ H ₁₀ Br ₆ O ₆]	UV, IR, MS, NMR	γ-Lactone	Cytotoxic	HCT116	NA	<i>Botryllus</i> sp.	SSW	[439]
(-)- <i>iso</i> -Cladospolide B 680 ^β [C ₁₂ H ₂₀ O ₄]	IR, MS, NMR, [α] _D , TS	γ-Lactone	Antibacterial	Gram-positive bacteria, Gram-negative bacteria	NA (250 µg/disk)	A fungus (symbiont) a sponge (host)	SSW	[397, 398, 440, 441]
(-)-Herbaric acid 681 ^β [C ₁₀ H ₈ O ₆]	MS, NMR, [α] _D , TS	γ-Lactone	Cytotoxic	HL-60	NA	<i>C. herbarum</i> (symbiont)	BLI	[440, 442]
				<i>A. salina</i>	NA	<i>A. aerophoba</i> (host)		
(+)-Biaketide 682 ^β [C ₁₇ H ₂₇ Cl ₂ O]	IR, MS, NMR, [α] _D	γ-Lactone*	Cytotoxic	NBT-T2	IC ₅₀ = 8.3 µg/mL	<i>Dysidea</i> sp.	PUA	[356]
(-)-Plakofuranaolactone 683 ^β [C ₈ H ₁₂ O ₄]	MS, NMR, [α] _D , QCC	γ-Lactone	Antibacterial (Quorum quenching agent)	<i>E. coli</i> pSB1075, <i>E. coli</i> pSB401, <i>C. violaceum</i> CV026), pyocyanin, total protease activity	0.781 – 200 µM	<i>Plakortis cf. lita</i>	NSW	[443]
Herbarin A 684 ^β [C ₁₂ H ₁₂ O ₅]	MS, NMR	δ-Lactone	Cytotoxic	<i>A. salina</i>	75% (50 µg) 85% (100 µg)	<i>C. herbarum</i> (symbiont) <i>C. aerizusa</i> (host)	BLI	[430]
Herbarin B 685 ^β [C ₁₀ H ₁₀ O ₅]	MS, NMR	δ-Lactone	Cytotoxic	<i>A. salina</i>	65% (50 µg) 80% (100 µg)	<i>C. herbarum</i> (symbiont) <i>C. aerizusa</i> (host)	BLI	[430]
686 ^β [C ₁₄ H ₂₀ O ₄]	UV, IR, MS, NMR	δ-Lactone	Growth restore activity	<i>S. cerevisiae</i> YNS17 (0.3 M CaCl ₂)	1.56-25 µg	<i>Fusarium</i> sp. (symbiont) <i>R. mucronata</i> (host)	JSCR	[444]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Helicascolide C 687 ^β [C ₁₂ H ₁₈ O ₃]	UV, IR, MS, NMR, CD, [α] _D , X-ray, TS	δ-Lactone	Antifungal	<i>C. cucumerinum</i>	78.5 mm ² (200 µg/disk)	<i>D. eschsholzii</i> (symbiont) <i>Gracilaria</i> sp. (host)	SSW	[445] – [447]
			Antibacterial	<i>C. maltosa</i> <i>E. coli</i> , <i>P. aeruginosa</i> <i>S. aureus</i> , <i>B. subtilis</i>	NA (200 µg/disk) NA (200 µg/disk) NA (300 µg/disk)			
			Cytotoxic	5637	NA (IC ₅₀ = 1.18 mM)			
			Cytotoxic	3Y1	10 µg/mL			
(+)–Bitungolide A 688 ^β [C ₂₅ H ₃₃ ClO ₅]	UV, IR, MS, NMR, [α] _D , X-ray	δ-Lactone	Antidiabetic	VHR	IC ₅₀ > 10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
			Cytotoxic	3Y1	10 µg/mL			
(+)–Bitungolide B 689a ^β [C ₂₅ H ₃₃ ClO ₅]	UV, IR, MS, NMR, [α] _D	δ-Lactone	Antidiabetic	VHR	IC ₅₀ > 10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
			Cytotoxic	3Y1	10 µg/mL			
(–)–Bitungolide B 689b ^γ [C ₂₅ H ₃₃ ClO ₅]	TS	δ-Lactone	Undetm.	Undetm.	Undetm.			[449]
(+)–Bitungolide C 690 ^β [C ₂₅ H ₃₃ ClO ₅]	UV, IR, MS, NMR, [α] _D	δ-Lactone	Cytotoxic	3Y1	10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
			Antidiabetic	VHR	IC ₅₀ > 10 µg/mL			
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
(+)–Bitungolide D 691 ^β [C ₂₅ H ₃₃ ClO ₅]	UV, IR, MS, NMR, [α] _D	δ-Lactone	Cytotoxic	3Y1	10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
			Antidiabetic	VHR	IC ₅₀ > 10 µg/mL			
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
(+)–Bitungolide E 692a ^β [C ₂₅ H ₃₄ O ₄]	UV, IR, MS, NMR, [α] _D	δ-Lactone	Cytotoxic	3Y1	10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
			Antidiabetic	VHR	IC ₅₀ > 10 µg/mL			
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
(–)–Bitungolide E 692b ^γ [C ₂₅ H ₃₄ O ₄]	TS	δ-Lactone	Undetm.	Undetm.	Undetm.			[449, 450]
(+)–Bitungolide F 693a ^β [C ₂₄ H ₃₂ O ₅]	UV, IR, MS, NMR, [α] _D	δ-Lactone	Cytotoxic	3Y1	10 µg/mL	<i>T. cf. swinhoei</i>	NSW	[448]
			Antidiabetic	VHR	IC ₅₀ > 10 µg/mL			
				PTP-S2, PP1 and PP2A	IC ₅₀ > 30 µg/mL			
(–)–Bitungolide F 693b ^γ [C ₂₅ H ₃₄ O ₄]	TS	δ-Lactone	Undetm.	Undetm.	Undetm.			[451]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)–Xestodecalactone A 694 ^β [C ₁₄ H ₁₆ O ₅]	UV, IR, MS, NMR, [α] _D , ECD, QCC, TS	Macrolactone	Antibacterial	<i>B. subtilis</i>	NA	<i>P. cf. montanense</i> (symbiont)	BLI	[452 – 454]
				<i>S. aureus</i>	NA	<i>X. exigua</i> (host)		
				<i>E. coli</i>	NA	<i>P. cf. montanense</i> (symbiont)		
(+)–Xestodecalactone B 695 ^β [C ₁₄ H ₁₆ O ₆]	UV, IR, MS, NMR, [α] _D , ECD, QCC, TS	Macrolactone	Antifungal	<i>C. albicans</i>	7 mm (20 μmol) 12 mm (50 μmol) 25 mm (100 μmol)	<i>X. exigua</i> (host)	BLI	[452, 455]
			Antibacterial	<i>B. subtilis</i> , <i>S. aureus</i> , <i>E. coli</i>	NA	<i>P. cf. montanense</i> (symbiont)		
(+)–Xestodecalactone C 696 ^β [C ₁₄ H ₁₆ O ₆]	UV, IR, MS, NMR, [α] _D , ECD, QCC, TS	Macrolactone	Antibacterial	<i>B. subtilis</i> , <i>S. aureus</i> , <i>E. coli</i>	NA	<i>X. exigua</i> (host)	BLI	[452, 456 – 458]
(–)-Pandangolide 1 697 ^β [C ₁₂ H ₂₀ O ₅]	UV, IR, MS, NMR, [α] _D	Macrolactone	Antibacterial	Gram-positive bacteria, Gram-negative bacteria	NA (250 μg)	A fungus (symbiont) a sponge (host)	SSW	[397, 459]
(–)-Pandangolide 2 698 ^β [C ₁₄ H ₂₂ O ₆ S]	UV, IR, MS, NMR, [α] _D	Macrolactone	Antibacterial	Gram-positive bacteria, Gram-negative bacteria	NA (250 μg)	A fungus (symbiont) a sponge (host)	SSW	[397]
(–)-Pandangolide 3 699 ^β [C ₁₆ H ₂₆ O ₇ S]	MS, NMR, [α] _D	Macrolactone	Antibacterial	Gram-positive bacteria, Gram-negative bacteria	NA	<i>C. herbarum</i> (symbiont)	BLI	[420]
(–)-Pandangolide 4 700 ^β [C ₂₄ H ₃₈ O ₈ S]	MS, NMR, [α] _D	Macrolactone [★]	Antibacterial	Gram-positive bacteria, Gram-negative bacteria	NA	<i>C. aerizusa</i> (host)	BLI	[420]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Callyspongiolide 701a ^β [C ₃₃ H ₄₄ BrNO ₆]	UV, IR, MS, NMR, [α] _D , CT	Macrolactone*	Cytotoxic	L5178Y Jurkat, Ramos B Induc. hypodiploid nuclei (Jurkat), (Ramos B)	IC ₅₀ = 320 nM IC ₅₀ = 60 – 70 nM (48 h) IC ₅₀ = 50 – 80 nM (48 h)	<i>Callyspongia</i> sp.	MLU	[460]
(-)-Callyspongiolide 701b ^γ [C ₃₃ H ₄₄ BrNO ₆]	TS	Macrolactone	Cytotoxic	MCF7, SH-SY5Y, HT-29, H1299, PC3 HeLa, RKO Jurkat	IC ₅₀ = 0.119 – 0.467 μM IC ₅₀ = 3.55 – 3.56 μM IC ₅₀ = 0.0111 μM			[460, 461, 462]
(+)-Phormidolide 702 ^β [C ₅₉ H ₉₇ BrO ₁₂]	UV, IR, MS, NMR, [α] _D , CT, Mol. Mod.	Macrolactone*	Cytotoxic	<i>A. salina</i> NCI <i>in vitro</i> 60-cell line	LD ₅₀ = 1.5 μM NA	<i>Phormidium</i> sp.	UEP	[463]
(+)- 703 ^{α,β} [C ₄₈ H ₈₀ O ₁₄]	IR, MS, NMR, [α] _D , CT	Macrolactone	Cytotoxic	NBT-T2	IC ₅₀ = 4.7 ng/mL	<i>Candida-spongia</i> sp.	ENT	[464]
(+)- 704 ^β [C ₃₂ H ₅₀ O ₁₃]	IR, MS, NMR, [α] _D , CT	Macrolactone	Cytotoxic	NBT-T2	IC ₅₀ = 19 ng/mL	<i>Candida-spongia</i> sp.	ENT	[464]
(-)-Laulimalide = (-)-Fijianolide B 705 ^β [C ₃₀ H ₄₂ O ₇]	UV, IR, MS, NMR, [α] _D , X-ray, CT, TS	Macrolactone*	Cytotoxic	P-388, KB, A549, MEL-28, HT HT-29	IC ₅₀ = 10-50 ng/mL IC ₅₀ = 6.9 nM (72 – 96 h) IC ₅₀ = 5.74 ± 0.58 nM (48 h)	<i>Hyattella</i> sp. <i>C. lochi</i>	NSW	[465 – 478]
				MDA-MB-435	IC ₅₀ = 2.3 ± 0.2 nM (72 – 96 h)			
				SK-OV-3	IC ₅₀ = 11.53 ± 0.53 nM (48 h)			
				MCF7	IC ₅₀ = 7 ng/mL, IC ₅₀ = 3.8 nM IC ₅₀ = 11.6 ± 0.5 nM (72 h)			
				MaTu	IC ₅₀ = 3.8 nM			
				HCT116, PC-3M	IC ₅₀ = 5.9 ± 0.3 – 7.8 ± 0.8 nM (72 h)			
				NCI/ADR	IC ₅₀ = 36 nM			
				MaTu/ADR	IC ₅₀ = 6.0 nM			
				SKVLB-1	IC ₅₀ = 1210 ± 490 nM (48 h)			
				Potency tubulin polymerization	EC ₅₀ = 4.0 ± 0.5 μM EC ₅₀ = 4.32 ± 0.4 μM			

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Isolaulimalide = (-)-Fijianolide A 706 ^β [C ₃₀ H ₄₂ O ₇]	UV, IR, MS, NMR, [α] _D , CT, TS	Macrolactone	Cytotoxic	MDA-MB-435 MCF7, PC-3M, HCT116	IC ₅₀ = 1970 ± 97 – 2650 ± 1384 nM (48 h) IC ₅₀ = 4400 ± 300 – 4900 ± 200 nM (72 h)	<i>Hyatella</i> sp. <i>C. lochi</i>	NSW	[465, 471, 476, 479]
(-)-Isoswinholide B 707 ^β [C ₇₈ H ₁₃₂ O ₂₀]	MS, NMR, [α] _D	Macrolactone	Cytostatic	HepG2	IC ₅₀ = 1500 nM	<i>T. swinhoei</i>	NSW	[480]
(-)-Swinholide K 708 ^β [C ₇₈ H ₁₃₂ O ₂₁]	MS, NMR, [α] _D	Macrolactone	Cytostatic	HepG2	IC ₅₀ = 15 nM	<i>T. swinhoei</i>	NSW	[480]
(-)-Xestosaprol L 709 ^β [C ₂₀ H ₁₈ O ₃]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 98 ± 8 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol I 710 ^β [C ₂₀ H ₁₈ O ₃]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 163 ± 11 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol J 711 ^β [C ₂₁ H ₂₀ O ₄]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 90 ± 5 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol K 712 ^β [C ₂₀ H ₁₈ O ₄]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 93 ± 4 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol G 713 ^β [C ₂₀ H ₁₈ O ₄]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 155 ± 15 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol H 714 ^β [C ₂₂ H ₂₂ O ₅]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 82 ± 3 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol F 715 ^β [C ₂₂ H ₂₂ O ₅]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 135 ± 11 μM	<i>Xestospongia</i> sp.	EKM	[481]
(-)-Xestosaprol D 716 ^β [C ₂₀ H ₁₈ O ₄]	UV, IR, MS, NMR, [α] _D	Quinone	Antibacterial Cytotoxic Anticancer Alzheimer	VRE, <i>E. coli</i> , MRSA, <i>S. aureus</i> SKOV-3 PKCδ BACE1	IC ₅₀ > 50 μg/mL IC ₅₀ > 50 μg/mL NA IC ₅₀ = 30 μg/mL	<i>Xestospongia</i> sp.	EKM	[482]
(-)-Xestosaprol E 717 ^β [C ₂₀ H ₁₈ O ₄]	UV, IR, MS, NMR, [α] _D	Quinone	Antibacterial Cytotoxic Anticancer	VRE, <i>E. coli</i> , MRSA, <i>S. aureus</i> SKOV-3 PKCδ	IC ₅₀ > 50 μg/mL IC ₅₀ > 50 μg/mL NA	<i>Xestospongia</i> sp.	EKM	[482]
(+)-Xestosaprol M 718 ^β [C ₂₀ H ₁₆ O ₂]	UV, IR, MS, NMR, [α] _D	Quinone	Alzheimer	BACE1	IC ₅₀ = 104 ± 8 μM	<i>Xestospongia</i> sp.	EKM	[481]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-14-Carboxy-xestoquinol sulfate 719 ^β [C ₂₁ H ₁₆ O ₉ S]	UV, IR, MS, NMR, [α] _D	Quinone	Anticancer	CDK1, CDK2, CDK5, CDK9, CK1, CLK1, DYRK1A, GSK3	IC ₅₀ > 10 μM	<i>Xestospongia</i> sp.	NSW	[483]
			Antibacterial	<i>S. aureus</i> ATCC 6538, <i>E. coli</i> ATCC 8739	NA			
			Antioxidant	DPPH	NA			
(-)-1-(2-Hydroxyethyl)-xestoquinone 720 ^β [C ₂₂ H ₁₈ O ₅]	UV, IR, MS, NMR, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 1.4 μM	<i>P. alfi</i>	NSW	[484]
1-(1-hydroxyethyl)-xestoquinone 721/722 ^{α,β} [C ₂₂ H ₁₈ O ₅]	MS, NMR	Quinone	Undetm.	Undetm.	Undetm.	<i>P. alfi</i>	NSW	[484]
(-)-3S-3-Hydroxy-xestoquinone 723 ^β [α] _D , ECD	UV, IR, MS, NMR, [α] _D , ECD	Quinone	Undetm.	Undetm.	Undetm.	<i>P. alfi</i>	NSW	[484]
(+)-Noelaquinone 724 ^β [C ₂₁ H ₁₅ N ₃ O ₅]	UV, IR, MS, NMR, [α] _D	Quinone*	Undetm.	Undetm.	Undetm.	<i>Xestospongia</i> sp.	EKM	[485]
(+)3-Ketoadociquinone B 725 ^β [C ₂₁ H ₁₅ N ₃ O ₅]	UV, IR, MS, NMR, [α] _D	Quinone	Anticancer	Recombinant human Cdc25B catalytic domain	IC ₅₀ = 0.13 ± 0.02 μM	<i>Xestospongia</i> sp.	NSW	[486]
				Recombinant human Cdc25B full length	IC ₅₀ = 0.21 ± 0.01 μM			
				Recombinant human VHR	IC ₅₀ = 9.0 ± 0.2 μM			
				Recombinant human PTP1B	IC ₅₀ = 3.9 ± 0.2 μM			
Xestoadociaquinones A 726/ B 727 ^{α,β} [C ₂₀ H ₁₉ NO ₈ S]	MS, NMR	Quinone	Undetm.	Undetm.	Undetm.	<i>Xestospongia</i> sp.	NSW	[483]
(-)-Xestoadociaminal A = Petroquinone I 728 ^β [C ₂₄ H ₂₁ NO ₇ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone*	Anticancer	USP7	IC ₅₀ > 5.0 μM	<i>Xestospongia</i> sp.	NSW	[483, 484]
Xestoadociaminal B = (+)-Petroquinone J 729 ^β [C ₂₄ H ₂₁ NO ₇ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ > 5.0 μM	<i>Xestospongia</i> sp.	NSW	[483, 484]
(+)-Petroquinone K 730 ^β [C ₂₄ H ₂₁ NO ₇ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ > 5.0 μM	<i>P. alfi</i>	NSW	[484]
(+)-Petroquinone L 731 ^β [C ₂₄ H ₂₁ NO ₇ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ > 5.0 μM	<i>P. alfi</i>	NSW	[484]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
Xestoadociaminals C 732 ^D 733 ^{α,β} [C ₂₄ H ₂₃ NO ₇ S]	MS, NMR	Quinone	Undetm.	Undetm.	Undetm.	<i>Xestospongia</i> sp.	NSW	[483]
(+)-Petroquinone A 734 ^β [C ₆₀ H ₃₆ O ₁₂]	UV, IR, MS, NMR, ECD, [α] _D	Quinone ^Δ	Anticancer	USP7	IC ₅₀ = 0.75 μM	<i>P. alfiani</i>	NSW	[484]
(-)-Petroquinone B 735 ^β [C ₆₀ H ₃₆ O ₁₂]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 0.36 μM	<i>P. alfiani</i>	NSW	[484]
(-)-Petroquinone C 736 ^β [C ₄₀ H ₃₀ O ₁₃ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone ^Δ	Anticancer	USP7	IC ₅₀ = 2.0 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Petroquinone D 737 ^β [C ₄₀ H ₃₂ O ₁₅ S ₂]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ > 5.0 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Petroquinone E 738 ^β [C ₄₀ H ₂₈ O ₁₁ S]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 1.2 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Petroquinone F 739 ^β [C ₄₀ H ₂₆ O ₈]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 0.35 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Petroquinone G 740 ^β [C ₄₀ H ₂₆ O ₈]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 0.47 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Petroquinone H 741 ^β [C ₄₂ H ₃₀ O ₈]	UV, IR, MS, NMR, ECD, [α] _D	Quinone	Anticancer	USP7	IC ₅₀ = 0.49 μM	<i>P. alfiani</i>	NSW	[484]
(+)-Biakamide A 742 ^β [C ₂₆ H ₄₂ ClN ₃ O ₃ S]	UV, IR, MS, NMR, [α] _D , ECD, CT, TS	Halogenated polyketide ^Δ	Cytostatic	PANC-1	IC ₅₀ = 1.0 μM (Glu.-Def. Med.) IC ₅₀ > 100 μM (Gen. Glu. Med.)	<i>Petrosaspongia</i> sp.	PUA	[487]
(+)-Biakamide B 743 ^β [C ₂₆ H ₄₂ ClN ₃ O ₃ S]	UV, IR, MS, NMR, [α] _D , CT, TS	Halogenated polyketide	Cytostatic	PANC-1	IC ₅₀ = 4.0 μM (Glu.-Def. Med.) IC ₅₀ > 100 μM (Gen. Glu. Med.)	<i>Petrosaspongia</i> sp	PUA	[487]
(-)-Biakamide C 744 ^β [C ₂₇ H ₄₂ ClN ₃ O ₃ S]	UV, IR, MS, NMR, [α] _D , CT, TS	Halogenated polyketide	Cytostatic	PANC-1	IC ₅₀ = 0.5 μM (Glu.-Def. Med.) IC ₅₀ = 50 μM (Gen. Glu. Med.)	<i>Petrosaspongia</i> sp	PUA	[487]

Table S21: Cont.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(-)-Biakamide D 745 ^β [C ₂₇ H ₄₂ ClN ₃ O ₃ S]	UV, IR, MS, NMR, [α] _D , CT, TS	Halogenated polyketide	Cytostatic	PANC-1	IC ₅₀ = 0.5 μM (Glu.-Def. Med.) IC ₅₀ = 35 μM (Gen. Glu. Med.)	<i>Petrosas- pongia</i> sp	PUA	[487]

Footnote: 1. **Activity** (3Y1 murine normal fibroblast, **H1299** human lung carcinoma, **MaTu** human breast adenocarcinoma, **MaTu/ADR** human multi-drug resistant breast adenocarcinoma, **MG63** human osteosarcoma, **NCI/ADR** human multi-drug resistant breast adenocarcinoma, **PC-3M** human prostate carcinoma, **RKO** human colorectal adenocarcinoma, **SH-SY5Y** human neuroblastoma, **SK-OV-3** human ovary adenocarcinoma, **CDK1** protein kinase, **CD45** protein tyrosine phosphatase, **PP1** protein serine/threonine phosphatase, **PKCδ** protein kinase C, **PP2A** protein serine/threonine phosphatase, **PTP-S2** protein tyrosine phosphatase, **TCPTP** protein tyrosine phosphatase, **VHR** protein tyrosine phosphatase, **D10 clone** *Plasmodium falciparum* chloroquine-sensitive, **VRE** Vancomycin-resistant enterococci); 2. **Geography** (PRC People's Republic China).

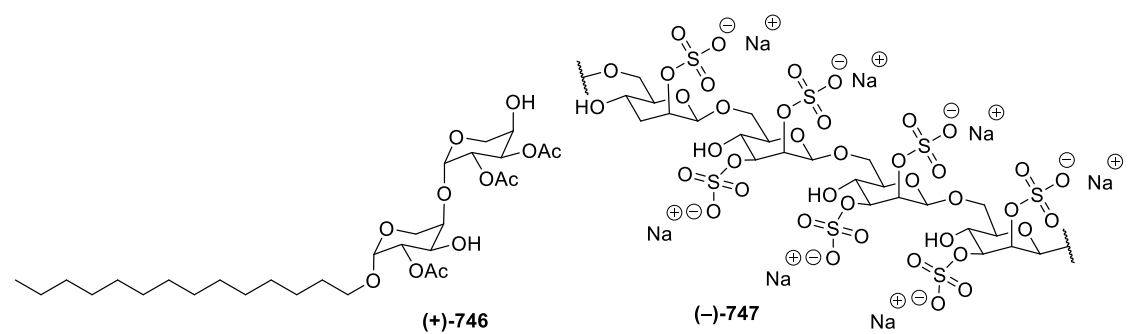


Figure S22: Structures of carbohydrates from Indonesian waters found in 1970–2017.

Table S22: Marine carbohydrates from Indonesian waters found in 1970–2017.

Compound	Structure Elucidation	Chemistry Type	Drug Class	Biological Activity		Source of Organism	Province	Ref
				Cell/Enzyme/Micro-organism/Insect/Others	Activity			
(+)-Sinularioside 746 ^β [C ₃₀ H ₅₂ O ₁₂]	MS, NMR, [α] _D , CT	Carbohydrate	Anti-inflammatory	JJI (LPS/ NO ₂ ⁻)	58% (30 μM)	<i>Sinularia</i> sp.	NSW	[88]
(-)-Kakelokelose 747 ^β	IR, MS, NMR, CT	Carbohydrate	Antiviral	HIV-1	100% (0.3 μg/mL)	<i>D. molle</i>	NSW	[488]

Footnote: 1. Activity (JJJ murine macrophage, LPS lipopolysaccharide).