



Screening of Fifteen Mangrove Plants Found in Sri Lanka for *in-vitro* Cytotoxic Properties on Breast (MCF-7) and Hepatocellular Carcinoma (HepG2) Cells

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Authors' contributions

This work was carried out in collaboration between all authors. Authors SRS, KHT and MKE were involved in designing the study. Authors SRS, CS, MKE and PP were involved in conducting the experiments, writing the manuscript and analyzing data. Manuscript was revised by authors SRS, KHT and IT. All authors read and approved the final manuscript.

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ABSTRACT

Aims: Evaluation of cytotoxic potential on the leaves and stem bark extracts of 15 mangrove plants grown in Sri Lanka on breast cancer (MCF -7) and hepatocellular carcinoma (HepG2) cells.
Place and Duration of Study: At the Institute of Biochemistry, Molecular Biology and Biotechnology, University of Colombo between 1st of February 2014 to April 2015.
Methodology: Leaves and stem barks of 15 mangrove plants were extracted with hexane, chloroform, ethyl acetate and methanol. Resulting extracts were screened for cytotoxic activity against MCF-7 and HepG2 cells using the Sulforhodamine B (SRB) assay.

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Results: *Phoenix paludosa*, *Avicennia officinalis* and *Scyphiphora hydrophyllacea* showed highest cytotoxic properties on cancer cells. Chloroform extract of stem bark of *S. hydrophyllacea*, *Bruguiera gymnorrhiza* (chloroform, ethyl acetate and methanol extracts of leaves), hexane and ethyl acetate extracts of leaves of *Aegiceras corniculatum*, methanol extracts of leaves and stem bark of *Nypa fruticans* and *Rhizophora mucronata*, methanol extract of stem bark of *Sonneratia alba* and *Rhizophora apiculata* and methanol extract of bark of *A. officinalis* exerted selective cytotoxicity to HepG2 cells. The hexane extract of leaves of *B. gymnorrhiza*, chloroform extract of leaves of *N. fruticans*, ethyl acetate extract of stem bark of *Lumnitzera littorea*, chloroform extract of leaves of *Rhizophora apiculata* and chloroform extract of leaves of *Pemphis acidula* showed selective cytotoxic effects against MCF-7 cells. Out of the 116 mangrove extracts tested, 82 extracts showed no significant cytotoxic effects ($IC_{50} > 100 \mu\text{g/mL}$) against MCF 7 or HepG2 cells.

Conclusion: The cytotoxic activities demonstrated by some of the solvent extracts of some mangrove plants provide scientific evidence for their therapeutic potentials and further studies are needed to identify active compounds responsible for cytotoxic effects.

Keywords: Mangrove; cytotoxicity; MCF-7; HepG2.

1. INTRODUCTION

Mangroves belong to twelve plant families and they are botanically diverse. Almost all the mangroves are holophytic species, well adapted to grow in wet soil conditions and usually possess some amount of viviparity [1,2]. The mangroves grown in Sri Lanka belong to true mangroves (14 species) and mangrove associates (12 species) [3]. In Sri Lanka mangroves are extensively found in Puttalam, Kalpitiya, Koggala, Kalamatiya and Kokilai areas in association with estuaries [4]. Mangroves have diverse uses: for example, they are used to obtain timber and tannins; they behave as coastal stabilizers; root system in mangroves provides shelter for many commercially important fishes and prawns, etc. [5]. These mangrove plants can survive in extremely high salinity, high temperature, high moisture, strong winds and high and low tides of water. In order to survive in these hostile environments, changes in their physiological activities have occurred ensuing in the bio-synthesis of novel secondary metabolites [6]. These secondary metabolites provide proper protection to these mangrove plants against various biotic and abiotic stress conditions [7]. A wide range of natural compounds, including novel chemical compounds have been isolated from mangroves and mangrove associates. Alkaloids, alcohols, amino acids, fatty acids, lipids, phenolic compounds, steroids, tannins, flavonoids, halogenated compounds, pheromones, phorbol esters and triterpenes are among these isolated compounds [8,9]. Some isolated compounds from mangroves are considered to have bioactivities that may be beneficial to improve human health and these compounds might be very useful in new drug

discovery process [10]. Mangrove plants have also been used as a folklore medicine and extracts from mangroves have been reported to have biological activities including cytotoxic, anti-bacterial, anti-viral and anti-inflammatory, etc as shown in Table 1. However, there is no data available on *in-vitro* cytotoxic properties of leaves and stem bark of most of the mangroves grown in Sri Lanka on breast (MCF-7) and hepatocellular carcinoma (HepG2) cells. Therefore, the main aim of this study was to evaluate the cytotoxic potential of leaves and stem bark of 15 selected mangroves/mangrove associates grown in Sri Lanka by evaluating of their effects on breast (MCF-7) and hepatocellular carcinoma (HepG2) cells.

2. MATERIALS AND METHODS

2.1 Chemicals

Fetal bovine serum (FBS), trypsin-EDTA, streptenicillin, HepG2 cells, MCF-7 cells and Dulbecco's modified Eagle's medium (DMEM) were purchased from American Type Culture Collection (ATCC), USA. All the chemicals used in the study were purchased from Sigma-Aldrich (St Louis, MO, USA) unless otherwise specified.

2.2 Plant Material

Healthy leaves and bark of 15 selected mangrove plants were collected from the mangrove park, Kadolkele, Negombo in the Western Province of Sri Lanka and Kalpitiya area in the North Western Province of Sri Lanka. Plants were identified by the Botanists at the National Herbarium, Royal Botanical Garden, Peradeniya, Sri Lanka and by Mr. W.A. Sumanadasa, of the National Aquatic Resources

Research and Development Agency (NARA), Negombo. Voucher specimens were deposited in the National Herbarium, Royal Botanical Garden, Peradeniya, Sri Lanka and Institute of Biochemistry Molecular Biology and Biochemistry, University of Colombo Sri Lanka (Table 2).

2.3 Preparation of Plant Extracts

Collected mangrove leaves and barks were dried at room temperature for 4-7 days and ground into

powder. Ground leaves and stem bark samples (10 g each) were extracted sequentially in to hexane, chloroform, ethyl acetate and methanol respectively by sonication at room temperature. All the resulting extracts (sixty leaf extracts and fifty six stem bark extracts) were filtered and concentrated under vacuum in a rotary evaporator (Rotavapor® R-/ BUCHI, Switzerland). Stock solutions were prepared by dissolving all extracts in dimethyl sulfoxide (DMSO).

Table 1. Botanical names, traditional uses and reported biological activities of studied mangroves grown in Sri Lanka

Botanical name	Traditional use of the plant	Reported biological activities	References
<i>Aegiceras corniculatum</i> ***	Used to treat for diabetes, asthma and in fish poisoning.	Toxic effects to fish, influence on the growth of fungi and some anti-viral activity	[11-17]
<i>Avicennia officinalis</i> ***	Used to treat for leprosy, hepatitis and as a diuretic.	Bio-toxic effects on fingerlings of fish and some anti-viral activity	[18-24]
<i>Bruguiera gymnorrhiza</i> ***	Used to treat for eye diseases.	Tested as growth hormones on plants	[25-28]
<i>Excoecaria Indica</i> ***	As a pain killer and in fish poisoning.	-	[29]
<i>Heritiera littoralis</i> ***	Used to treat for diarrhea and to control mosquitoes.	Antifungal, antifeedant activity and bio-toxic effects on some fish	[30-33]
<i>Lumnitzera littorea</i> **	Used to treat for Celiac disease.	-	[34]
<i>Lumnitzera racemosa</i> **	Used to treat for asthma, diabetes and infertility.	Antiviral activity	[35-38]
<i>Nypa fruticans</i> ***	Used to treat for diabetes, asthma, snake bites leprosy and rheumatoid arthritis.		[39,40]
<i>Pemphis acidula</i> **	Used to treat for reproduction related diseases.	Spasmolytic and oestrogenic activity	[41,42]
<i>Phoenix paludosa</i> *	Used to treat for diarrhea	Cytotoxic activity	[43,44]
<i>Rhizophora apiculata</i> ***	Used to treat for diarrhea, nausea, vomiting, typhoid, and hepatitis.	Antimicrobial activity, antiviral activity, antifungal, antifeedant and studies on HIV.	[45-48]
<i>Rhizophora mucronata</i> ***	Used to treat for elephantiasis, hepatitis, ulcers and hematoma.	Studies on HIV, as growth hormone on plants, bio-toxicity on fish.	[49-52]
<i>Scyphiphora hydrophyllacea</i> **	-	Cytotoxic properties of isolated compounds	[53,54]
<i>Sonneratia alba</i> ***	Used to treat swellings and sprains.		[55-57]
<i>Sonneratia caseolaris</i> ***	Used to treat for hemorrhages.	Toxic effects mosquito larvae	[58-60]

***mangroves; ** mangrove minors; * mangrove associates

2.4 Cell Culture Maintenance

HepG2 and MCF-7 cells were maintained in Dulbecco's modified Eagle's medium supplemented with 10% (v/v) fetal bovine serum and 50 IU/mL penicillin and 50 µg/mL streptomycin at 37°C in a humidified environment (95% air; 5% CO₂). At 80% confluency, cancer cells were trypsinized and seeded (5x10³ cells/well) in 96-well cell culture plates and incubated for 24 h.

2.5 Assessment of Cytotoxicity

After 24 h incubation, cells were exposed to leaves and stem bark extracts (doses ranging from 25 to 400 µg/mL and in triplicates) of mangroves and positive control paclitaxel for 24 and 48 h and cytotoxicity assessed by Sulforhodamine B assay (SRB) as previously described by us [61,62]. Briefly, treated cells were washed three times with PBS and fixed with Trichloroacetic acid (10%). Fixed cells were washed with tap water five times and then SRB (0.4%) was added to each well and incubated for 20 min. Unbound dye was removed by washing with acetic acid and bound dye was solubilized with Tris base (10 mM; pH 7.5). Plates were then kept on a plate shaker for 1 h and absorbance was taken at 540 nm using Synergy™ HT Multi-Mode Microplate Reader (BioTek, USA).

2.6 Statistical Analysis

All the experiments in this study were carried out at least three times in triplicate. Data were analyzed using Prism 5.0 (Graph pad Prism) statistical software package and results were expressed as mean ± standard deviation (SD).

3. RESULTS

A total of 116 solvent extracts (hexane, chloroform, ethyl acetate and methanol extracts of leaves and stem bark) representing 15 mangrove/mangrove associates and mangrove minors collected from Sri Lanka were tested for their cytotoxic effects on MCF 7 and HepG2 cells. The cytotoxic activities of mangrove extracts have been summarized in Table 2. Extracts with IC₅₀ value < 100 µg/mL were considered to be cytotoxic, while those with IC₅₀ value > 100 µg/mL were considered to be low/non-cytotoxic at 24 h or 48 h post incubation [63].

3.1 Selective Cytotoxic Effects of Mangrove Extracts

Among the 116 extracts tested in MCF-7 and HepG2 cells, chloroform extract of stem bark of *Scyphiphora hydrophyllacea*, *Bruguiera gymnorrhiza* (chloroform, ethyl acetate and methanol extracts of leaves), hexane and ethyl acetate extracts of leaves of *Aegiceras corniculatum*, methanol extracts of leaves and stem bark of *Nypa fruticans* and *Rhizophora mucronata*, methanol extracts of stem bark of *Sonneratia alba* and *Rhizophora apiculata* showed selective cytotoxicity to HepG2 cells. However, the hexane extract of leaves of *B. gymnorrhiza*, chloroform extracts of leaves of *N. fruticans*, ethyl acetate extract of stem bark of *Lumnitzera littorea*, chloroform extract of leaves of *R. apiculata* and chloroform extract of leaves of *Pemphis acidula* showed selective cytotoxic effects against MCF-7 breast cancer cells.

3.2 Non-selective Cytotoxic Effects of Mangrove Extracts

Hexane extracts of leaves and stem bark of *S. hydrophyllacea*, methanol extract of leaves of *P. paludosa* and ethyl acetate extract of stem bark of *A. officinalis* showed non-selective cytotoxic activity against both cancer cell lines tested.

3.3 Low or No Cytotoxic Effects of Mangrove Extracts

Out of the 116 mangrove extracts tested, 84 extracts showed no significant cytotoxic effects (IC₅₀>100 µg/mL) against MCF 7 and HepG2 cancer cells (Table 2).

4. DISCUSSION

Mangrove forests are considered to be the most productive ecosystems in the world [64]. However, mangroves grow under conditions such as high salinity, strong winds, extreme tides, high temperatures and extreme muddy soils. Thus, mangrove plants possess physiological, biological, ecological and morphological adaptations to extreme conditions [65]. Even though mangrove ecosystems have been studied broadly, there is a critical need to understand them better and care must be taken to prevent degradation and destruction of mangrove ecosystems.

Table 2. Cytotoxic effects of leaves and stem bark of mangrove plants and their voucher specimen numbers: IC₅₀ values of plant extracts on MCF 7 and HepG2 cells as determined by SRB assay at 24 and 48 h post incubation periods

Botanical name of the mangrove plant and voucher specimen number	Part used	Extract	IC ₅₀ value (µg/mL) at 24 h post incubation		IC ₅₀ value (µg/mL) at 48 h post incubation	
			MCF-7	HepG2	MCF-7	HepG2
<i>Aegiceras corniculatum</i> (C-7)	Leaves	Hexane	145.1	73.99	139.1	64.0
		Chloroform	109.6	98.72	93.8	80.2
		Ethyl acetate	166.9	82.91	133.2	71.6
		Methanol	283.1	426.2	200.5	408.1
	Stem/bark	Hexane	256.1	951.2	148.4	867.6
		Chloroform	189.0	260.2	44.69	159.7
		Ethyl acetate	466.7	148.1	302.8	100.7
<i>Avicennia officinalis</i> (C-11)	Leaves	Hexane	259.78	158.7	198.3	109.4
		Chloroform	418.89	184.7	336.1	132.2
		Ethyl acetate	267.39	241.1	227.9	180.5
		Methanol	198.9	221.2	153.2	182.3
	Stem/bark	Hexane	230.6	>1000	189.9	>1000
		Chloroform	81.38	155.4	67.2	140.0
		Ethyl acetate	65.08	54.8	56.3	27.91
<i>Bruguiera gymnorrhiza</i> (C-3)	Leaves	Hexane	98.43	764.9	90.2	730.3
		Chloroform	146.8	74.98	130.5	69.2
		Ethyl acetate	154.5	98.08	142.3	88.4
		Methanol	296.6	63.37	230.6	60.3
	Stem/bark	Hexane	>1000	>1000	576.5	152.9
		Chloroform	240.1	143.3	245.9	97.07
		Ethyl acetate	180.5	105.9	167.9	89.71
<i>Excoecaria Indica</i> (S-15)	Leaves	Hexane	>1000	>1000	>1000	>1000
		Chloroform	733.5	654.4	680.9	554.1
		Ethyl acetate	437.1	303.7	378.3	247.7
		Methanol	239.3	260.4	221.3	200.2
	Stem/bark	Hexane	499.9	>1000	419.9	>1000
		Chloroform	252.0	143.2	222.5	100.9
		Ethyl acetate	286.3	200.4	146.2	141.7
<i>Heritiera littoralis</i> (C-5)	Leaves	Hexane	939.9	>1000	883.7	>1000
		Chloroform	859.8	>1000	820.0	>1000
		Ethyl acetate	904.5	969.4	829.8	907.2
		Methanol	>1000	>1000	>1000	>1000
	Stem/bark	Hexane	>1000	>1000	>1000	>1000
		Chloroform	>1000	>1000	>1000	>1000
		Ethyl acetate	>1000	>1000	>1000	>1000
<i>Lumnitzera littorea</i> (S-2)	Leaves	Hexane	>1000	>1000	>1000	>1000
		Chloroform	323.0	377.7	298.2	358.0
		Ethyl acetate	286.3	288.9	247.2	260.1
		Methanol	200.3	264.3	148.6	234.1
	Stem/bark	Hexane	>1000	>1000	>1000	>1000
		Chloroform	395.2	402.5	355.9	337.8
		Ethyl acetate	120.2	250.2	88.4	251.0
Methanol	166.6	394.5	114.6	344.8		

Botanical name of the mangrove plant and voucher specimen number	Part used	Extract	IC ₅₀ value (µg/mL) at 24 h post incubation		IC ₅₀ value (µg/mL) at 48 h post incubation	
			MCF-7	HepG2	MCF-7	HepG2
<i>Lumnitzera racemosa</i> (C-4)	Leaves	Hexane	347.6	700.2	300.8	640.0
		Chloroform	170.1	230.1	123.3	169.4
		Ethyl acetate	541.6	440.7	499.0	394.0
		Methanol	765.87	777.98	752.1	701.9
	Stem/bark	Hexane	820.7	1020.5	582.9	66.46
		Chloroform	180.7	150.2	128.7	66.98
		Ethyl acetate	>1000	>1000	>1000	>1000
		Methanol	231.0	321.8	136.3	235.6
<i>Nypa fruticans</i> (S-3)	Leaves	Hexane	139.2	264.2	108.9	210.4
		Chloroform	90.36	277.7	77.2	240.3
		Ethyl acetate	166.9	203.2	154.8	164.2
		Methanol	390.98	74.27	120.4	64.2
	Stem/bark	Hexane	744.4	329.3	687.2	324.2
		Chloroform	460.5	566.8	419.0	417.2
		Ethyl acetate	>1000	>1000	>1000	>1000
		Methanol	160.2	103.6	140.3	71.52
<i>Pemphis acidula</i> (C-14)	Leaves	Hexane	154.5	135.9	122.3	109.4
		Chloroform	163.5	132.7	129.6	112.9
		Ethyl acetate	395.1	368.9	377.5	362.2
		Methanol	632.1	734.6	597.3	704.6
	Stem/bark	Hexane	189.7	138.4	154.6	111.8
		Chloroform	73.03	188.3	60.9	152.8
		Ethyl acetate	195.4	305.8	149.7	245.5
		Methanol	414.8	197.9	402.1	157.6
<i>Phoenix paludosa</i> (S-10)	Leaves	Hexane	369.2	889.3	319.9	840.1
		Chloroform	590.5	732.9	550.0	769.2
		Ethyl acetate	487.2	621.3	465.2	602.8
		Methanol	36.71	49.0	33.19	44.2
<i>Rhizophora apiculata</i> (C-12)	Leaves	Hexane	567.98	278.2	540.0	244.3
		Chloroform	678.89	598.4	629.3	509.7
		Ethyl acetate	765.87	1272	700.2	>1000
		Methanol	765.99	720.5	678.3	667.9
	Stem/bark	Hexane	>1000	303.6	>1000	235.5
		Chloroform	104.9	465.3	83.5	355.6
		Ethyl acetate	1004	460.5	980.4	420.2
		Methanol	245.9	133.6	204.5	96.91
<i>Rhizophora mucronata</i> (C-13)	Leaves	Hexane	908.78	801.2	820.0	720.6
		Chloroform	547.67	491.6	480.3	405.5
		Ethyl acetate	>1000	>1000	>1000	>1000
		Methanol	176.98	82.93	140.3	67.0
	Stem/bark	Hexane	716.6	821.8	640.5	745.1
		Chloroform	>1000	>1000	>1000	>1000
		Ethyl acetate	358.6	679.0	283.0	613.2
		Methanol	204.3	89.4	162.2	46.59
<i>Scyphiphora hydrophyllacea</i> (C-10)	Leaves	Hexane	87.97	83.74	66.3	62.16
		Chloroform	109.4	118	94.2	62.01
		Ethyl acetate	752.6	528.5	730.6	651.0
		Methanol	544.6	791.0	503.4	>1000
	Stem/bark	Hexane	90.3	84.74	80.9	56.7
		Chloroform	120.5	89.93	106.3	80.2
		Ethyl acetate	>1000	>1000	1031.0	>1000
		Methanol	900.4	861.4	474.2	803.5
<i>Sonneratia alba</i> (S-4)	Leaves	Hexane	764.92	678.9	673.9	598.2
		Chloroform	879.9	675.89	855.2	597.3
		Ethyl acetate	>1000	>1000	>1000	>1000
		Methanol	987.12	560.78	900.7	507.0

Botanical name of the mangrove plant and voucher specimen number	Part used	Extract	IC ₅₀ value (µg/mL) at 24 h post incubation		IC ₅₀ value (µg/mL) at 48 h post incubation	
			MCF-7	HepG2	MCF-7	HepG2
	Stem/ bark	Hexane	695.4	332.5	565.6	255.6
		Chloroform	152.6	159.0	112.4	100.2
		Ethyl acetate	326.2	>1000	300.3	>1000
		Methanol	201.4	144.7	172.3	94.92
<i>Sonneratia caseolaris</i> (C-16)	Leaves	Hexane	>1000	630.6	>1000	600.3
		Chloroform	632.6	420.3	583.5	301.3
		Ethyl acetate	365.8	399.2	366.4	320.5
		Methanol	600.6	487.7	580.4	434.4
	Stem/ bark	Hexane	>1000	>1000	>1000	>1000
		Chloroform	289.3	184.6	250.3	115.4
		Ethyl acetate	500.1	308.4	457.2	105.4
		Methanol	636.8	425.3	602.8	292.8
Paclitaxel		As positive control	4.7	6.3	3.8	2.8

Results of the present study with the leaves and stem bark extracts of fifteen mangrove species grown in Sri Lanka indicate that some of them have cytotoxic properties in breast (MCF-7) and hepatocellular carcinoma (HepG2) cells. Some mangrove plant extracts showed selective cytotoxic effects whereas some extracts showed non-selective cytotoxicity against both cancer cell lines or were not active (IC₅₀ >100 µg/mL) against any of the cell lines tested at 24 h or 48 h post incubations. Among the extracts tested, the methanol extract of *Phoenix paludosa* leaves showed the highest cytotoxicity in both cancer cell lines tested at 24 and 48 h post incubations. We have previously shown cytotoxic activity of different leaf extracts (hexane, chloroform, ethyl acetate and methanol) of *P. paludosa* in several cancer cell lines and normal cell lines [66]. Some known cytotoxic phytochemicals such as lupeol, epi-lupeol, and β-sitosterol have been isolated by other investigators from methanol extract of the leaves of *P. paludosa* [67-69]. In the present study, cytotoxic effects of stem bark extracts of *P. paludosa* were not tested due to difficulties in the collection of plant material. Ethyl acetate extract of *Avicennia officinalis* stem bark showed second highest cytotoxic properties in the two cancer cell lines tested and cytotoxic activity increased significantly at 48 h post incubation. Previous studies by other researchers have shown that, *A. officinalis* leaves extracts have cytotoxic effects with known phytochemicals such as triterpenes, betulinic acid, naphthoquinones, and avicequinone to Ehrlich ascites carcinoma (EAC) and human promyelocytic leukemia cell lines (HL 60) [70]. Hexane extract of *S. hydrophyllacea* leaves showed third highest and time dependent

cytotoxic properties in the two cancer cell lines tested and several cytotoxic compounds have been reported to be isolated from *S. hydrophyllacea* such as scyphiphin C, hopenone I, α-amyrin, β-amyrin, β-sitosterol and stigmasterol [53,54,71].

S. hydrophyllacea (hexane and chloroform extracts of stem bark), *B. gymnorrhiza* (chloroform, ethyl acetate and methanol extracts of leaves), *Aegiceras coniculatum* (hexane and ethyl acetate extracts of leaves), *N. fruticans* (methanol extract of leaves and stem bark), *S. alba* (methanol extract of stem bark), *A. officinalis* (methanol extract of bark), *R. apiculata* (methanol extract of stem bark) and *R. mucronata* (methanol extracts of leaves and stem bark) showed selective cytotoxic properties to HepG2 cells. Moreover, *B. gymnorrhiza* (hexane extract of leaves), *N. fruticans* (chloroform extract of leaves), *L. littorea* (ethyl acetate extract of stem bark), *R. apiculata* (chloroform extract of leaves) and *P. acidula* (chloroform extract of leaves) showed selective cytotoxic effects against MCF-7 breast cancer cells (IC₅₀ < 100 µg/mL). Among these plants, *A. coniculatum*, which was cytotoxic against HepG2 cells, has been used as a medicinal plant in Bangladesh for asthma, diabetes and rheumatism [72]. Extracts of this plant have reported to be cytotoxic to human gastric adenocarcinoma cells (AGS), colorectal adenocarcinoma cells (HT-29) and breast carcinoma cells (MDA-MB-435S) [72]. *A. officinalis* and *B. gymnorrhiza* which was cytotoxic to HepG2 cells in the present study have been used in traditional medicine to treat for leprosy, hepatitis, as a diuretic and for eye

disease respectively [73]. Extracts of these plants have also shown cytotoxic properties in cancer cells [73]. None of the extracts obtained from *Lumnitzera racemosa*, *Heritiera littoralis*, *Excoecaria indica* and *Sonneratia caseolaris* showed significant cytotoxic properties ($IC_{50} > 100 \mu\text{g/mL}$) on the two cancer cell lines tested.

This is the first study on screening of cytotoxic properties of leaves and bark of 15 listed mangrove plants grown in Sri Lanka against human breast and hepatocellular cancer cell lines. This study supports the reported cytotoxic activities of *S. hydrophyllacea*, *A. corniculatum*, *A. officinalis* and *B. gymnorrhiza*. Cytotoxic properties of *B. gymnorrhiza*, *P. paludosa*, *N. fruticans*, *S. alba*, *L. littorea*, *R. apiculata*, *R. mucronata* and *P. acidula* have not been reported previously. This study offers baseline data to focus on further studies into the isolation and characterization of novel secondary metabolites and to determine anti-cancer mechanism of such metabolites from mangrove plants grown in Sri Lanka. Mangrove plants that were found to be cytotoxic in the present study will be very useful as a source of new anti-cancer drug leads for drug discovery to fight against cancer.

5. CONCLUSION

Screening of leaves and stem barks of 15 selected mangrove plants growing in Sri Lanka, for cytotoxic activity in breast cancer cells (MCF-7) and hepatocellular carcinoma cells (HepG2) have demonstrated. Some mangrove plant extracts can exert selective cytotoxic properties to MCF-7 and HepG2 cells, whereas a few plant extracts showed non-selective cytotoxic properties, while a few others demonstrated no cytotoxic properties. The overall results indicate that some mangrove species found in Sri Lanka have the potential to be used in cancer therapy.

CONSENT

It is not applicable.

ETHICAL APPROVAL

It is not applicable.

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COMPETING INTERESTS

Authors have declared that no competing interests exist.

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