#### **REVIEW**

# Exploring gastropods in drug discovery: a review of their bioactive compounds and pharmacological applications

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Abstract Mollusc species are an abundant source of secondary metabolites with numerous biological activities. Approximately eighty percent of mollusks are gastropods, which are used in various ways, including as food and traditional medicine, due to their high protein content. The use of various gastropods has grown rapidly in the last decades, particularly in the pharmaceutical, nutraceutical, and cosmetic products industries. This review focuses on the various bioactive compounds derived from gastropods and their corresponding biological activities. A list of essential factors for discovering and developing novel therapeutic drugs has also been discussed, along with a roadmap for further research on marine-derived products. This review provides a detailed discussion of the various metabolites isolated from different gastropod species, including 40 bioactive compounds and their reported biological activities. However, we aim to analyze the evidence for the traditional use of gastropods in controlling various clinical problems. This study suggests that isolated bioactive compounds are valid leads for medical biotechnologists and pharmaceutical chemists to design novel drug molecules.

Keywords Bioactive compounds . Gastropods . Biological activities . Natural product . Pharmaceutical

#### Introduction

Gastropods are a significant class of mollusks found in both terrestrial and aquatic ecosystems, characterized by remarkable biological and chemical diversity. The marine environment is a huge source for discovering bioactive natural products. Traditional human populations have utilized numerous natural bioactive compounds from marine animal species to treat various ailments and enhance their healthcare. These bioactive compounds are regarded as potential candidates for affordable drug therapy worldwide. Mollusks are considered the second-most-diverse phylum of living creatures; they are characterized into eight classes, including Bivalvia, Scaphopoda, Cephalopoda, Polyplacophora, Monoplacophora, Caudofoveata, and Solenogastrea (Lydeard et al. 2004). They are known to be pharmacologically significant organisms of the Animal Kingdom. Among them, gastropod species are the most abundant class of mollusks, including various snails and slugs. Gastropod extracts are usually a complex combination of biochemically and pharmacologically active compounds, including proteins, peptides, lipids, steroids, alkaloids, terpenoids, and other organic compounds, which exhibit significant pharmacological properties, such as antimicrobial, anti-inflammatory, and anticancer activities. An in vitro antibacterial assay was conducted using the agar well diffusion method to analyze the hemolymph of *Buccinum undatum* and *Crepidula fornicata* against

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gram-positive and gram-negative bacterial strains. Furthermore, in vitro tests using Vero cells have demonstrated the antiviral effects of acidic extracts from gastropod species against Herpes simplex virus type 1 (HSV-1) (Blunt et al. 2009; Margret et al. 2013). Despite growing interest, many gastropod species remain underexplored in terms of their bioactive potential and therapeutic applications. This review aims to consolidate current findings on gastropod-derived bioactive compounds, highlighting their mechanisms of action and potential roles in modern drug development. Gastropod-derived natural metabolites have generally provided significant leads to the development of pharmaceutical medications that may exhibit more effective activity and benefits with fewer side effects. This study covers various scientific research on gastropods and their importance in pharmacology. The Scopus database, PubMed, ScienceDirect, Google Scholar, MarinLit, Web of Science, and other online scientific literature search engines were used to retrieve all published peer-reviewed scientific papers. Information was collected from published research articles, books, review articles, and regional-specific ethnomedical research about the significant use of gastropods' natural bioactive compounds. According to the World Register of Marine Species (WoRMS) database, the gastropod specimen nomenclature and family classification were described in this review article. The following information about gastropod-isolated compounds was obtained from chemical structure databases, such as PubChem and ChemSpider. In this study, Table 1 presents numerous species that contain various bioactive compounds and their corresponding bioactivities, including antimicrobial, anti-inflammatory, antioxidant, and anticancer properties, as determined through in vitro and in vivo experiments. This review is unique in that it offers the first comprehensive perspective on bioactive compounds derived from gastropods. It connects their pharmacological mechanisms and structural diversity with potential future directions in translational applications, high-throughput screening, metabolomics, and sustainable bioprospecting.

#### **Bioactive compounds**

Bioactive compounds are metabolites extracted from natural sources that can be protective against various diseases and metabolic disorders. These compounds provide significant leads for the development of pharmaceutical agents. Jiménez-Romero et al. (2014) have evaluated that a diterpene secondary metabolite, dactyloditerpenol acetate, was isolated from sea hare Aplysia dactylomel, which was screened for its in vitro anti-neuroinflammatory activity by inhibiting the production of thromboxane B2 (TXB2) and superoxide anion (O2(-)). This substance was discovered to have cytotoxic action against the prostate cancer cell lines DU-145 and A2058 melanoma cells. It was also shown to have antituberculosis activity against Mycobacterium tuberculosis H37Rv, with a minimum inhibitory concentration (MIC) of 59.4 μg/mL. Furthermore, Cadlina luteomarginata is a prevalent type of nudibranch, commonly found in the habitats of British Columbia. Secondary metabolites, such as sesterterpenoids and Ansellone A, were identified in the skin extract of C. luteromarginata, which stimulates the cAMP signaling pathway in HEK293 cells (Daoust et al. 2010). Muricidae family gastropods are an excellent source for producing synthetic brominated indoles and choline ester compounds. Bioactive compounds, including tyrindoxyl, 6-bromoindole, tyrindolinone, murexine, enecioylcholine, dihydromurexine, tyrindoxyl sulphate, tyriverdin, 6,6'-dibromoindigo, indirubin, 6-bromoindirubin, isatin, 5-bromoisatin, 6-bromoisatin, and 7-bromoisatin, were derived from both hypobranchial gland extracts and egg masses of Dicathais orbita species. These compounds have been shown to exhibit potent anti-inflammatory effects both in vitro and in vivo, inhibiting the production of tumor necrosis factor-alpha (TNF- $\alpha$ ), nitric oxide (NO), and prostaglandin E2 (PGE2). In addition, brominated isatin and 6-bromoisatin have also demonstrated both in vitro and in vivo antiproliferative activity against HT29 and Caco-2 cancer cells (Valles-Regino et al. 2016; Ahmad et al. 2017). Besides this, Chakraborty et al. (2019) investigated another secondary metabolite, ramosane, a sesquiterpenoid derivative obtained from the organic extract of the muricid gastropod Chicoreus ramosus. This isolated metabolite exhibits in vitro anti-inflammatory and antioxidant properties, making it beneficial for use in nutritional supplements and medicinal formulations. Dhahri et al. (2020) demonstrated that an active sulfated polysaccharide isolated from sea hare Bursatella leachii viscera exhibited in vitro anticoagulant activity using partial thromboplastin time and thrombin time.



#### Pharmaceutical applications of bioactive compounds

### Wound healing properties

In a study, Tsoutsos et al. (2009) reported Elicina® (Locafar, Chile), a cosmetic skin repair cream made from *Cornu (Helix) aspersum* mucus extracts. This cream offers effective treatment with skin-healing properties, working more quickly than other burn ointments used by adult patients to improve tissue regeneration and healing. Various components, including polyunsaturated fatty acids, amino acids, sterols, vitamin E, and aromatic compounds, were found in the flesh of the muricid *Rapana venosa* lipid extracts. These compounds induced skin burns in animal models by regenerating the skin's dermis and epidermis tissues, along with the formation of new blood vessels, epithelium, and collagen fibers. Such compounds are also considered as potential therapeutic anti-inflammatory agents (Badiu et al. 2008). In another study, El Mubarak et al. (2013) demonstrated that the mucus extract from *Helix aspersa* is utilized as a component of cosmetic skin treatments, which is a rich source of bioactive substances like glycolic acid and allantoin.

#### Anti-cancer properties

Considering the prevalence of cancer worldwide, anticancer substances are extremely important therapeutics. Marine gastropods have been identified as a novel source of bioactive compounds with promising anticancer potential. Numerous gastropod species' whole-body extracts have been tested for their activity on lymphocyte proliferation and antiproliferative assay on various human cancer cell lines. A drug named Brentuximab vedotin, isolated from the marine gastropod *Dollabella auricularia*, was FDA-approved and used as a medicine to treat Hodgkin's disease and lymphoma (Ciavatta et al. 2017). Nevertheless, achacin, a glycoprotein separated from the Achatina fulica snail's body mucus, has demonstrated anticancer efficacy against kidney epithelial and breast cancer cell lines in vitro (Ehara et al. 2002). Furthermore, bioactive cyclic and linear peptides were derived from the D. auricularia species, which exhibit anticancer properties against several cancer cell lines. Peptides such as dolastatins 10 and dolastatins 15, isolated from D. auricularia extracts. Both metabolites prevent the proliferation of liver and breast cancer cell lines. Dolastatin 10 is a unique structure with a pentapeptide subunit, while dolastatin 15 is a seven-subunit depsipeptide, and both peptides have been clinically tested for cancer therapy. Dolastatin 10 interferes with tubulin polymerization, leading to cell cycle arrest. LU-103793 (cematodin) and ILX651 (synthadotin) are derivatives of dolastatin 15; these two synthetic analogs have been formulated as anticancer medications for cancer therapy (Kang et al. 2018). Furthermore, Pla et al. (2006) reported that most lamellarin compounds were identified in the marine gastropod Lamellaria species, which have been reported to exhibit various pharmacological activities, including antimicrobial, antioxidant, and cytotoxic properties. For example, Lamellarin D, a bioactive alkaloid lead, exhibited in vitro cytotoxicity in human cancer cells, such as A-549 lung, MB-231 breast, and HT-29 colon cell lines. It was also shown to have potent inhibitory effects on human topoisomerase I. Some other cytotoxic bioactive metabolites, including jorunnamycin A, jorunnamycin B, jorunnamycin C, renieramycin M, renieramycin O, renieramycin Q, and mimosamycin, were isolated from the mantle, egg ribbons, and visceral organs of the nudibranch Jorunna funebris. These compounds were found to show cytotoxic activity in P388 lymphoma, A549 lung cancer, HT29 colon cancer, and MEL28 human skin cancer cell lines (Charupant et al. 2007). Similarly, Van et al. (2008) have reported diterpenes 6β,7α-diacetoxylabda-8,13E-dien-15-ol, 2α,6β,7α-triacetoxylabda-8,13E-dien-15-ol, 6β,7α,15-triacetoxylabda-8,13E-diene, 3α,11-dihydroxy-9,11-seco-cholest-4,7-dien-6,9-dione, Cholest-7-en-3,5,7-triol, which were isolated from *Timusculus costatus*. The findings reported that compounds such as 6β,7α-diacetoxylabda-8,13E-dien-15-ol (IC50 value: 25 μM), 2α,6β,7α-triacetoxylabda-8,13E-dien-15-ol (IC50 value: 24 μM), 6β,7α,15-triacetoxylabda-8,13E-diene (IC50 value: 84 μM), and 3α,11-dihydroxy-9,11-seco-cholest-4,7-dien-6,9-dione (IC50 value: 3 μM) have shown cytotoxicity against the human esophageal cancer cell line WHCO1. Besides this, the extracts of this nudibranch specimen yielded compound epoxygoniolide-1 and compounds of the gracilin family, including aplytandiene-3, gracilin A, gracilin B, gracilin C, gracilin G, and gracilin M. All of these compounds were screened against human colorectal cancer and lung and liver cancer cell lines (Hirayama et al. 2016; Forster et al. 2017).



#### Anti-inflammatory properties

Gastropods' whole-body, flesh, and shell extracts contain bioactive compounds that are utilized as anti-inflammatory agents to treat osteoarthritis, rheumatoid arthritis, fever, lethargy, and joint discomfort. Marine gastropod Cypraea arabica methanolic extracts have shown potent analgesic, antipyretic, and anti-inflammatory activity in Wistar albino rats (Subavathy and Thilaga 2018). The venom of cone snails provides pain-relieving toxins. For example, Ziconotide (Prialt) is the first painkiller derived from the conotoxin in the venom of the marine carnivorous cone snail *Conus magus*. The Food and Drug Administration (FDA) has approved it after clinical testing, and it is used to treat neurological diseases and chronic pain. Ziconitide is a drug designed from naturally occurring conopeptides (Joseph et al. 2011). According to Joshi et al. (2016), the hexapeptide C-II is derived from *Harpa ventricosa* visceral mass extracts and acts as an anti-inflammatory peptide. This protein fraction exhibited an anti-inflammatory assay that inhibits the synthesis of TNF- $\alpha$  and IL-1 $\beta$ . Furthermore, the purified peptide also showed cytotoxicity against the THP-1 leukemia cell line. Thus, the peptide is utilized in medicinal applications for the treatment of inflammatory and cancer diseases. Moreover, Bhattacharya et al. (2014) reported that lipid extracts were made from the footpads of the edible snail Bellamya bengalensis, which is used by tribal people as a potential medicine to prevent rheumatism-like joint and bone inflammation, inhibit macrophage activation, and reduce cellular hypersensitivity in a rat model. Similarly, Oliveira et al. (2015) have investigated the digestive gland extracts of Aplysia depilans Gmelin, which consist of eight carotenoids and twenty-two long-chain polyunsaturated fatty acids. These compounds have been shown to have significant anti-inflammatory effects and reduce nitric oxide (NO) production.

#### Antimicrobial properties

An antimitotic and antiviral agent, kelletinin A (KA), is a naturally occurring substance extracted from the marine gastropod Buccinulum corneum. In addition to its antimitotic action on HTLV-1-infected MT2 cells, KA demonstrated antiviral activity against the human T-cell leukemia virus type 1 (HTLV-1) and suppressed the production of cellular DNA and RNA (Silvestri et al. 1995). Benkendorff et al. (2001) extracted brominated indole metabolites, such as tyriverdin, 6-bromoisatin, and tyrindolenin, from the egg mass of the Dicathais orbita species. Additionally, another compound, 6-bromo-2-methylmercaptoindoxyl-3-sulfate, was obtained from the hypobranchial glands of this species. These compounds have exhibited antimicrobial properties against various pathogens, including Staphylococcus aureus, Escherichia coli, Pseudomonas aeruginosa, Candida albicans, Enterococcus sericolicida, and Vibrio angillarium, as determined by a disk diffusion assay. Similarly, Zhong et al. (2013) reported that mytimacin-AF peptide was identified from the mucus extract of the giant African snail Achatina fulica. These compounds have shown promising antimicrobial activity against gram-positive pathogens, including Staphylococcus aureus, Bacillus megatherium, and gram-negative pathogens such as Escherichia coli, Bacillus pyocyaneus, Bacillus dysenteriae, and Klebsiella pneumoniae, as well as the fungal strain C. albicans, by using disk diffusion methods. The results showed that S. aureus was the most sensitive to mytimacin-AF among all the strains, with a minimum inhibitory concentration (MIC) of 1.9 µg/ml. Furthermore, other antimicrobial compounds, such as pseudoephedrine, hexanal-2-methyl, 2,2-dimethylpropionic acid, hexadecyl ester, and 1,2-benzenedicarboxylic acid diisooctyl ester, were isolated from *Phalium glaucum* whole-body tissue extracts. The extracted bioactive compounds were tested for antibacterial activity using the disc diffusion method against the pathogens Escherichia coli, Klebsiella pneumoniae, Proteus vulgaris, Salmonella typhi, Shigella flexneri, and Staphylococcus aureus. The results showed that the highest activity was exhibited against the S. typhi strain, with an MIC of 12 mm, and the lowest activity was observed against the E. coli strain, with an MIC of 5 mm (Thilaga et al. 2014).

As shown in Table 1, gastropod-derived metabolites exhibit a broad spectrum of biological activities, with the majority demonstrating cytotoxic and anticancer potential (e.g., aplyronines, elisidepsin, zalypsis, dendocarbins). Several compounds also display antimicrobial and antifungal effects (e.g., deoxymanoalide, scutinins, aglajne-1), while bromotriterpenes such as aplysqualenols broaden this scope to include antiviral and antiplasmodial activity. Notably, certain alkaloids and terpenes exhibit anti-inflammatory, antifouling, and antimalarial properties, underscoring the structural and functional diversity



 Table 1 Pharmacological activity of bioactive compounds reported from gastropod species.

SL. No.	Compound Name (Gastropod Species); Chemical Class	Molecular Formula; Chemical Database ID	Chemical Properties	Biological Effect	Reference
1	Aurilol	MF:C <sub>30</sub> H <sub>53</sub> BrO <sub>7</sub>	MW:605.6[g/mol]	Cytotoxic	Suenaga et al. (1998)
	(Dolabella auricularia);	PubChem ID:11365474	XLogP3:3.6		
	Bromotriterpene		H-bd:3; H-ba:7		
2	Laurinterol	$MF: C_{15}H_{19}BrO$	MW: 295.21[g/mol]	Cytotoxi	Tsukamoto et al. (2005)
	(Aplysia kurodai);	PubChem ID: 11471955	XLogP3:5.3	Antibacterial	
	Sesquiterpene		H-bd:1; H-ba:1		
3	Debromolaurinterol	MF: C <sub>15</sub> H <sub>20</sub> O	MW: 216.32[g/mol]	Cytotoxic	Tsukamoto et al. (2005)
	(Aplysia kurodai);	PubChem ID: 466442	XLogP3:4.6	Antibacterial	
	Sesquiterpene		H-bd:1; H-ba:1		
4	Aplysqualenol A	MF: C <sub>31</sub> H <sub>53</sub> BrO <sub>7</sub>	MW: 617.7 [g/mol]	Antivira	Vera et al. (2009)
	(Aplysia dactylomela);	PubChem ID: 44551049	XLogP3:4.5	Antitumor	
	Bromotriterpene		H-bd:2; H-ba:7		
	Aglajne-1	MF: C <sub>25</sub> H <sub>40</sub> O <sub>3</sub>	MW: 388.6 [g/mol]	Antibacterial	Arrieche et al. (2019)
5	(Bulla occidentalis);	PubChem ID: 13917442	XLogP3:6.8		( /
	Polypropionate		H-bd:0; H-ba:3		
6	Deoxymanoalide (Chromodoris willani);	MF: C <sub>25</sub> H <sub>36</sub> O <sub>4</sub>	Avg. mass: 400.551 [Da]	Antimicrobial	Uddin et al. (2009)
	Sesterterpene	ChemSpider ID:27024351	XLogP3:6.07		()
	Sesterterpene	Chemophael 13.2702 1331	H-bd:1; H-ba:4		
	Fusaripyrone A	MF: C <sub>32</sub> H <sub>48</sub> O <sub>3</sub>	MW: 480.7[g/mol]	Antifouling	Cutignano et al. (2007)
	(Haminoea fusari);	PubChem ID: 101446903	XLogP3:10.7	Antilouning	Cutignano et al. (2007)
		rubellelli ID. 101440303			
	Pyrone	ME C. H. O.	H-bd:1; H-ba:3	A ('C 1'	G (' (2007)
	Fusaripyrone B	MF: C <sub>32</sub> H <sub>48</sub> O <sub>4</sub>	MW: 496.7[g/mol]	Antifouling	Cutignano et al. (2007)
	(Haminoea fusari);	PubChem ID: 101446904	XLogP3:9.3		
9	Pyrone		H-bd:2; H-ba:4		
	Malyngamide S	MF: C <sub>26</sub> H <sub>42</sub> ClNO <sub>5</sub>	MW: 484.1 [g/mol]	Anti-inflammatory	Appleton et al. (2002)
	(Bursatella leachii);	PubChem ID: 643654	XLogP3:4.6	Antimicrobial	
	Alkaloid		H-bd:2; H-ba:5	Cytotoxicity	
10	Scutinin A	MF: $C_{48}H_{38}O_{18}$	MW: 902.8 [g/mol]	Antimicrobial	Chand et al. (2017)
	(Scutus antipodes);	PubChem ID: 132850622	XLogP3:7.7H-bd:6; H-ba:18		
	Epimer				
11	Glandulaurencianol A	$MF: C_{20}H_{33}BrO_2$	Avg. mass:385.379 [Da]	Antibacterial	Kladi et al. (2014)
	(Aplysia punctata);	ChemSpider ID:32675113	XLogP3:6.10		
	Diterpene		H-bd:1; H-ba:2		
12	Glandulaurencianol C	MF: $C_{20}H_{32}O_2$	Avg. mass:304.467 [Da]	Antibacterial	Kladi et al. (2014)
	(Aplysia punctata);	ChemSpider ID:32675115	XLogP3:5.77		
	Diterpene	_	H-bd:1; H-ba:2		
13	Diemenensin A	MF: C <sub>21</sub> H <sub>32</sub> O <sub>3</sub>	MW: 332.5 [g/mol]	Antimicrobial	Hochlowski and
	(Siphonaria diemenensis);	PubChem ID: 54715959	XLogP3:6.5		Faulkner (1983)
	Polypropionate		H-bd:1; H-ba:3		
14	Actinofide	MF: C <sub>21</sub> H <sub>35</sub> N <sub>3</sub> O	MW: 345.5 [g/mol]	Anticancer	Carbone et al. (2017)
	(Actinocyclus papillatus);	PubChem ID: 132606933	XLogP3:6.2		( ,
	Terpenoid		H-bd:2; H-ba:2		
15	Kulokekahilide-2	MF: C <sub>44</sub> H <sub>67</sub> N <sub>5</sub> O <sub>10</sub>	MW: 826.0 [g/mol]	Anticancer	Nakao et al. (2004)
	(Philinopsis speciosa);	PubChem ID: 11491350	XLogP3:6	7 11110411001	1 tantao et an. (2001)
	Cyclic depsipeptide	1 doction 15. 11451550	H-bd:4; H-ba:10		
6	Keenamide-A	MF: C <sub>30</sub> H <sub>48</sub> N <sub>6</sub> O <sub>6</sub> S	Avg. mass: 620.804 [Da]	Anticancer	Wesson et al. (1996)
,	(Pleurobranchus forskalii);	ChemSpider ID:10200275	XLogP3: -0.95	Anticancei	wesson et al. (1990)
	Cyclic hexapeptide	Chemspider iD:10200275	H-bd:4; H-ba:12		
7	Zalypsis	MF: C <sub>37</sub> H <sub>38</sub> F <sub>3</sub> N <sub>3</sub> O <sub>8</sub>	MW: 709.7 [g/mol]	Antitumor	Malve (2016)
/				Anutumoi	Maive (2010)
	(Joruna funebris);	PubChem ID: 16061448	XLogP3:5		
18	Alkaloid	ACC H N C	H-bd:3; H-ba:13		11 (1000)
5	Elisidepsin	MF: C <sub>75</sub> H <sub>124</sub> N <sub>14</sub> O <sub>16</sub>	MW: 1477.9 [g/mol]	Anticancer	Hamann et al. (1996)
	(Elysia rufescens);	PubChem ID: 9855343	XLogP3:7.4		
	Cyclic depsipeptide		H-bd:14; H-ba:17		
9	Tyrindoleninone (Dicathais orbita);	MF: C <sub>9</sub> H <sub>6</sub> BrNOS	MW: 256.12 [g/mol]	Anticancer	Edwards et al. (2012)
	Brominated indole	PubChem ID: 618037	XLogP3:2.6		
			H-bd:0; H-ba:3		
0	Aplyronine A	$MF: C_{59}H_{101}N_3O_{14}$	MW: 1076.4 [g/mol]	Cytotoxic	Yamada et al. (1993)
	(Aplysia kurodai);	PubChem ID: 11840920	XLogP3:9.1		
	Alkaloid		H-bd:2; H-ba:16		
21	Aplyronine H	MF: $C_{58}H_{99}N_3O_{14}$	MW: 1062.4 [g/mol]	Cytotoxic	Yamada et al. (2000)
	(Aplysia kurodai);	PubChem ID: 102037441	XLogP3:8.7		
	Alkaloid		H-bd:3; H-ba:16		
22	Neoaplaminone	MF: C <sub>26</sub> H <sub>40</sub> BrNO <sub>4</sub>	Avg. mass:510.504 [Da]	Cytotoxic	Kigoshi et al. (1990)
	(Aplysia kurodai);	ChemSpider ID8431248	XLogP3:4.99		
	Alkaloid	•	H-bd:2; H-ba:5		
3	Phidianidine A	MF: C <sub>17</sub> H <sub>22</sub> BrN <sub>7</sub> O	MW: 420.3 [g/mol]	Cytotoxic	Carbone et al. (2011)
-	(Phidiana militaris);	PubChem ID: 59053149	XLogP3:2.8	•	` ′
	Indole alkaloid		H-bd:4; H-ba:5		
24	Phidianidine B	MF: C <sub>17</sub> H <sub>23</sub> N <sub>7</sub> O	MW: 341.4 [g/mol]	Cytotoxic	Carbone et al. (2011)
				Cytotoxic	Carbone et al. (2011)
	(Phidiana militaris);	PubChem ID: 60150771	XLogP3:2.1		
25	Indole alkaloid	ME, C. H. N.O.	H-bd:4; H-ba:5	Cutatorii	Walring 4 1 (2012)
,	Ergosinine	MF: C <sub>30</sub> H <sub>37</sub> N <sub>5</sub> O <sub>5</sub>	MW: 547.6 [g/mol]	Cytotoxic	Wakimoto et al. (2013)
	(Pleurobranchus forskalii);	PubChem ID: 10030389	XLogP3:1.8		
	Ergot alkaloid		H-bd:3; H-ba:6		
26	Cycloforskamide	$MF: C_{54}H_{86}N_{12}O_{11}S_3$	MW: 1175.5 [g/mol]	Cytotoxic	Tan et al. (2013)
	(Pleurobranchus forskalii);	PubChem ID: 71747326	XLogP3:4.8		
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27	Reticulidin A	MF: C <sub>16</sub> H <sub>22</sub> C <sub>13</sub> NO	MW: 350.7 [g/mol]	Cytotoxic	Tanaka and Higa (1999)
	(Reticulidia fungia);	PubChem ID: 10337976	XLogP3:5.6		
	Sesquiterpene		H-bd:1; H-ba:2		
28	Jorumycin	MF: $C_{27}H_{30}N_2O_9$	MW: 526.5 [g/mol]	Cytotoxicity	Fontana et al. (2000)
	(Jorunna funebris);	PubChem ID: 9849761	XLogP3:0.1		
	Alkaloid		H-bd:1; H-ba:11		
29	Tambjamine K	MF: C <sub>15</sub> H <sub>21</sub> N <sub>3</sub> O	MW: 259.35 [g/mol]	Cytotoxic	Carbone et al. (2010)
	(Tambja ceutae);	PubChem ID: 135934866	XLogP3:2.8		
	Alkaloid		H-bd:2; H-ba:2		
30	Dendocarbin A	MF: C <sub>15</sub> H <sub>22</sub> O <sub>3</sub>	MW: 250.33 [g/mol]	Cytotoxicity	Sakio et al. (2001)
	(Dendrodoris carbunculosa);	PubChem ID: 10911949	XLogP3:3.5		
	Sesquiterpene		H-bd:1; H-ba:3		
31	Dendocarbin N	MF: C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	MW: 268.35 [g/mol]	Cytotoxicity	Sakio et al. (2001)
	(Dendrodoris carbunculosa);	PubChem ID: 21592441	XLogP3:1.8		
	Sesquiterpene		H-bd:2; H-ba:4		
	Isodrimeninol	MF: $C_{15}H_{24}O_2$	MW: 236.35 [g/mol]	Cytotoxicity	Sakio et al. (2001)
32	(Dendrodoris carbunculosa);	PubChem ID: 11322321	XLogP3:3.1		
	Sesquiterpene		H-bd:1; H-ba:2		
33	11-epivaldiviolide	MF: C <sub>15</sub> H <sub>22</sub> O <sub>3</sub>	MW: 250.33 [g/mol]	Cytotoxicity	Sakio et al. (2001)
	(Dendrodoris carbunculosa);	PubChem ID: 10354740	XLogP3:3.2		
	Sesquiterpene		H-bd:1; H-ba:3		
	Furodysinin	MF: C <sub>15</sub> H <sub>20</sub> O	MW: 216.32 [g/mol]	Cytotoxicity	Mudianta et al. (2016)
34	(Hypselodoris infucata);	PubChem ID: 155517	XLogP3:3.9		
	Sesquiterpene		H-bd:0; H-ba:1		
35	Lovenone	MF: C <sub>29</sub> H <sub>48</sub> O <sub>4</sub>	Avg. mass: 460.689 [Da]	Cytotoxicity	Graziani et al. (1995)
	(Adalaria loveni);	ChemSpider ID:8225484	XLogP3:5.60		
	Triterpenoid		H-bd:2; H-ba:4		
36	Aplykurodin A	MF: $C_{20}H_{34}O_3$	MW: 322.5 [g/mol]	Cytotoxicity	Lee et al. (2020)
	(Aplysia kurodai);	PubChem ID: 21674181	XLogP3:5.1		
	Sesquiterpene		H-bd:1; H-ba:3		
	Aplysiasecosterol A	MF: C <sub>27</sub> H <sub>44</sub> O <sub>7</sub>	MW: 480.6 [g/mol]	Cytotoxicity	Kawamura et al. (2015)
37	(Aplysia kurodai);	PubChem ID: 102367614	XLogP3:1.9		
	Steroid		H-bd:5; H-ba:7		
38	Tritoniopsin A	MF: C <sub>24</sub> H <sub>38</sub> O <sub>5</sub>	MW: 406.6 [g/mol]	Cytotoxicity	Ciavatta et al. (2011)
	(Tritoniopsis elegans);	PubChem ID: 101960706	XLogP3:3.3		
	Diterpenoid		H-bd:1; H-ba:5		
39	Tritoniopsin C	MF: C <sub>24</sub> H <sub>38</sub> O <sub>6</sub>	MW: 422.6 [g/mol]	Cytotoxicity	Ciavatta et al. (2011)
	(Tritoniopsis elegans);	PubChem ID: 54671808	XLogP3:2.3		
	Diterpenoid		H-bd:2; H-ba:6		
40	Tritoniopsin D	MF: C <sub>26</sub> H <sub>40</sub> O <sub>7</sub>	MW: 464.6 [g/mol]	Cytotoxicity	Ciavatta et al. (2011)
	(Tritoniopsis elegans);	PubChem ID: 54671895	XLogP3:2.8		
	Diterpenoid		H-bd:1; H-ba:7		

MF: Molecular formula; MW: Molecular weight; Avg. mass: Average mass; H-bd: Hydrogen bond donor; H-ba: Hydrogen bond acceptor

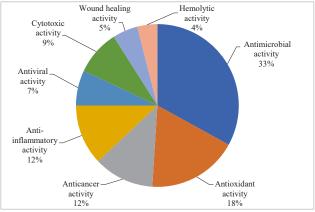


Fig. 1 Distribution of bioactivities studied from various species of gastropod

of these metabolites. collectively, these findings highlight gastropods as a rich reservoir of bioactive compounds with strong promise for drug discovery and therapeutic development.

## Limitations and challenges

Despite the impressive pharmacological potential of secondary metabolites produced from gastropods, their translational potential is limited by several significant constraints. Initially, many of these metabo-



lites are found in extremely low natural yields, making it challenging to isolate them on a large scale and frequently impossible to replicate due to seasonal or ecological fluctuations. A reliable supply is further challenged by the reliance on wild harvesting, which raises concerns about sustainability and ethics. Furthermore, the structural complexity of many compounds derived from gastropods, particularly peptides and alkaloids, poses significant challenges to their scalable manufacture, as it limits chemical modification and synthetic replication. Environmental effects, such as the overuse of marine resources, underscore the need for synthetic biology techniques, aquaculture systems, and non-lethal sampling methods. Lastly, regulatory obstacles that hinder the transition from discovery to therapeutic application include the lack of thorough toxicological data, inconsistent pharmacokinetic investigations, and stringent clinical approval requirements. Addressing these barriers will be crucial for advancing gastropod-derived compounds from the bench to the bedside.

#### **Future perspectives**

This study provides information on the bioactive characteristics of gastropod species and their extracted chemicals, which are essential for developing new, powerful medications to treat various infectious disorders and inflammations. In the future, drugs such as antibiotics and anticancer agents may be discovered from compounds derived from gastropods. Several studies have reported the presence and potential importance of bioactive compounds, while the bioactive compounds of some species and their biological applications have yet to be explored. Therefore, further studies are required to isolate and structurally elucidate the bioactive compounds of gastropods. Detailed structural characterization facilitates rational drug design and enables structure-activity relationship (SAR) studies, which are critical for optimizing potency, selectivity, and safety. Researchers can employ advanced techniques, such as proteomics and transcriptomics, to identify rare and novel genes and proteins. However, recombinant techniques can also be used to express the proteins and peptides in gastropod species that may serve as a potential source for synthesizing novel drug formulations. There are limited silico investigations and human clinical trials on gastropod-derived bioactive compounds. This highlights the need for computational modeling, pharmacokinetic studies, and well-designed clinical evaluations to validate their therapeutic potential. Future investigations should also focus on in silico approaches using advanced bioinformatics tools and quantitative structure-activity relationship (QSAR) methods, which can enhance the identification, characterization, and function of unknown bioactive proteins or peptides within a shorter timeframe. This investigation will enable researchers to explore the critical aspects of the medicinal properties of gastropods and provide new insights into their pharmacological properties. Furthermore, to ensure long-term resource availability, future studies on compounds derived from gastropods should focus on advanced metabolomic profiling to identify new metabolites, combining this with high-throughput screening to link metabolites to their corresponding bioactivities. Employing sustainable bioprospecting techniques will also be crucial in translating promising leads into applications in pharmaceuticals, cosmeceuticals, and nutraceuticals.

#### Conclusion

This review highlights the increasing importance of gastropods as a promising yet underexplored source of bioactive compounds with diverse pharmacological applications. Particularly, we reviewed key compounds, including peptides, alkaloids, sterols, and polysaccharides, many of which exhibit potent mechanisms of action, such as antimicrobial, anticancer, anti-inflammatory, and cytotoxic effects. The evidence underscores their therapeutic value in addressing pressing health challenges ranging from infectious diseases to cancer and metabolic disorders. Future efforts integrating advanced metabolomics, high-throughput bioactivity screening, structural elucidation, sustainable bioprospecting, and translational studies will be essential to unlock their full potential. In conclusion, gastropods represent an efficient source of natural products that could be of tremendous therapeutic value in the new millennium. Therefore, by bridging traditional natural product research with modern technologies, gastropods could emerge as a vital resource in the discovery and development of next-generation therapeutics.



Acknowledgments We express our gratitude to the academic facilities and the Honorable President, Siksha 'O' Anusandhan (Deemed to be University), Odisha, Bhubaneswar. We are particularly grateful for the extended research facilities provided by the Dean of IMS and SUM Hospital in Odisha, Bhubaneswar.

Conflict of interest The authors have no conflicts of interest associated with this publication.

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