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# BIOACTIVE COMPOUND OF HOLOTHOROIDEA

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## PREFACE

Praise be to Allah, gratitude is rightly poured forth to His presence. With the blessings, guidance, and favors from Him, the author has successfully completed the book entitled "Bioactive Compound of Holothuroidea" This book represents the author's concern for the utilization of natural resources derived from the marine biota of Indonesia.

Therefore, the existence of this book is undoubtedly a result of the contributions and assistance from all parties. Thus, on this occasion, the author extends appreciation and expresses gratitude to all those who have contributed significantly to the process of creating this book.

As imperfect beings, the contents of this book are still far from perfection. Thus, we earnestly welcome criticisms and suggestions from all parties. Finally, we hope that the presence of this book will be beneficial to all of us. Aamiin.

Bandung, August 2023

**Author Team** 

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## **OVERVIEW BIOACTIVE HOLOTHOROIDEA**

#### A. INTRODUCTION

Changes in lifestyle, such as diet and physical activity, increase the risk of developing diseases, which include diabetes, obesity, hypertension, cancer and others. Several prevention strategies have been tested, including the use of multiple bioactive components found in living things in the form of functional food ingredients. Sea cucumbers are a source of bioactive components that can be used in biopharmaceuticals, health foods, and industrial raw materials. Sea cucumbers have a high protein content and a variety of bioactive components. Sea cucumbers are invertebrates or thorn-skinned animals (Echinodermata) with an elongated cylindrical body and oral and aboral lines connecting the anterior and posterior parts as the axis. The potential for sea cucumbers from capture fisheries in Indonesia is quite significant, representing a 51.37% increase in average production.1

Sea cucumbers have been used and studied for a long time. The Chinese have known sea cucumbers as a food with medicinal properties since the Ming dynasty. Sea cucumber bioactive ingredients act as antioxidants (reduce cell and tissue damage), antibacterial, antifungal, antinociceptive (painkillers), and anti-inflammatory agents (fight inflammation and reduce swelling). Among the bioactive components are mucopolysaccharides, glucosamine and chondroitin sulfate, minerals and trace minerals, steroids, collagen, Omega 3 - DHA, and

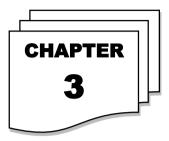


### COMPOUND CONCEPTS

#### A. DEFINITIONS OF COMPOUNDS

Compounds are substances that can be formed in combination with elements with these subdivisions. A compound must be made by a chemical reaction between two or more elements in order to go through the reactions in formation. Compounds are substances formed by two or more elements. Through chemical reactions, compounds can be broken down into their constituent elements. In addition, the compound is also defined as a single substance that can be broken down into other simpler substances through chemical reactions. An example of a compound that we often encounter in everyday life is water. Water is a combination of the elements hydrogen (H) and oxygen (O) with the chemical formula, namely H20. Through chemical reactions, water can be decomposed back into hydrogen and oxygen. Even though at atmospheric pressure, hydrogen and oxygen are both gaseous, when they unite and bond with each other, their form can turn into a liquid. (Alex, 2023),

A compound is a single substance that can be broken down into two or more elements. Compounds have several elements that combine chemically with each other, so that the symbol for a compound consists of several element symbols. Analysis to find the constituent elements of a compound is usually called qualitative analysis, whereas if the analysis finds a comparison of each

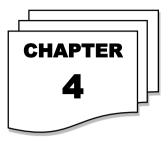


### HOLOTHOROIDEA

#### A. THE DEFINITION OF HOLOTHUROIDEA

Holothuroidea in Indonesian is called sea cucumbers or sea cucumbers. Is a group of invertebrate animals that are found in many oceans around the world. They move slowly and live on the ocean floor. In the marine ecosystem, sea cucumbers are a group of animals that play a very important role in the ecosystem. They are deposit and suspension feeders.

Sea cucumbers or better known as sea cucumbers are one of the organisms from the phylum Echinodermata class Holothuroidea. Sea cucumbers (Holothuroidea) can be found or found throughout coastal waters, from shallow tidal areas to deeper waters for their life, sea cucumbers prefer polluted free waters, and the water is relatively calm. In general, each species has a specific habitat, for example, the white sea cucumber (Holothuria scabra) is found in seagrass-covered waters, while the Koro sea cucumber (Muelleria leconoro) and sand sea cucumber are found in deeper waters (Martoyo et al. al., 2007). The main habitat of sea cucumbers is on seagrasses and corals. The spread of sea cucumbers in Indonesia is very wide, including; coastal waters of Madura, East Java, Bali, Sumba, Lombok, Aceh, Bengkulu, Bangka, Riau and its surroundings, Belitung, Kalimantan (west, east and south), Sulawesi, Maluku, Papua and the Thousand Islands (Martoyo et al., 2007) . In (Handayani et al., 2017).



## STUDY ANALYSIS BIOACTIVE HOLOTHOROIDEA

### A. LINKAGE AND CLUSTERING OF THEMES IN POTENTIAL BIOACTIVE COMPOUND HOLOTHOROIDEA

This category describes the ideas contained in 132 visualization techniques discovered in 148 articles that are related to the study's theme. A VOSviewer review also revealed 12 concept clusters (see table 1). Figure 4.1 depicts the concept identities deduced from the cluster density view. Furthermore, the color code for each cluster is used to identify the key concepts of each cluster. The goal is to identify as many themes as possible that have been frequently discussed in previous research and to make them available for future use. Figure 4.1 depicts the density of clusters as indicated by the different colors of each cluster.



## **CONCLUDING CONCLUSION**

#### A. CONCLUSION

Sea cucumbers, particularly H.atra, H.scabra, C.frondosa, and A.japonicus, contain antioxidants such as phenolic acids, flavonoids, peptides, fucosylated chondroitin sulfate (FCS), fucoidan, and triterpene glycosides. These compounds may also have anticancer, anti-inflammatory, anti-glycation, anti-tyrosinase, anti-hypertension, antithrombotic, anti-diabetic, and antimicrobial properties. As a result, sea cucumber antioxidants have the potential to be used in nutraceuticals, pharmaceuticals, cosmetics, and functional foods. More research is needed to understand the detailed chemical structures, mechanisms of action, and bioaccessibility and bioavailability of sea cucumber-derived value-added products through in vivo analysis and clinical trials in order to support the health claims and commercialize sea cucumber-derived value-added products.

Sea cucumbers are marine invertebrates that have potential as anticancer. Sea cucumber contains a significant amount of protein, vitamins, fatty acid, collagen, essential and non-essential amino acids, and minerals that are beneficial to the body. Active compound as anticancer such as philinopside A and B, patagonicosides, holothurin A and echinosides, colohiroside A, Intercedenside A, Okhotosides and Frondoside A sticoposide originating from Thelenota anax, sticoposide, bivittoside A, Holocosinos A and Holotox, cucumariosides produced from different types of sea cucumbers. Each active

## REFERENCES

Alex (2023), Compound, In https://pengajar.co.id/senyawa-Jadi/

- Bitar, (2023), Compounds: Definition, Characteristics, Properties, Kinds, Differences, Formulas, Abbreviations, Roles in https://www.gurudinding.co.id/senvawa/
- Dakrory AI, Fahmy SR, Soliman AM, Mohamed AS, Amer SAM. Protective and curative effects of the sea cucumber Holothuria atra extract against DMBA-induced Hepatorenal diseases in rats. Biomed Res Int. 2015;2015.
- Daluningrum Wahyu Pranata Ika, (2009) Early Screening of Bioactive Components from Blood Clams (Anadara granosa) As Antibacterial Compounds, Agricultural Institute: Bogor
- Dang HF, Han X, Guo Y, Li Q, Ye SG, Liu J, et al. Expression profiles of immunerelated genes in coelomocytes during regeneration after evisceration in Apostichopus japonicus. ... Surviv J [Internet]. 2020; Available from: https://www.isj.unimore.it/index.php/ISJ/article/view/610
- Detikedu, (2021) Compounds, in https://www.detik.com/edu/detikpedia/d-5876913/senyawa-pengertian-tipe-dan-example.
- EREGUERO MG, CORDERO MA, DALAYAP R, ... Antifungal activity of selected sea cucumber species from Tukuran, Zamboanga del Sur, Mindanao, Philippines using modified SPOTi assay [Internet]. Biodiversitas Journal of .... smujo.id; 2022. Available from:

https://www.smujo.id/biodiv/article/download/12677/6301

Fonseca VJA, Braga AL, Filho JR, ... A review on the antimicrobial properties of lectins. Int J ... [Internet]. 2022; Available from: https://www.sciencedirect.com/science/article/pii/S014181302102614

3

Gueadi (2020) Holothuroidea, in https://tipe.net/holothuroidea/

Guo K, Su L, Wang Y, Liu H, Lin J, Cheng P, et al. Antioxidant and anti-aging effects of a sea cucumber protein hydrolyzate and bioinformatic characterization of its composing peptides. Food Funct. 2020;11(6):5004–16. Hossain A, Dave D, Shahidi F. Antioxidant Potential of Sea Cucumbers and Their Beneficial Effects on Human Health. Mar Drugs. 2022;20(8):1–22.

- Hossain A, Dave D, Shahidi F. Northern sea cucumber (Cucumaria frondosa): A potential candidate for functional food, nutraceutical, and pharmaceutical sector. Mar Drugs. 2020;18(5).
- Kalinin VI, Silchenko AS, Avilov SA, Stonik VA. Progress in the Studies of Triterpene Glycosides From Sea Cucumbers (Holothuroidea, Echinodermata) Between 2017 and 2021. Nat Prod Commun. 2021;16(10).
- Karthikeyan A, Joseph A, Nair BG. Promising bioactive compounds from the marine environment and their potential effects on various diseases. J Genet Eng Biotechnol [Internet]. 2022;20(1). Available from: https://doi.org/10.1186/s43141-021-00290-4
- Khotimchenko Y. Pharmacological potential of sea cucumbers. Int J Mol Sci [Internet]. 2018; Available from: https://www.mdpi.com/288380
- Kim B, Han JW, Thi Ngo M, Le Dang Q, Kim JC, Kim H, et al. Identification of novel compounds, oleanane- and ursane-type triterpene glycosides, from Trevesia palmata: their biocontrol activity against phytopathogenic fungi. Sci Rep [Internet]. 2018;8(1):1–11. Available from: http://dx.doi.org/10.1038/s41598-018-32956-4
- Kim S, Lee J, Oh DB, Kwon O. Marine invertebrate sialyltransferase of the sea squirt Ciona savignyi sialylated core 1 O-linked glycans [Internet]. ... Journal of Biological Macromolecules. Elsevier; 2022. Available from: https://www.sciencedirect.com/science/article/pii/S014181302102482 X
- Mondol MAM, Shin HJ, Rahman MA, Islam MT. Sea cucumber glycosides: Chemical structures, producing species and important biological properties. Mar Drugs. 2017;15(10).
- Naglah AM, Al-omar MA, Almehizia AA, Obaidullah AJ, Bhat MA, Kalmouch A, et al. Synthesis, characterization, and anti-diabetic activity of some novel vanadium-folate-amino acid materials. Biomolecules [Internet]. 2020;10(5). Available from:

https://www.scopus.com/inward/record.uri?eid=2-s2.0-85085175228&doi=10.3390%2Fbiom10050781&partnerID=40&md5=ff 2173ba1a4634f3adcfe90ac32ef688

42 | Bioactive Compound of Holothoroidea

Nugroho A, Harahap IA, Ardiansyah A, Bayu A, ... Antioxidant and antibacterial activities in 21 species of Indonesian sea cucumbers. J Food Sci ... [Internet]. 2022; Available from:

https://link.springer.com/article/10.1007/s13197-021-05007-6

- OECD. Sustainable Ocean Economy Country Diagnostics of Indonesia. Sustainable Ocean for All Series. 2021.
- Panagos CG, Thomson DS, Moss C, Hughes AD, Kelly MS, Liu Y, et al. Fucosylated chondroitin sulfates from the body wall of the sea cucumber Holothuria forskali: Conformation, selectin binding, and biological activity. J Biol Chem. 2014;289(41):28284–98.
- Pangestuti R, Arifin Z. Medicinal and health benefit effects of functional sea cucumbers [Internet]. Journal of traditional and complementary medicine. Elsevier; 2018. Available from: https://www.sciencedirect.com/science/article/pii/S222541101730069

х

- Pangestuti R, Arifin Z. Medicinal and health benefit effects of functional sea cucumbers. J Tradit Complement Med. 2018;8(3):341–51.
- Pankey. Prospect of Sea Cucumber Culture in Indonesia A Potential Food Sources. J Coast Development [Internet]. 2012;15(2):114–24. Available from: Sulawesi Utara
- Pranweerapaiboon K, Garon A, Seidel T, ... In vitro and in silico studies of holothurin A on androgen receptor in prostate cancer. J ... [Internet]. 2021; Available from:

https://www.tandfonline.com/doi/abs/10.1080/07391102.2021.19755 62

- Purnomo Gery, (2020) Teripang Atau Timun Laut; Klasifikasi, Morfologi, Habitat, dalam https://www.melekperikanan.com/2020/01/teripang.html
- Quilliot D, Coupaye M, Ciangura C, Czernichow S, Sallé A, Gaborit B, et al. Recommendations for nutritional care after bariatric surgery: Recommendations for best practice and SOFFCO-MM/AFERO/SFNCM/ expert consensus. J Visc Surg [Internet]. 2021;158(1):51–61. Available from:

https://www.scopus.com/inward/record.uri?eid=2-s2.0-

85099132196&doi=10.1016%2Fj.jviscsurg.2020.10.013&partnerID=40& md5=f96e537bbcae4caa1f6eb429ded90548

- Rasyid A, Wahyuningsih T, Ardiansyah A. PROFIL METABOLIT SEKUNDER, AKTIVITAS ANTIBAKTERI DAN KOMPOSISI SENYAWA YANG TERKANDUNG DALAM EKSTRAK METANOL TERIPANG Stichopus horrens. J Ilmu dan Teknol Kelaut Trop. 2018;10(2):333–40.
- Reithuber E, Nannapaneni P, Rzhepishevska O, Lindgren AEG, Ilchenko O, Normark S, et al. The bactericidal fatty acid mimetic 2CCA-1 selectively targets pneumococcal extracellular polyunsaturated fatty acid metabolism. MBio. 2020;11(6):1–16.
- Senadheera TRL, Hossain A, Dave D, Shahidi F. In Silico Analysis of Bioactive Peptides Produced from Underutilized Sea Cucumber By-Products—A Bioinformatics Approach. Mar Drugs. 2022;20(10).
- Shi S, Feng W, Hu S, Liang S, An N, Mao Y. Bioactive compounds of sea cucumbers and their therapeutic effects. Chinese J Oceanol Limnol. 2016;34(3):549–58.
- Suryaningrum TD. Teripang : Potensinya sebagai bahan nutraceutical. Squalen. 2008;3(2):63–69.
- Telahigue K, Ghali R, Nouiri E, Labidi A, ... Antibacterial activities and bioactive compounds of the ethyl acetate extract of the sea cucumber Holothuria forskali from Tunisian coasts. J Mar ... [Internet]. 2020; Available from: https://www.cambridge.org/core/journals/journal-of-the-marinebiological-association-of-the-united-kingdom/article/antibacterialactivities-and-bioactive-compounds-of-the-ethyl-acetate-extract-ofthe-sea-cucumber-holothuria-forskali-from-tunisian-coasts/8
- Telahigue K, Ghali R, Nouiri E, Labidi A, ... Antibacterial activities and bioactive compounds of the ethyl acetate extract of the sea cucumber Holothuria forskali from Tunisian coasts. J Mar ... [Internet]. 2020; Available from: https://www.cambridge.org/core/journals/journal-of-the-marinebiological-association-of-the-united-kingdom/article/antibacterialactivities-and-bioactive-compounds-of-the-ethyl-acetate-extract-ofthe-sea-cucumber-holothuria-forskali-from-tunisian-coasts/8
- Thongjuy R, ... Effects of scabraside D on cell viability inhibition and apoptosis promotion of human hepatocellular carcinoma [Internet]. Journal of Applied Animal .... vs.mahidol.ac.th; 2018. Available from: https://vs.mahidol.ac.th/jaas/Files/Vol11No2/RS K.Rassamee Final.pdf

- Vien LT, Hanh TTH, Quang TH, Thanh N Van, Thao DT, Cuong NX, et al. Triterpene Tetraglycosides From Stichopus Herrmanni Semper, 1868. Nat Prod Commun. 2022;17(5):5–9.
- Wu Y, Jiang H, Lin JS, Liu J, Wu CJ, Xu R. Antioxidant, hypolipidemic and hepatic protective activities of polysaccharides from phascolosoma esculenta. Mar Drugs. 2020;18(3).
- Wulandari DA, Syahputra G, ... The Bioactive Compound and Mechanism of Action of Sea Cucumber (Holothuridae) as Anticancer: A Review. J Pure ... [Internet]. 2020;9(3):153–70. Available from:

https://www.researchgate.net/profile/Gita-Syahputra/publication/346886210\_The\_Bioactive\_Compound\_and\_Me chanism\_of\_Action\_of\_Sea\_Cucumber\_Holothuridae\_As\_Anticancer\_A \_Review/links/5fd1bd3492851c00f8623b65/The-Bioactive-Compoundand-Mechanism-of-Action-of-S

Xu C, Zhang R, Wen Z. Bioactive compounds and biological functions of sea cucumbers as potential functional foods. J Funct Foods [Internet]. 2018; Available from:

https://www.sciencedirect.com/science/article/pii/S175646461830433 X

- Yano A, Abe A, Aizawa F, Yamada H, Minami K, Matsui M, et al. The effect of eating sea cucumber jelly on Candida load in the oral cavity of elderly individuals in a nursing home. Mar Drugs. 2013;11(12):4993–5007.
- Yoon BK, Jackman JA, Valle-González ER, Cho NJ. Antibacterial free fatty acids and monoglycerides: Biological activities, experimental testing, and therapeutic applications. Vol. 19, International Journal of Molecular Sciences. 2018.
- Yuan WH, Yi YH, Tang HF, Liu BS, Wang ZL, Sun GQ, et al. Antifungal triterpene glycosides from the sea cucumber Bohadschia marmorata. Planta Med. 2009;75(2):168–73.
- Yuniati R, Sulardiono B. Effectivity of Holothuria scabra and Spirulina platensis extract combination as an Antiinflammatory Agent Measured by Carrageenan-induced Rat Paw Edema [Internet]. ILMU KELAUTAN: Indonesian Journal .... scholar.archive.org; 2020. Available from: https://scholar.archive.org/work/pdw3lboyszemlc76i546salblm/access

/wayback/https://ejournal.undip.ac.id/index.php/ijms/article/downloa d/28728/pdf

- Zheng Koh DH, Saheki Y. Regulation of Plasma Membrane Sterol Homeostasis by Nonvesicular Lipid Transport. Contact. 2021;4:251525642110424.
- Zhou X, Zhou D-Y, Yin F-W, Song L, Liu Y-X, Xie H-K, et al. Glycerophospholipids in sea cucumber (Stichopus japonicus) and its processing by-products serve as bioactives and functional food ingredients. J Food Bioact. 2018;1:134–42.

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Irena Ujianti is a highly accomplished and respected lecturer in medical physiology at the Faculty of Medicine, University of Muhammadiyah Prof. Dr. Hamka. With a Doctorate degree in physiology from the prestigious Faculty of Medicine, University of Indonesia, Irena's research interests primarily revolve around nutrition and

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Irena's expertise lies in the application of physiological and biomolecular methods to investigate complex medical conditions. In addition to her academic responsibilities, Irena is an active member of several professional organizations, including the Indonesian Doctors Association (IDI), the Indonesian Association of Clinical Physiologists (PDFKI), and the Jakarta Association of Indonesian Physiological Sciences (IAIFI).

Irena's dedication to her field is further demonstrated through her involvement in several ongoing research projects related to nutrition and metabolic diseases. Her contributions to the field have been widely recognized, making her a highly sought-after speaker and consultant in the medical community.

## BIOACTIVE COMPOUND OF HOLOTHOROIDEA

Holothoroidea (sea cucumbers) are a commercially significant variety of marine invertebrates that are widely consumed in Asian countries. Sea cucumbers contain various bioactive substances such as lipids, phenols, peptides, triterpene glycosides, and polysaccharides, which have a wide range of beneficial biological effects. Our research aims to provide a thorough understanding of these bioactive compounds. We discuss natural stock fishery and aquaculture as management strategies for ensuring sustainable sea cucumber populations. We also describe the extraction and purification of bioactive compounds, providing insights into the preparation of functional ingredients derived from sea cucumbers. This review aims to give academics and industry a better understanding of sea cucumbers and their potential for the development of high-value nutraceutical products.

The goal of our study is to categorize themes related to the study of Holothoroidea bioactive components and mechanisms. We analyzed 200 research articles using descriptive analysis and the Nvivo-12 software. Our findings revealed 151 relationships between the bioactive components of Holothoroidea, which could be classified into 11 groups. The discovery of the mechanism of action of the bioactive components of Holothoroidea is significant as it can aid in the development of a conceptual framework for the study of these organisms. However, a limitation of this study is the lack of references to the articles reviewed. Future research should employ a comparative analysis approach to address this limitation.



