

Undaria pinnatifida and Cancer: A Literature Review

Ana Rita Rente ¹, Patrícia Correia ^{1,2}, Cláudia Pinho ^{1,2}

¹ Escola Superior de Saúde, Instituto Politécnico do Porto, R. Dr. António Bernardino de Almeida 400, 4200-072 Porto, Portugal

² REQUIMTE/LAQV, Escola Superior de Saúde, Instituto Politécnico do Porto, Rua Dr. António Bernardino de Almeida, 4200-072 Porto, Portugal

* rita-rente@hotmail.com

Background: Cancer is a prevalent chronic disease and a leading cause of death worldwide, with treatment options having limited effectiveness and severe side effects [1,2]. Searching for new therapeutic options has been conducted over the years, including research with natural compounds [1]. *Undaria pinnatifida* (known as wakame), presents high nutritional quality and bioactivity due to its composition mainly of polysaccharides (fucoidan), carotenoids (fucoxanthin) and proteins [1–4]. Some research has been performed on the potential biological activities of the *U. pinnatifida* constituents, including its anti-cancer properties [5,6]. **Objective:** The main objective of this study is to discuss the *in vitro* studies regarding cytotoxic activity of wakame and its compounds against different cancer cell lines. **Methods:** A narrative review was conducted using the PubMed® database and a combination of keywords with Boolean search terms, to find articles published between 2015 and 2025, related to *in vitro* cytotoxic activity of *U. pinnatifida* and its compounds. **Results:** Twenty articles were selected. Wakame and its compounds (polysaccharides or fucoidan and fucoxanthin) have demonstrated *in vitro* cytotoxic effects on several cancer cell lines, including prostate (PC-3), cervical (HeLa 229 and SiHa), ovarian (A2780 and SKOV3), pancreatic (PANC-1), breast (MCF7 and MDA-MB-231), lung (A549) and melanoma [7–16]. Nanoformulations have shown considerable potential, since enhanced delivery of fucoidan and quinacrine resulting in a reduction in IC₅₀ values [11]. Fucoidan administered concomitantly with other drugs demonstrated synergistic (aspirin, paclitaxel, lapatinib or tamoxifen) and additive (topotecan) effects [7,14,17]. Additionally, anti-cancer effect may be explained through the induction of apoptosis and inhibition of cell proliferation [7,10–13,16]. **Conclusions:** *U. pinnatifida* and its compounds showed anticancer properties and potential enhanced efficacy of other compounds when administered concomitantly. Future research should focus on pharmacokinetics, toxicity assessment, mechanism of action and *in vivo* efficacy to establish its potential as a therapeutic agent.

keywords: *Undaria pinnatifida*; wakame; cancer; cytotoxic

Acknowledgements: This research did not receive external funding.

References

1. Nadeeshani H, Hassouna A, Lu J. Proteins extracted from seaweed *Undaria pinnatifida* and their potential uses as foods and nutraceuticals. *Crit Rev Food Sci Nutr*. 2021 Mar;1–17.
2. Phull AR, Kim SJ. *Undaria pinnatifida* a Rich Marine Reservoir of Nutritional and Pharmacological Potential: Insights into Growth Signaling and Apoptosis Mechanisms in Cancer. *Nutr Cancer*. 2018;70(6):956–70.
3. Hwang JA, Islam MM, Ahmed ST, Mun HS, Kim GM, Kim YJ, et al. Seamustard (*Undaria pinnatifida*) Improves Growth, Immunity, Fatty Acid Profile and Reduces Cholesterol in Hanwoo Steers. *Asian-Australas J Anim Sci*. 2014 Aug;27(8):1114–23.
4. Neri TAN, Rohmah Z, Ticar BF, Palmos GN, Choi BD. Evaluation of sea mustard (*Undaria pinnatifida*) sporophylls from South Korea as fucoidan source and its corresponding antioxidant activities. *Fish Aquatic Sci* [Internet]. 2019 Dec 19;22(1):24. Available from: <https://fas.biomedcentral.com/articles/10.1186/s41240-019-0141-4>
5. Gammone M, D'Orazio N. Anti-Obesity Activity of the Marine Carotenoid Fucoxanthin. *Mar Drugs*. 2015 Apr 13;13(4):2196–214.
6. Pereira AG, Otero P, Echave J, Carreira-Casais A, Chamorro F, Collazo N, et al. Xanthophylls from the Sea: Algae as Source of Bioactive Carotenoids. *Mar Drugs*. 2021 Mar 27;19(4):188.
7. Zhou R, Zhong L, Jia S, Luo Y, Li Y, Tang Y. Preparation, and characterization of aspirin–fucoidan complex and its admirable antitumor activity on human non-small cell lung cancer cells. *Int J Biol Macromol*. 2024 Apr; 263:130163.
8. Qiao F, Yu X, Tie S, Chen Y, Hou S, Tan M. Zinc delivery system constructed from food-borne nanoparticles derived from *Undaria pinnatifida*. *Food Funct*. 2021;12(18):8626–34.

9. Queffelec J, Flórez-Fernández N, Domínguez H, Torres MD. Microwave hydrothermal processing of *Undaria pinnatifida* for bioactive peptides. *Bioresour Technol.* 2021 Dec, **342**,125882.
10. Park AY, Nafia I, Stringer DN, Karpiniec SS, Fitton JH. Fucoïdan Independently Enhances Activity in Human Immune Cells and Has a Cytostatic Effect on Prostate Cancer Cells in the Presence of Nivolumab. *Mar Drugs.* 2021 Dec 22, **20**(1),12.
11. Etman SM, Mehanna RA, Bary AA, Elnaggar YSR, Abdallah OY. *Undaria pinnatifida* fucoidan nanoparticles loaded with quinacrine attenuate growth and metastasis of pancreatic cancer. *Int J Biol Macromol.* 2021 Feb, **170**, 284–97.
12. Etman SM, Abdallah OY, Elnaggar YSR. Novel fucoidan based bioactive targeted nanoparticles from *Undaria Pinnatifida* for treatment of pancreatic cancer. *Int J Biol Macromol.* 2020 Feb, **145**,390–401.
13. Wu J, Li H, Wang X, Zhang X, Liu W, Wang Y, et al. Effect of polysaccharide from *Undaria pinnatifida* on proliferation, migration, and apoptosis of breast cancer cell MCF7. *Int J Biol Macromol.* 2019 Jan, **121**,734–42.
14. Mathew L, Burney M, Gaikwad A, Nyshadham P, Nugent EK, Gonzalez A, et al. Preclinical Evaluation of Safety of Fucoïdan Extracts from *Undaria pinnatifida* and *Fucus vesiculosus* for Use in Cancer Treatment. *Integr Cancer Ther.* 2017 Dec 21, **16**(4),572–84.
15. Corban M, Ambrose M, Pagnon J, Stringer D, Karpiniec S, Park A, et al. Pathway Analysis of Fucoïdan Activity Using a Yeast Gene Deletion Library Screen. *Mar Drugs.* 2019 Jan 14, **17**(1).
16. Lu J, Shi KK, Chen S, Wang J, Hassouna A, White LN, et al. Fucoïdan Extracted from the New Zealand *Undaria pinnatifida*-Physicochemical Comparison against Five Other Fucoïdians: Unique Low Molecular Weight Fraction Bioactivity in Breast Cancer Cell Lines. *Mar Drugs.* 2018 Nov 22, **16**(12),461.
17. Thakur V, Lu J, Roscilli G, Aurisicchio L, Cappelletti M, Pavoni E, et al. The natural compound fucoidan from New Zealand *Undaria pinnatifida* synergizes with the ERBB inhibitor lapatinib enhancing melanoma growth inhibition. *Oncotarget.* 2017 Mar 14, **8**(11),17887–96.