

Clinical Applications of Oligo Fucoidan in Translational Medicine for Adjuvant Cancer Therapy



OliFuco[®] Oligo Fucoidan





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Health Ingredients Verified with **Human Clinical Trials**

Fucoidan is a kind of polysaccharide substance which mainly exists in the brown seaweed. Hi-Q Marine Biotech has developed OLIFUCO® Oligo Fucoidan by the advanced hydrolysis technology to extract the bioactive Low Molecular Weight Fucoidan (Oligo Fucoidan, 500-1500 Daltons) from natural oceanic brown seaweed. Oligo Fucoidan is verified by scientific studies and human clinical trials with multi-health benefits. Hi-Q Oligo Fucoidan has various pharmacological and biological activities, such as anti-cancer, anti-inflammatory, and immunomodulatory functions. In addition, it is proven to protect against or reduce toxicity from chemotherapeutic agents and radiation for cancer treatments. Oligo Fucoidan has been recently marketed as a dietary supplement or nutraceutical for various diseases, including cancer.



Making Difference in Cancer with Clinically Tested Adjunct Supplement

With the passion of empowering people to lead healthier lives, Hi-Q has been investing in R&D and product development of Oligo Fucoidan in dietary health supplement as adjunct supplement to cancer patients to relieve the side effects from cachexia. Hi-Q has joined collaboration with PhD scientists and medical doctors in leading research institutions, universities hospitals for continue Oligo Fucoidan scientific studies and human clinical trials.

Hi-Q Oligo Fucoidan, the health ingredient is marketed under the tradename OLIFUCO®. Hi-Q has its own branded (Hi-Q® Health, FucoHiQ®) dietary supplements the products are researched and developed by using this scientific, clinically proven ingredient. Hi-Q is a leading fucoidan company and has 13 years of successful sales history in Taiwan since 2009. Hi-Q's Oligo Fucoidan products are recommended by medical doctors and pharmacists.

Anti-Cancer Effects of Fucoidan Combinations As An Adjuvant

Several studies have reported that fucoidan and its combinations can induce apoptosis in a variety of cancers. Studies of fucoidan and combinations of fucoidan with clinical drugs in adjuvant settings have suggested that fucoidan may reduce the toxicity of certain anti-cancer drugs. These studies suggest that fucoidan could be used in adjuvant settings for cancer management. Synergistic effects of fucoidan with clinically used drugs in the inhibition of various tumors and/or cancers in mice have been reported. Synthetic drugs used as chemotherapeutics agents are frequently limited by their side effects and drug resistance. Fucoidan has been investigated as a dietary supplement or synergetic anti-tumor agent to reduce the toxicity and/or enhance the efficacy of chemotherapeutic.

Positive synergistic cytotoxic effects of fucoidan combined with cisplatin on both LLC1 mouse lung carcinoma cells and mice have been reported. A role of the Smurf2-dependent ubiquitin-proteasome pathway in TGF receptor degradation has been proposed. Fucoidan upregulates TLR4/CHOP-mediated caspase-3 and PARP activation and enhances cisplatin-induced cytotoxicity in human lung cancer cells. Moreover, in human clinical studies, combined cisplatin and fucoidan treatment effectively increased the survival rate of patients with lung cancer in Taiwan. The results indicated that fucoidan exerts a greater anti-tumorigenic effect as an adjuvant, second-line anti-cancer diet supplement and may also regulate immune functions, while cisplatin remains a first-line anti-cancer drug for chemotherapy in clinical human lung cancer treatment.

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- Immune-modulatory effects
- · Enhance proliferation of natural killer (NKs) cells and cytotoxic
- · Inhibit proliferation/metastasis of cancer cell
- · Inhibit angiogenesis of cancer
- · Induce apoptosis of cancer cell
- · Prevent tumor progression, alter tumor microenvironment (TME)
- Improve outcome of chemotherapy & radiotherapy
- · Reduce side effects of chemotherapy & radiotherapy
- · Reduce radiation-induced fibrosis
- Improve cancer cachexia
- Improve diseases control rate (DCR), be used as adjunct therapy supplement alongside conventional oncology treatments

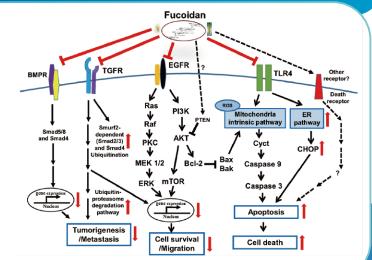
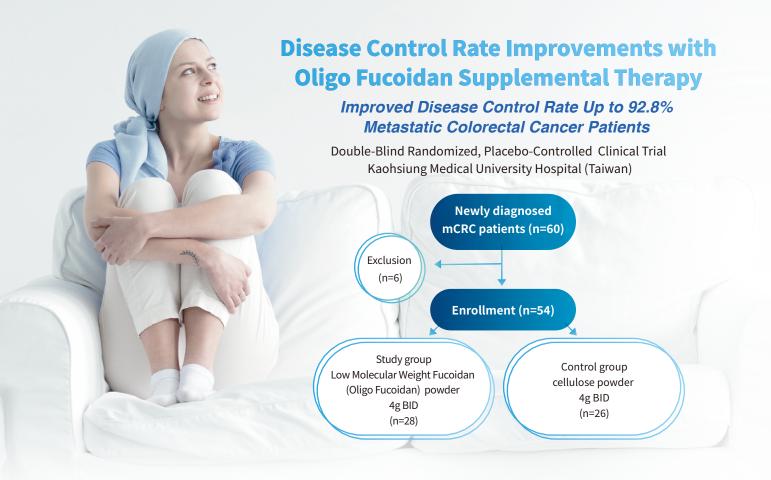


Fig. Fucoidan interacts receptors-mediated anticancer pathways and reactions

The scientific studies result show that Oligo fucoidan, whether through in vitro to in vivo research studies or clinical trials in humans, has been proven to produce positive bioactive and synergy effects of adjuvant therapy on cancer treatment. This allows molecular mechanisms in cancer research to be applied in adjuvant cancer treatment, in line with the pursuit of translational medicine and the mindset of establishing a direct link between basic medical research and clinical application. Oligo Fucoidan also has attained its inclusion in the Dictionary of Drugs of the U.S. National Cancer Institute (NCI) (Code C170752).

More about Hi-Q's research articles





In 2017, Hi-O cooperated with medical expert team from Kaohsiung Medical University Hospital and performed the first prospective, randomized, double-blind, controlled trial to investigate the efficacy of Low Molecular Weight Fucoidan (Oligo Fucoidan) as a supplemental therapy to chemotarget agents in patients with metastatic colorectal cancer (mCRC). The results indicated that Oligo Fucoidan combined with chemotarget agents significantly improved the Disease Control Rate (DCR). This study provided insights into the development of cancer treatments, particularly in the combination of natural fucoidan with chemotarget agents.

Primary Endpoints

The DCR (Disease Control Rate), defined as the sum of the CR, PR, and SD rates, was significantly higher (by 23.6%) in the study group than in the control group (92.8% vs. 69.2%; p = 0.026).

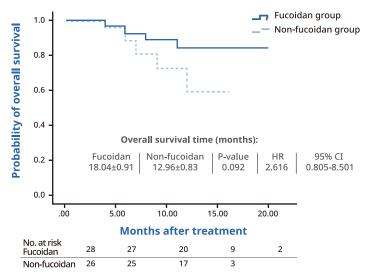
Total Study Group Control Group

	lotal	Study Group	Control Group	
	(n=54)	(n=28)	(n=26)	p-value
Disease control rate				0.026
Yes (CR+PR+SD) *	44	26 (92.8)	18 (69.2)	
No (PD) *	10	2 (7.2)	8 (30.8)	
Objective response rate				0.284
Yes (CR+PR) *	29	17 (60.7)	12 (46.2)	
No (SD+PD) *	25	11 (39.3)	14 (53.8)	

^{*} CR: complete response; PR: partial response; SD: stable disease; PD: progressive disease classified using RECIST criteria. Version1.1

Secondary Endpoints

Secondary Outcome CR and PR rates) was comparable in the study group and the control group (60.7% vs. 46.2%; p = 0.284). Compared with the control group, the study group exhibited a trend of improved Overall Survival Rate (18.04 \pm 0.91 vs. 12.96 \pm 0.83 months: p = 0.092).



No adverse effects were observed in both groups during the trial period.

Full Article:

2017 [Marine Drugs], Efficacy of Low-Molecular-Weight Fucoidan as a Supplemental Therapy in Metastatic Colorectal Cancer Patients: A Double-Blind Randomized Controlled Trial

Advanced research in Mechanism:

2021 [International Journal of Molecular Sciences], Low-Molecular-Weight Fucoidan as Complementary Therapy of Fluoropyrimidine-Based Chemotherapy in Colorectal Cancer







Hi-Q is Leading in Human Clinical Trials of Oligo Fucoidan in Cancer Studies and Mechanism Research

Hi-Q collaborates with leading research institutions, hospitals and universities for advanced research and solution. More than 50 international scientific articles were published. Among the articles, over 30 explore oncology, immunomodulation and inflammation studies. Not only animal model-based pre-clinical studies, Hi-Q also invests in more than 10 human clinical trials to prove the efficacy of Oligo Fucoidan.



Oncology

- 2022 [Journal of Biomedical Science], Oligo-Fucoidan Supplementation Enhances the Effect of Olaparib on Preventing Metastasis and Recurrence of Triple-Negative Breast Cancer
- 2021 [Topics in Companion Animal Medicine], The Use of Oligo Fucoidan in Cancer Bearing Dogs Undergoing Chemotherapy: A Double-Blinded Study
- 2021 [International Journal of Molecular Sciences], Low-Molecular-Weight Fucoidan as Complementary Therapy of Fluoropyrimidine-Based Chemotherapy in Colorectal Cancer
- 2021 [Frontiers in Oncology], Fucoidan Inhibits the Progression of Hepatocellular Carcinoma via Causing IncRNA Linc00261 Overexpression
- 2020 [Clinical and Translational Medicine], Low Molecular Weight Fucoidan Inhibits Hepatocarcinogenesis and Nonalcoholic Fatty Liver Disease in Zebrafish via ASGR/STAT3/HNF4A Signaling
- 2020 [Cancers], Low Molecular Weight Fucoidan Prevents Radiation-Induced Fibrosis and Secondary Tumors in a Zebrafish Model
- 2020 [Marine Drugs], Protective Effect of Low Molecular Weight Fucoidan on Radiation-Induced Fibrosis Through TGF-\(\beta\)1Smad Pathway Mediated Inhibition of Collagen I
- 2020 [Cancers], Oligo-Fucoidan Prevents M2 Macrophage Differentiation and HCT116 Tumor Progression
- 2020 [Pharmaceutics], Radioprotective Effect of Self-assembled Low Molecular Weight Fucoidan-Chitosan Nanoparticles
- 2019 [Clinical and Translational Medicine], Clinical Applications of Fucoidan in Translational Medicine for Adjuvant Cancer Therapy
- 2019 [Cytotechnology], The Anti-Tumor Activity of Brown Seaweed Oligo-Fucoidan via IncRNA Expression Modulation in HepG2 cells
- 2019 [Marine Drugs], Epigenetic Modification and Differentiation Induction of Malignant Glioma Cells by Oligo-Fucoidan
- 2018 [Marine Drugs], Fucoidan Inhibits Radiation-Induced Pneumonitis 2 and Lung Fibrosis by Reducing Inflammatory 3 Cytokine Expression in Lung Tissues
- 2018 [Cancer Letters], Fucoidan Upregulates TLR4/CHOP-Mediated Caspase-3 and PARP Activation to Enhance Cisplatin-Induced Cytotoxicity in Human Lung Cancer Cells
- 2017 [Scientific Reports], Fucoidan Induces Toll-like Receptor 4-Regulated Reactive Oxygen Species and Promotes Endoplasmic Reticulum Stress-Mediated Apoptosis in
- 2017 [Scientific Reports], Oligo-Fucoidan Prevents IL-6 and CCL2 Production and Cooperates with p53 to Suppress ATM Signaling and Tumor Progression

Human Clinical Trial

- 2017 [Marine Drugs], Efficacy of Low-Molecular-Weight Fucoidan as a Supplemental Therapy in Metastatic Colorectal Cancer Patients: A Double-Blind Randomized Controlled Trial
- 2016 [Journal of Cancer], Brown Seaweed Fucoidan Inhibits Cancer Progression by Dual Regulation of mir-29c/ADAM12 and miR-17-5p/PTEN Axes in Human Breast Cancer Cells
- 2016 [Oncotarget], Combined Administration of Fucoidan Ameliorates Tumor and Chemotherapy-Induced Skeletal Muscle Atrophy in Bladder Cancer-Bearing Mice
- 2015 [Marine Drugs], Fucoidan Elevates MicroRNA-29b to Regulate DNMT3B-MTSS1 Axis and Inhibit EMT in Human Hepatocellular Carcinoma Cells
- 2015 [Marine Drugs], Low Molecular Weight Fucoidan Inhibits Tumor Angiogenesis through Downregulation of HIF-1/VEGF Signaling under Hypoxia
- 2014 [Oncotarget], Fucoidan Inhibition of Lung Cancer in vivo and in vitro: Role of the Smurf2-dependent Ubiquitin Proteasome Pathway in TGFβ Receptor Degradation
- 2013 [CarcinoGenesis], Fucoidan Induces Changes in the Epithelial to Mesenchymal Transition and Decreases Metastasis by Enhancing Ubiquitin-Dependent TGF\$ Re+B18:C28ceptor Degradation in Breast Cancer



Immunomodulation and Anti-Inflammation

Human Clinical Trial

- 2022 [Scientific Reports], Effects of Oligo-Fucoidan on the Immune Response, Inflammatory Status and Pulmonary Function in Patients with Asthma: A Randomized, Double-Blind, Placebo-Controlled Trial
- 2021 [Scientific Reports], Oligosaccharides Ameliorate Acute Kidney Injury by Alleviating Cluster of Differentiation 44-Mediated Immune Responses in Renal Tubular Cells
- 2020 [Biomolecules], Transcriptomically-Revealed Oligo-Fucoidan Enhances the Immune System and Protects Hepatocytes via the ASGPR/STAT3/HNF4A

Human Clinical Trial

- 2020 [Nutrients], AxisThe 25(OH)Vitamin D Status Affected the Effectiveness of Oligo Fucoidan in Patients with Chronic Hepatitis B Virus Infection with Immune Tolerance Phase
- 2019 [Marine Drugs], Fucoidan Prevents RANKL-Stimulated Osteoclastogenesis and LPS-Induced Inflammatory Bone Loss via Regulation of Akt/GSK3B/PTEN/NFATc1 Signaling Pathway and Calcineurin Activity

Human Clinical Trial

- 2018 [Experimental and Therapeutic Medicine], Oligo-Fucoidan Improved Unbalance the Th1Th2 and TregTh17 Ratios in Asthmatic Patients An ex vivo Study
- 2016 [Food & Nutrition], Low-Molecular-Weight Fucoidan and High-Stability Fucoxanthin from Brown Seaweed Exert Prebiotics and Anti-Inflammatory Activities in Caco-2 Cells
- 2016 [Marine Drugs], The Oligo Fucoidan Inhibits Platelet-Derived Growth Factor-Stimulated Proliferation of Airway Smooth Muscle Cells
- 2015 [Journal of Food and Drug Analysis], Inhibitory Activity of Sargassum hemiphyllum Sulfated Polysaccharide in Arachidonic Acid-Induced Animal models of Inflammation
- 2011 [Journal of Agricultural and Food Chemistry], Inhibition of Lipopolysaccharide (LPS)-Induced Inflammatory Responses by Sargassum hemiphyllum Sulfated Polysaccharide Extract in RAW 264.7 Macrophage Cells
- 2010 [Journal of Marine Science and Technology], Antioxidant and Immune-Stimulating activities of Hot-Water Extract from Seaweed Sargassum hemiphyllum

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