

Find authenticated court documents without watermarks at docketalarm.com.

		PPLICATI a provisional						1.5
			VENTOR	والمتلافة وبوين ويتؤاكم				_
· · · · · · · · · · · · · · · · · · ·					Residence			
Given Name (first and middle [if any])		Family Name or Sumame		(City and either State or Foreign Country) Laval, Quebec, Canada		Country)		
		ANTALIS		Lavan, Caupea,	Omilia			
					••••••••••••••••••			
Additional inventors	are being nam	ed on thesepa	rately numbe	rad sheets atlache	d hereto	والمراجع والمتحد والمراجع		
				BO characters may	()			
ATURAL MARINE SOUR OLYUNSATURATED FA	tty acids an	D THEIR APPLICA	ng flavon Tions	oids,		•	•	
		COPPLE	ONDENCE					
Direct all correspondence to: CORRESPONDED				ADDRESS Place Customer Num			lumber	
Customer Number					Bar Code Label here			
OR	Type Cu	stomer Number hei	9				·······	
Firm or Individual Name SMART & BIGGAR								
Address Suite 3400								
Address		le la Gauchetiere S	T	· · · · · · · · · · · · · · · · · · ·		1		
City		eal, Quebec	State	544.054.4500	ZIP	514-954-1396		
Country		W5 Canada CLOSED APPLICA		514-954-1500	Fax	014-304-1030	, 	
Specification Num		21		CD(s), Numi		1		
Drawing(s) Numb	er of Sheets	13			r			
Application Data Si		R 1.76		Other (speci	(y)			
METHOD OF PAYMENT			VISIONAL A	PPLICATION FOR	PATENT (	check one)		
Applicant claims	small entity sta	lus. See 37 CFR 1.	27.			FILING		
	•	sea to cover the fil	•		•			
		Inorized to charge f to Deposit Account		19-2550	]	\$75.	00	
		TO-2038 is attache						
The Invention was made United States Governme		of the United States	Government	or under a contrac	it with an ag	ency of the		
No.								
Yes, the name of the L	J.S. Government a	gency and the Govern	ment contract r	lumber ara:				
Respectfully submitted,		واليوانلي معوديت المالية البراني			07/27/01		والكفيدية بالبسية بالته	
	Wola			Date		\	46474	
TYPED or PRINTED NAME Hans Koenig				(if appropriate)			46474	
613-232-2486				Docket Number:			86187-1	

This collection of information is required by 37 CFR 1.51. The information is used by the public to file (and by the PTO to process) a provisional application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 8 hours to complete, including gathering, preparing, and submitting the complete provisional application to the PTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, Washington, D.C. 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Box Provisional Application, Assistant Commissioner for Patents, Washington, D.C.

.

NATURAL MARINE SOURCE PHOSPHOLIPIDS COMPRISING FLAVONOIDS, POLYUNSATURATED FATTY ACIDS AND THEIR APPLICATIONS

#### Field of the Invention

The present invention is directed to nutraceutical, pharmaceutical or cosmetic compositions, particularly to phospholipid compositions derived from natural marine or aquatic sources.

Background of the Invention

Phospholipids are complex lipids containing phosphorus. The phosphatides, known as phospholipids, are usually divided into groups on the basis of compounds from which they are derived. In addition to two chains of fatty acids they contain phosphoric acid, glycerol and nitrogenous bases such as choline. The phospholipids considered most important are phosphatidylcholine, phosphatidylethanolamine and phosphatidylinositol. Their nature as amphophilic molecules provides them with unique physicochamical properties. Their function as the principle components of cell membranes makes phospholipids essential for all vital cell processes. They are widespread as secretory and structural components of the body and can mimic or enhance natural physiological processes.

Phosphelipid production may be either synthetic or through extraction from natural tissues. The chief source of commercial natural phospholipids is soybean, egg yolk and cows (brain and liver). Since an individual phospholipid may contain a variety of fatty acid residues, it may be described as pure only with this limitation in mind. Naturally occurring essential polyunsaturated fatty acids, can contribute to the activation of cellular metabolism. The main fatty acid found in phospholipid products is lineleic acid, present in soybean at more than 65%. The longest chain polyunsaturated fatty acids found in commercial available phospholipids either as

DOCKET

Find authenticated court documents without watermarks at docketalarm.com.

preparations or individually are 20:4 among the elcosanoids, known as arachidonic acid and 22:6 known as decosahexanoic acid.

Arachidonic acid is a fatty acid that is found as part of the phospholipid membrane, generally as part of phosphatidylcholine and phosphatidylinositol. Adverse callular stimuli will activate enzymes (phospholipase) that cleave arachidonic acid from the phospholipid backbone in the cell membrane. Arachidonic acid, which serves as the precursor for prostaglandins and prostacyclin (PGs, PGI<sub>2</sub>) and thromboxane (TXs) can then be metabolized by one of two major pathways: the cyclooxygenase (COX) pathway or the ilpoxygenase pathway. The COX pathway products, PGG<sub>2</sub> and PGH<sub>2</sub> can then be acted upon by thromboxane synthase (in platelets) or prostacyclin synthase (in endothelium) to form TXs or PGI<sub>2</sub>, respectively. Arachidonic acid can also be acted upon by 5-lipoxygenase, primarily in leukocytes, to form leukotrienes (LTs). One or more of these metabolites can mediate all the signs and symptome associated with arachidonic acid, i.e. inflammatory disease and pain.

Platelets, leukocytes, smooth muscle, and endothelium can produce vesoactive substances, products of arachidonic acid metabolism such as prostaglendins (PG9), prostacyclin (PGI<sub>2</sub>), leukotrienes (LTs), and thromboxanes (TXs). These substances can either act as vesocilators or as vesoconstrictors. PGI<sub>2</sub> is essential in vescular function since it inhibits platelet adhesion to the vescular endothelium and has significant vesodilators or guardies. Damaged endothelial cells cannot produce PGI<sub>2</sub>, making the vessel more susceptible to thrombosis and vesospasm. Thromboxanes and leukotrienes serve a vescular function during inflammation, generally producing vesoconstriction. Prostaglandins have a vescular role during inflammation, and also play a more subtle role in normal flow regulation, most notably as modulators of other control mechanisms. Prostaglandins have both vesoconstrictor and vesocilator activities. Leukotrienes and prostaglandins can also increase the endothelial membrane permeability thus promoting edema during inflammation. Arachidonic activities and is naturally present in most phospholipid mixtures or emulsions available today.

Nervonic acid (C24:1) is also called selacholeic acid or tertracosenic acid. Nervonic acid is the symbol of white matter in glucoside, which is quantitatively contained in nerve tissue and white matter. The absence of nervonic acid may result in cerebral lesion, fatigue, hypodynamia, amentia, and senile dementia.

DOCKET

----

Find authenticated court documents without watermarks at docketalarm.com.

Nervonic acid, tertracosenic acid in another name, is monounsaturated, non-oxidable/decomposed and absorptive. It is called a rare tonic as it is rare existent in nature, may be micro-obtained by compounded in cerebral chondriosome. Therefore, the substance is far below the demand of human body. In foreign countries, nervonic acid mainly comes from shark brain and oil.

Flavonoids are polyphanolic compounds ubiquitous in nature. They are categorized into isoflavonea, anthocyanidins, flavans, flavonola, flavones, citrus flavonoids, hesparidin, chalcones, catechins, rutin, and flavanones. Essential flavonoids, such as quercetin in onions and genistein in soy are actually considered subcategories rather than independent categories. Over 4,000 flavonoids have been identified in fruits, vegetables and beverages (tea, coffee, beer, wine and fruit drinks). Even though they have a similar molecular structure between them, their functions are different from each other. Flavonoids have been shown to have antibacterial, anti-Inflammatory, antiallergic, antimutagenic, antiviral, antineoplastic, anti-thrombotic, and vascellatory activity. Quercetin has been proven to block the "sorbitol pathway" which is directly associated with diabetes as well as to prevent LDL-cholesterol exidative damage, which is essential for the maintenance of a healthy cardiovascular system.

Flavonoids are found in a wide range of fruits and vegetables. For example, Quercetin (a flavonol in vegetables, fruit and onions), Xanthohumol (a prenylated chalcone in beer). Isoxanthohumol (a prenylated flavanone in beer), Genistein (an isoflavone in soy), Chalconaringenin (a non-prenylated chalcone in citrus fruits) and Naringenin (a non-prenylated flavanone in citrus fruits).

In plants flavonoids have very well defined functions. First, the accumulation of pigment in flower petals, seeds and leafs. Flowers, as pollinators, must attract pollen carriers. Second, they protect plants from UV damage, by absorbing UV at the epidermal layer. Third, they protect the plants against insects and pathogens.

The fiavonoid biosynthetic pathway is one of the best understood plant secondary metabolism pathways (1992, Gerats). The key enzymes are phenylalanine-ammonia lyase and chalcone synthase. Phenylalanine-ammonia lyase converts phenylalanine into cinnamic acid as it controls the total flow of carbons into phenolics which is shown to be the limiting step in this pathway (1974, Cressy). Another key enzyme of the flavonoid pathway is the chalcone synthase. It condenses three molecules of

DOCKET

RM

## DOCKET A L A R M



# Explore Litigation Insights

Docket Alarm provides insights to develop a more informed litigation strategy and the peace of mind of knowing you're on top of things.

## **Real-Time Litigation Alerts**



Keep your litigation team up-to-date with **real-time alerts** and advanced team management tools built for the enterprise, all while greatly reducing PACER spend.

Our comprehensive service means we can handle Federal, State, and Administrative courts across the country.

## **Advanced Docket Research**



With over 230 million records, Docket Alarm's cloud-native docket research platform finds what other services can't. Coverage includes Federal, State, plus PTAB, TTAB, ITC and NLRB decisions, all in one place.

Identify arguments that have been successful in the past with full text, pinpoint searching. Link to case law cited within any court document via Fastcase.

## **Analytics At Your Fingertips**



Learn what happened the last time a particular judge, opposing counsel or company faced cases similar to yours.

Advanced out-of-the-box PTAB and TTAB analytics are always at your fingertips.

## API

Docket Alarm offers a powerful API (application programming interface) to developers that want to integrate case filings into their apps.

#### LAW FIRMS

Build custom dashboards for your attorneys and clients with live data direct from the court.

Automate many repetitive legal tasks like conflict checks, document management, and marketing.

#### FINANCIAL INSTITUTIONS

Litigation and bankruptcy checks for companies and debtors.

### E-DISCOVERY AND LEGAL VENDORS

Sync your system to PACER to automate legal marketing.