

TRIDOCOSAHEXANOÍNA-AOX®

30 YEARS

PROVIDING
SCIENTIFIC INNOVATION
IN HEALTH CARE

1995 – 2025



BRUDYLAB®

About us

BRUDY is a pioneering biotechnology company in Medical Nutrition. We believe in high-quality clinical research and are driven by our innovative spirit in the field of omega-3 fatty acids, particularly Docosahexaenoic Acid (DHA) and its impact on human health.

Our goal is to improve the quality of life for everyone and contribute to the treatment of diseases associated with oxidative stress and inflammation.



Our track record

BRUDY has been committed to researching and developing innovative DHA-based products since 1995. Our multidisciplinary team focuses on creating top-quality products. Our approach merges science and state-of-the-art techniques, providing tailored solutions with an ongoing commitment to excellence and innovation.

We have invested more than €10 million in research projects throughout our history and hold three international patents¹⁻³ on the use of our DHA-enriched triglyceride, Tridocosahecanoína-AOX®.

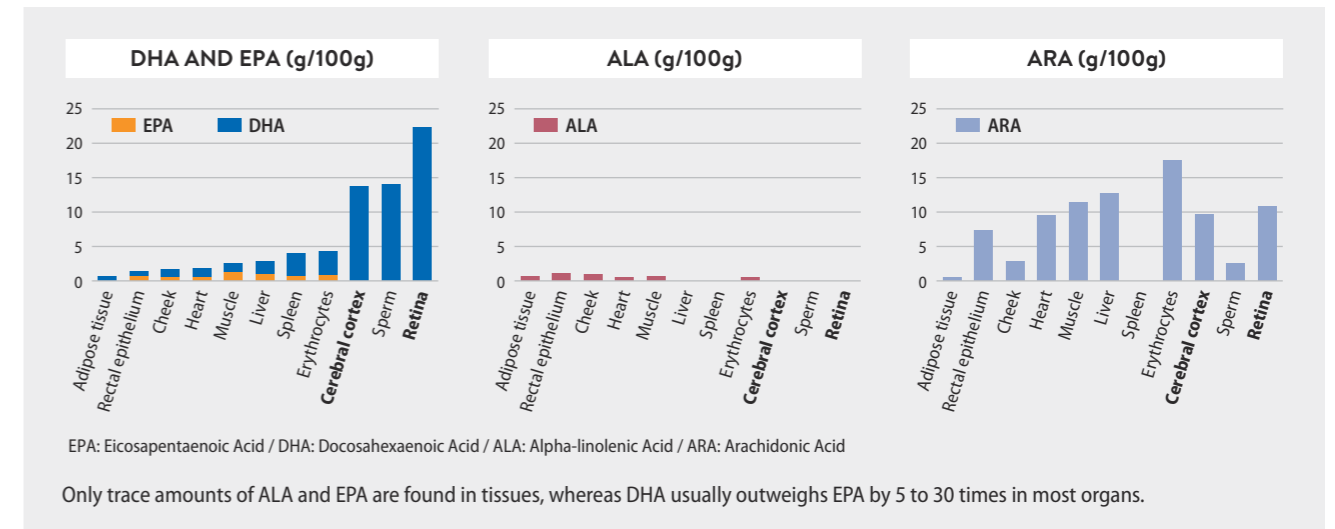
We meet the highest quality standards and have repeatedly provided optimal bioactivity, hence positioning us as global leaders in this field.

BRUDYLAB has focused on marketing our proprietary products with Tridocosahecanoína-AOX® as the main ingredient in food supplements, food for special medical purposes as well as cosmetic products on a national and international level since 2010.

DHA – an essential omega-3 for health

DHA is a very long-chain (C22:6) polyunsaturated fatty acid, an essential omega-3 for cell membrane structure and function.

Fatty acid concentration found in human tissues⁴



Health benefits of DHA acknowledged by the European Commission⁵

HEALTH BENEFITS OF DHA APPROVED BY THE EUROPEAN COMMISSION	CONSUMERS SHOULD BE INFORMED THAT BENEFITS ARE OBTAINED WITH THE INTAKE OF:
DHA (and DHA + EPA) contributes to normal heart function.	250 mg/day of DHA (EU 432/2012). For adults. EFSA Journal 2014;12(10):3840. For children aged 2 to 18.
DHA contributes to normal vision.	
DHA contributes to normal brain function.	
DHA contributes to normal visual development in children up to 12 months.	100 mg/day of DHA (EU 440/2011).
Maternal intake of DHA contributes to normal visual development of the foetus and of breastfed infants.	(Pregnant and breast-feeding women) 200 mg/day of DHA in addition to the 250 mg/day daily intake for adults (EU 440/2011).
Maternal intake of DHA contributes to normal brain development of the foetus and of breastfed infants.	
DHA (or DHA + EPA) helps maintain normal triglyceride levels in the blood.	2 g/day of DHA or DHA + EPA. Do not exceed 5 g/day of combined intake (EU 536/2013).
DHA and EPA help reduce blood pressure.	3 g/day of DHA or DHA + EPA. Do not exceed 5 g/day of combined intake (EU 536/2013).

EU Regulatory Consultations at: www.eur-lex.europa.eu (search by year and regulation no. mentioned in the right-hand table column)

BRUDY worldwide

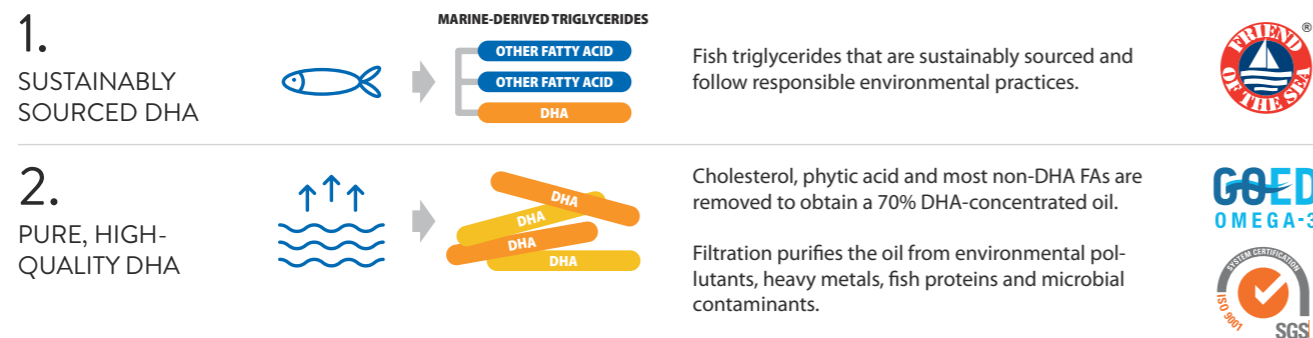
● BRUDYLAB ● BRUDYTECHNOLOGY



What is Tridocosahexanoína-AOX®?

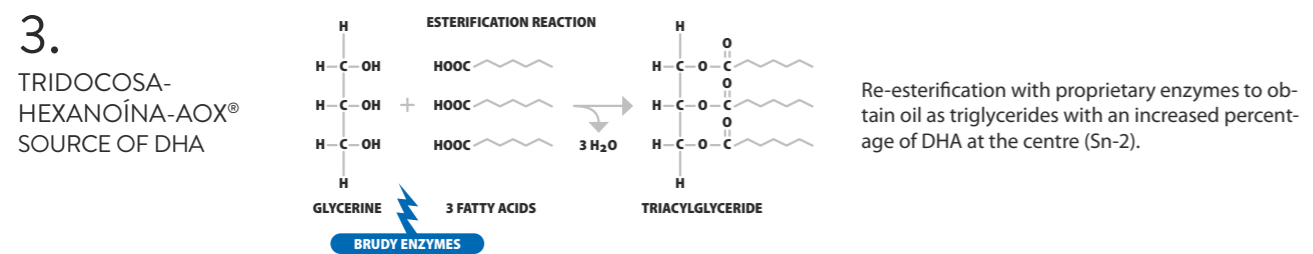
A DHA-enriched source developed by BRUDY as highly purified and structured re-esterified triglyceride to ensure bioavailability and biological efficacy.

Tridocosahexanoína-AOX® synthesis in 3 steps:

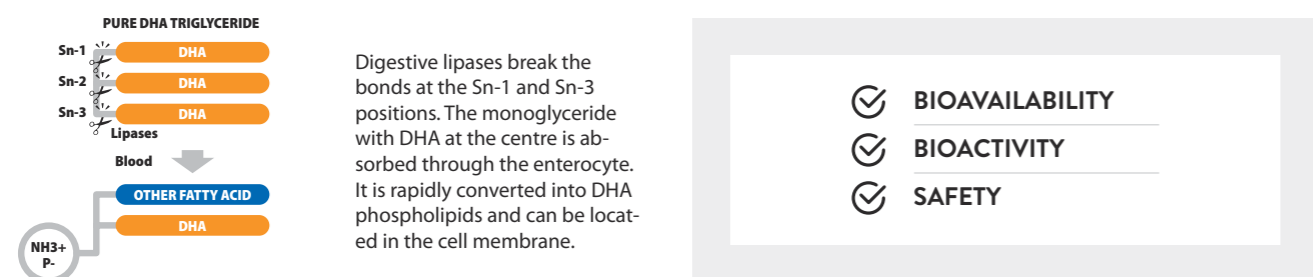


	European legislation ⁶	GOED	IFOS	Brudy's batch*
Purity, safety and cleanliness				
PCBs	200 ng/g	90 ng/g	<=45 ppb	<15 ng/g
Dioxins	1.75 pg/g	1.75 pg/g	<= 1 ppt	<0.4 pg/g
Dioxins and similar PCBs	6 pg/g	3 pg/g	<= 1.5 ppt	<0.5 pg/g
Heavy metals				
Mercury (Hg)	0.5 ppm	0.1 ppm	<= 0.1 ppm	<0.005 ppm
Lead (Pb)	0.1 ppm	0.05 ppm	<= 0.1 ppm	<0.02 ppm
Arsenic (As)	-	0.1 ppm	<= 0.1 ppm	<0.05 ppm
Cadmium (Cd)	0.1 ppm	0.1 ppm	<= 0.1 ppm	<0.05 ppm
Stability				
Acid value	-	-	<=3 mg KOH/g	0.1 mg KOH/g
Peroxide	-	-	<=5 meq/kg	2-4 meq O2/kg
Anisidine	-	-	<=20	4-10 U.A.

*Each batch of oil is analysed by Eurofins Analytics



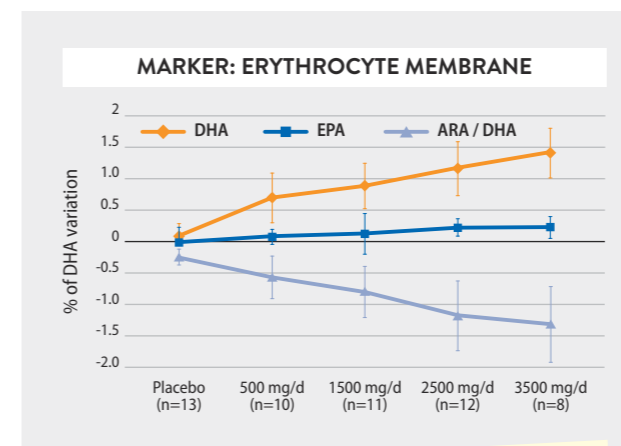
What happens in the body?



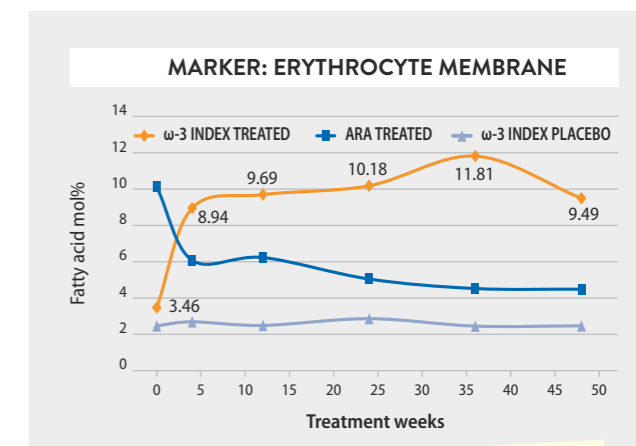
Proven short- and long-term bioavailability^{7, 8}

- ✓ In the short term, supplementation with Tridocosahexanoína-AOX® increases DHA levels in cell membranes, suggesting **efficient absorption** and **quick incorporation** into tissues.
- ✓ In the long term, ongoing supplementation keeps DHA levels high.

- ✓ From the outset, a dependent relationship between increased DHA and reduced arachidonic acid (ARA) levels was observed.



• n = 54 healthy volunteers
 • 5 groups: placebo and increasing doses of Tridocosahexanoína-AOX®
 • Duration = 1 month



• n = 73 HIV+ patients
 • Control group (n = 38) without supplementation
 • Experimental group (n = 35) with 4 g/day of Tridocosahexanoína-AOX®
 • Duration = 1 year

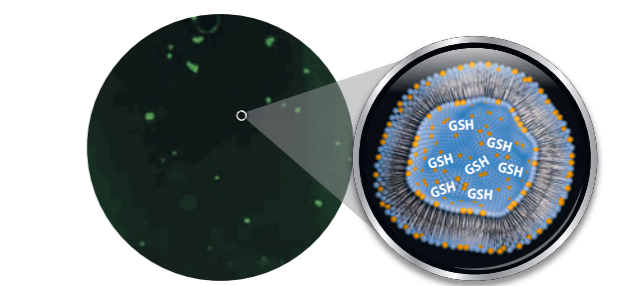
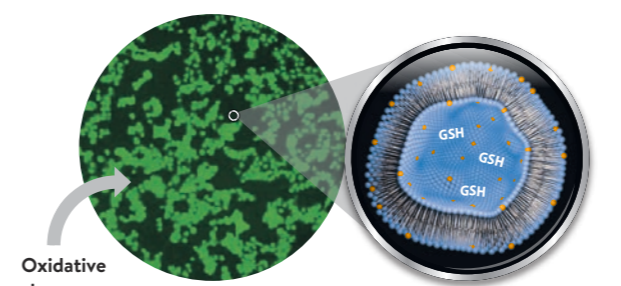
Patented antioxidant activity of Tridocosahexanoína-AOX®

Our analyses of human cell cultures revealed that, with Tridocosahexanoína-AOX®, cell membranes **increase glutathione synthesis by 200% to 300%**⁹. This powerful antioxidant, which is essential for regulating internal cellular balance, protects cells from oxidative stress by

neutralising free radicals that can damage cells and cause cell death. Glutathione protects cellular health, and its increase, which is stimulated by Tridocosahexanoína-AOX®, prepares cells to withstand external aggressions.

ARPE-19 RETINAL CELLS CONTROL GROUP CELL CULTURE

ARPE-19 RETINAL CELLS BRUDY TG-DHA CELL CULTURE



GSH: Glutathione

ARA: Arachidonic Acid Omega 6

DHA: Docosahexaenoic Acid Omega 3

↑ 200-300% increase of glutathione

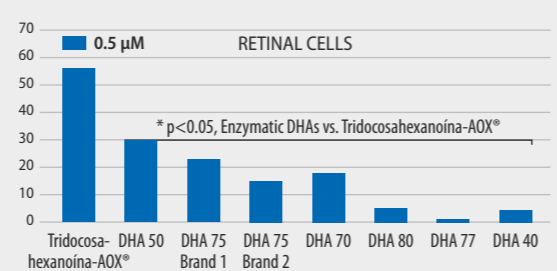
↑ Increase in endogenous antioxidant defence capacity

Bioactivity – our main Gold standard

At BRUDY, we produce clean, pure DHA as bioavailable triglyceride with **bioactivity that mimics human physiology**. Our production process preserves the natural DHA structure, while maintaining the cis double bonds that provide its antioxidant capacity and benefits for the body. Each batch of oil undergoes cellular bioactivity testing to ensure its **antioxidant protection activity (50-60%)**. Unlike other processes, our technology ensures that bioactivity does not only depend on DHA concentration, but also on the preservation of its structure, from oil extraction to the end product.



ANTIOXIDANT EFFECT OF TRIGLYCERIDES WITH VARYING DHA CONCENTRATIONS



Consolidated group "Drug Transportation and Delivery" (2009 SGR-367). Department of Biochemistry and Molecular Biology, University of Barcelona. Reproduction of graphs is prohibited © All rights reserved.

Clinical research

At BRUDY, our mission is to back up the benefits of our products with scientific evidence, while always seeking to provide innovative solutions based on current needs. Thus,

we work with national and international experts in several areas, such as ophthalmology, fertility, sports medicine or lipid metabolism disorders.

Evidence of Tridocosa-hexanoína-AOX® in other therapeutic areas

CHILD CARE



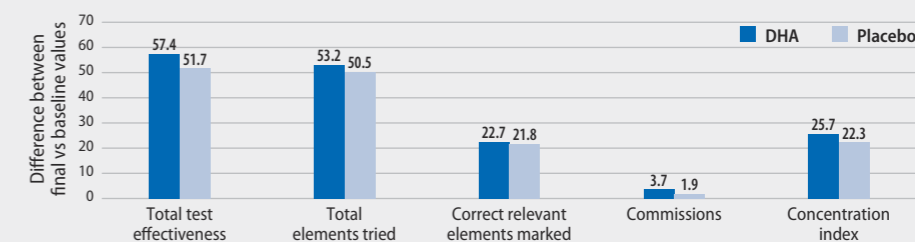
3 studies

- ✓ Improvement in cognitive and behavioural variables supporting its benefit in children and adolescents with attention deficit hyperactivity disorder (ADHD)
- ✓ Improvements in lung function (FVC and FEV1) and decrease in the number of exacerbations in children with cystic fibrosis

Child care¹

- N = 66 patients aged 6 to 18 with ADHD
- 1000-2000 mg/day of Tridocosa-hexanoína-AOX® (according to weight)
- Duration = 6 months

DIFFERENCE BETWEEN FINAL VS BASELINE VALUES



42 clinical studies + 5 review articles

Eye care

19 studies

3 review articles



9 studies on Dry eye*



5 studies Retina + review articles



4 studies on Glaucoma*



2 studies on Eye inflammation



1 study Topical effect + review article**

Other therapeutic areas

23 studies

2 review articles



6 studies Sports



3 studies Lipid metabolism disorder + review articles



4 studies Fertility



4 studies Children with ADHD



2 studies Cystic fibrosis



3 studies Neurological disorder



4 studies In vitro techniques

*Categories shared by one or more studies.
**Accepted article pending publication

DHA triglyceride concentration
70%

Antioxidant protection
50-60%

Glutathione increase
200-300%

100% bioactivity in each batch

3 patents for Tridocosa-hexanoína-AOX® use

FERTILITY



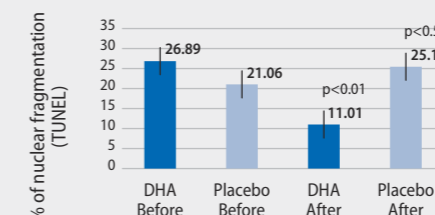
4 studies

- ✓ Increased antioxidant capacity
- ✓ Decreased DNA fragmentation
- ✓ Increased sperm motility

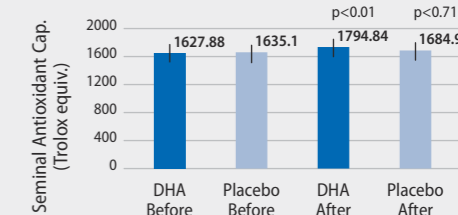
Fertility⁴

- N = 64 subjects undergoing fertility assessment
- Placebo group = 29
- Supplemented group = 35 (1050 mg/day of Tridocosa-hexanoína-AOX®)
- Duration = 10 weeks

DNA FRAGMENTATION



ANTIOXIDANT CAPACITY



Before and 10 weeks after supplement intake

SPORTS

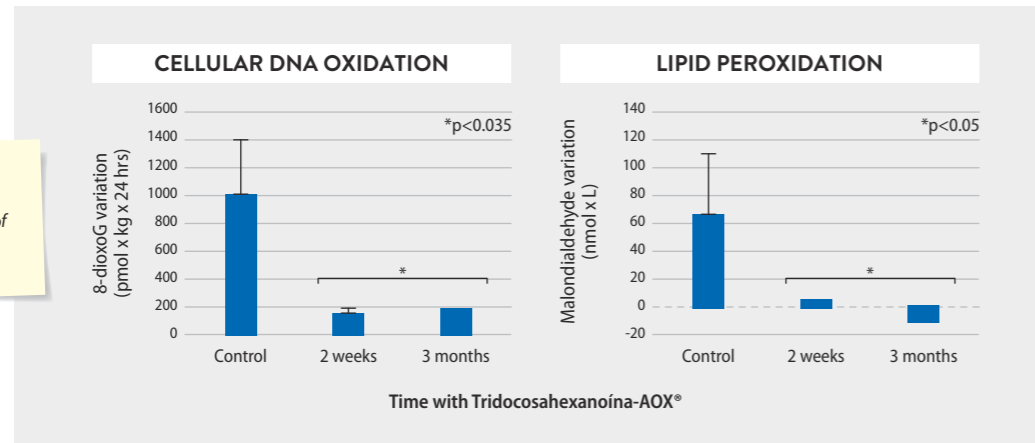


6 studies

- ✔ Significant protection against oxidative damage resulting from moderate-intense exercise
- ✔ Improved perceptual-motor reaction time
- ✔ It promotes low inflammation concentrations and muscle damage markers, while also decreasing muscle pain

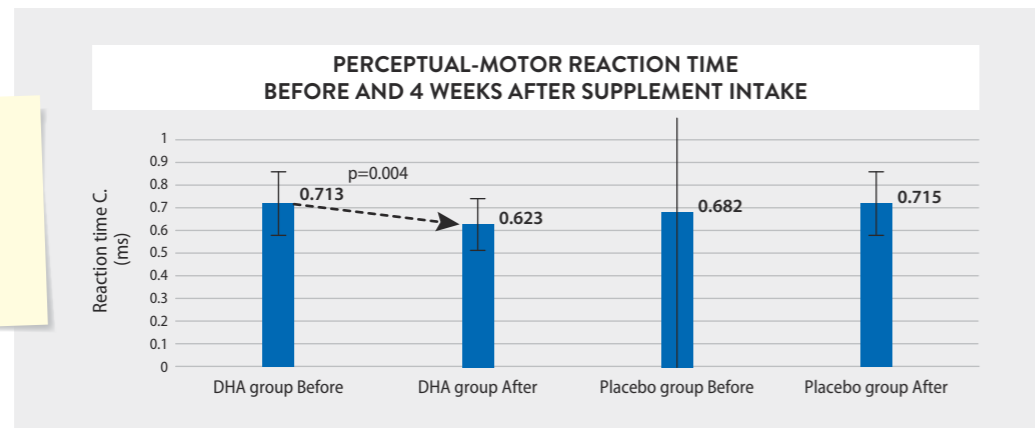
Sports⁶

- n = 40 healthy sportspeople
- Treatment with 2000 mg/day of Tridocosahexanoína-AOX[®]
- Duration = 3 months



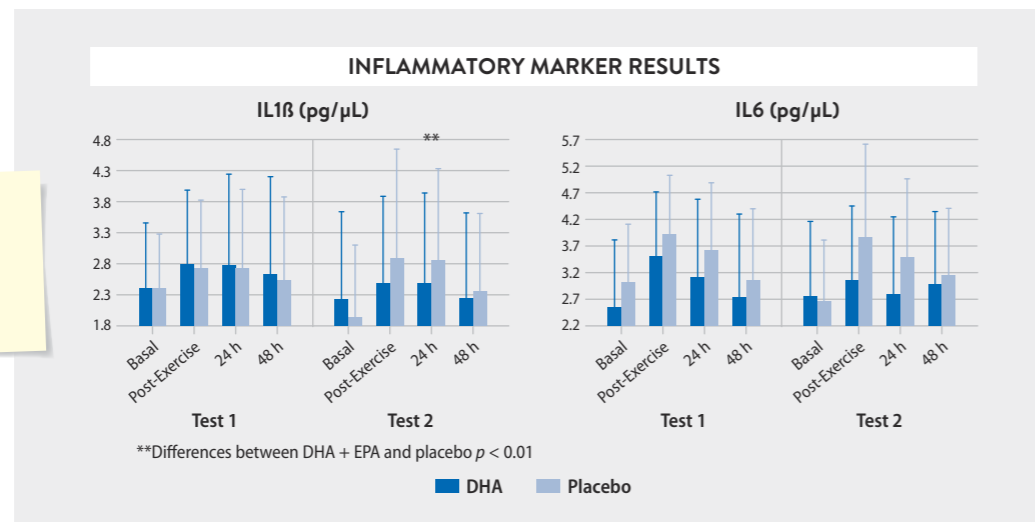
Sports¹

- N = 24 professional female football players
- Control group n = 12 placebo
- Supplemented group n = 12 3500 mg/day of Tridocosahexanoína-AOX[®]
- Duration = 4 weeks



Sports⁴

- n = 15 triathletes
- Supplemented group with 2000 mg/day of Tridocosahexanoína-AOX[®]
- Duration = 10 weeks



LIPID METABOLISM



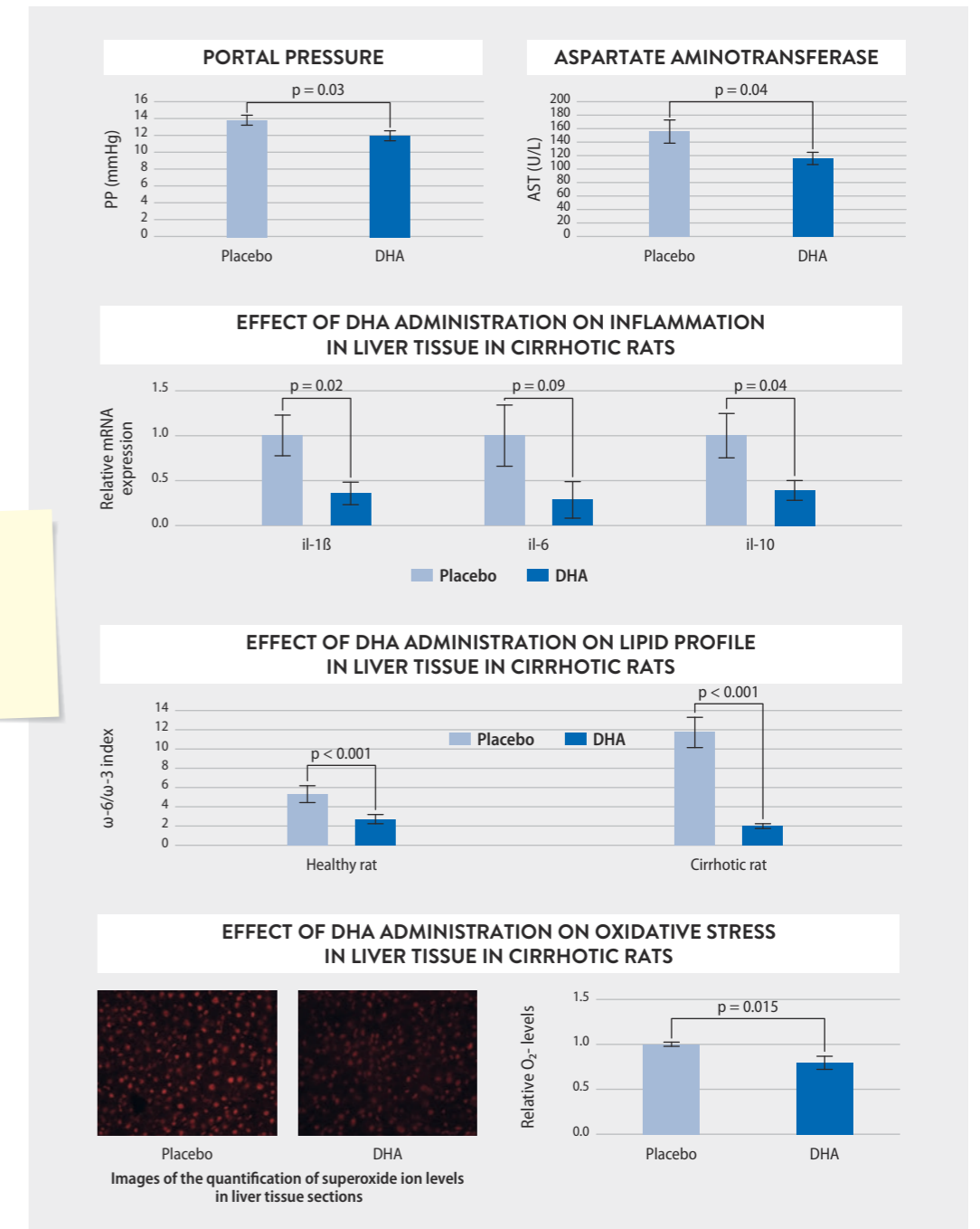
3 studies

2 review articles

- ✔ Correction of intrahepatic omega-6/omega-3 imbalance
- ✔ Reduction of intrahepatic oxidative stress: -20.7%
- ✔ Reduction in the expression level of circulating inflammatory markers: IL-1β: -65% | IL-10: -61% | IL-6: -71.63%
- ✔ Deactivation of mouse and human hepatic stellate cell and slowing down of the fibrotic process
- ✔ Significant reduction in portal pressure: -13.4%
- ✔ Reduction in circulating aspartate aminotransferase: -25.7%

Lipid metabolism³

- n = 12 healthy rats and 25 cirrhotic rats
- ½ with placebo and ½ supplemented with Tridocosahexanoína-AOX[®] (500 mg/kg)
- Duration = 2 weeks



Evidence of Tridocosahecanoína-AOX® in eye care

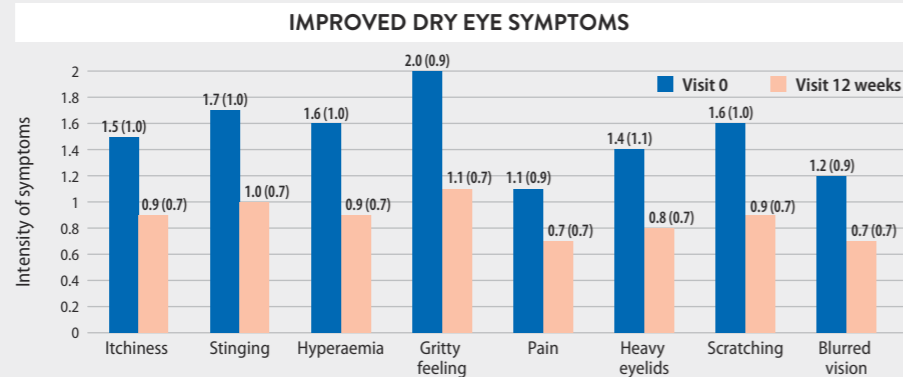
DRY EYE SYNDROME



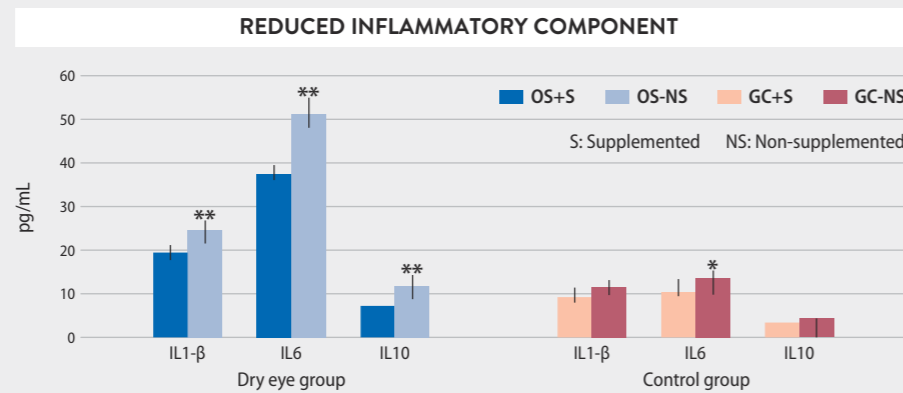
9 studies

- Improved dry eye symptoms
- Reduced inflammation
- Improved tear quantity and quality

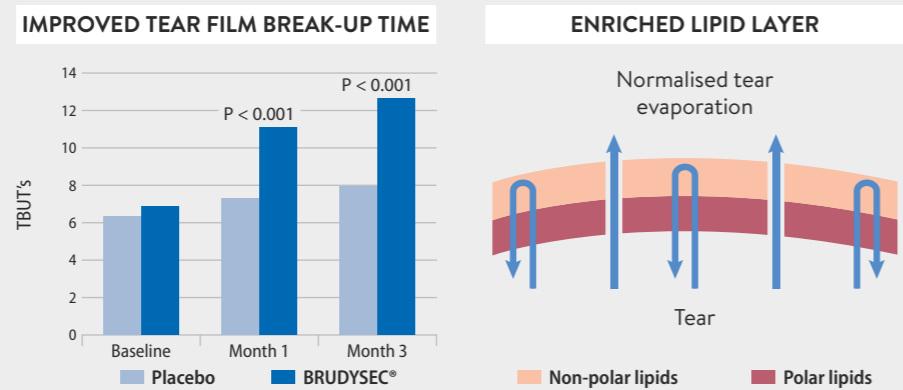
Dry eye disease¹
 • n = 905 patients with dry eye symptoms
 • 1050 mg/day of Tridocosahecanoína-AOX®
 • Duration = 3 months



Dry eye disease¹
 • n = 66 patients (132 eyes)
 • 1050 mg/day of Tridocosahecanoína-AOX®
 • Mild-moderate dry eye group > 1 year (n = 30)
 • ½ suppl. – ½ not suppl.
 • Healthy control group (n = 36)
 • ½ suppl. – ½ not suppl.
 • Duration = 3 months



Dry eye disease³
 • n = 61 patients with meibomian gland dysfunction (MGD)
 • Control group (n = 31) with placebo
 • Experimental group (n = 30) 1050 mg/day of Tridocosahecanoína-AOX®
 • Duration = 3 months



RETINAL/MACULAR DISORDERS

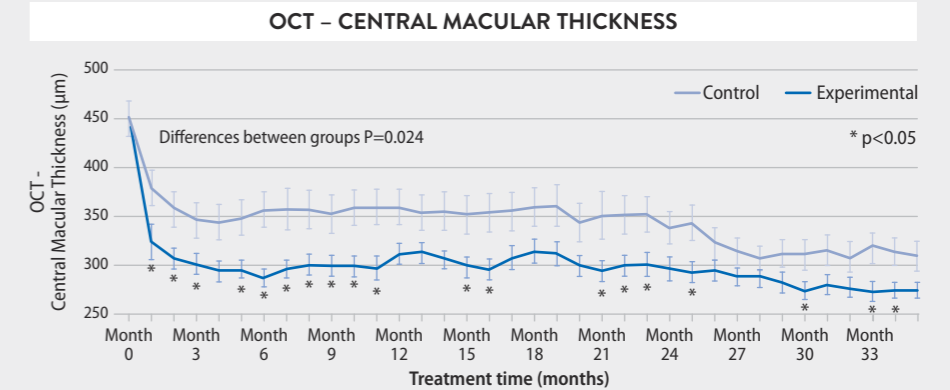


5 studies

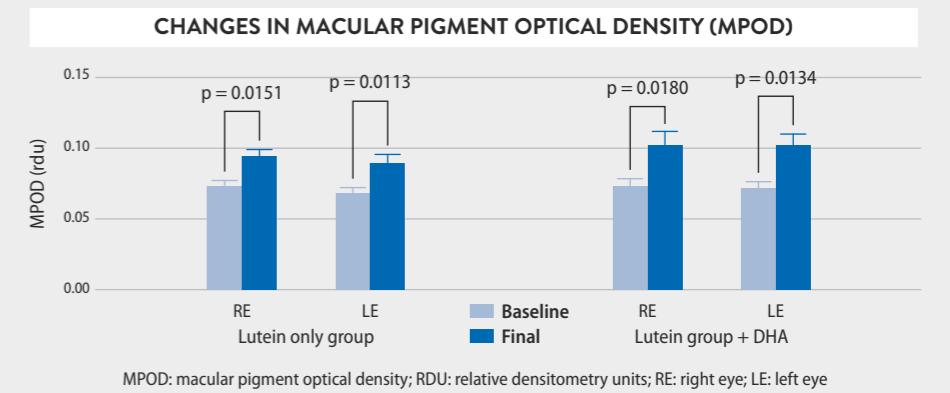
2 review articles

- Anti-oedematous effect in addition to conventional treatment of Age-related Macular Degeneration (AMD)
- Significant increase in Macular Pigment Optical Density (MPOD)
- Improved macular sensitivity and macular integrity assessment in Non-Proliferative Diabetic Retinopathy (NPDR)

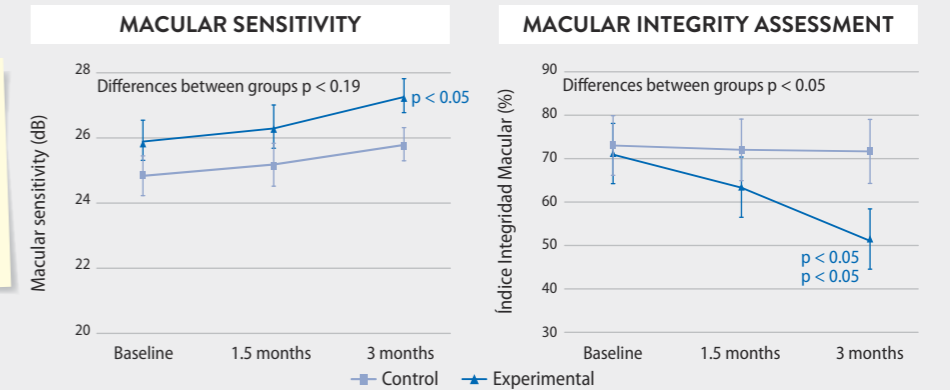
Retinal/macular disorders³
 • n = 77 AMD patients treated with ranibizumab IV
 • Control group (n = 29) without supplementation
 • Experimental group (n = 26) 1050 mg/day of Tridocosahecanoína-AOX®
 • Duration = 36 months



Retinal/macular disorders⁴
 • n = 100 healthy volunteers (200 eyes)
 • Control group (n = 49) Placebo with lutein
 • Experimental group (n = 51) 1050 mg/day of Tridocosahecanoína-AOX®
 • Duration = 3 months



Retinal/macular disorders²
 • n = 24 patients with NPDR
 • Control group (n = 12) without supplement
 • Experimental group (n = 12) 1050 mg/day of Tridocosahecanoína-AOX®
 • Duration = 3 months



GLAUCOMA



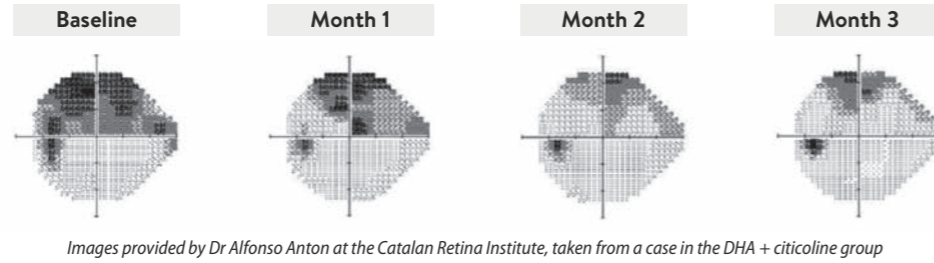
4 studies

- Improved visual field indices and progression rate with Tridocosahexanoína-AOX® + citicoline combined
- Improved dry eye symptoms and decrease in lacrimal gland inflammation
- Antioxidant and inflammation-regulating effects as mechanisms related to glaucoma



Glaucoma⁴

- n = 73 glaucoma patients with controlled IOP
- Control group (n = 17) vitamin C
- Group A (n = 16) 1050 mg/day of Tridocosahexanoína-AOX®
- Group B (n = 20) 326 mg of citicoline
- Group C (n = 20) 1050 mg/day of Tridocosahexanoína-AOX® + 326 mg of citicoline
- Duration = 3 months



VISUAL FIELD PARAMETERS AT BASELINE AND AFTER 3 MONTHS OF TREATMENT

	Baseline MD (dB)	Month 3 MD (dB)	p Value	Baseline VFI (%)	Month 3 VFI (%)	p Value
Global (mean ± SD)	-8,96 ± 3,91	-8,42 ± 4,29	0,025	77,47 ± 12,83	78,52 ± 13,94	0,096
Vitamin C (mean ± SD)	-8,46 ± 3,70	-8,32 ± 3,87	0,727	78,35 ± 12,97	78,53 ± 13,24	0,871
DHA (mean ± SD)	-9,08 ± 4,07	-8,86 ± 5,07	0,685	78,13 ± 12,83	77,56 ± 15,72	0,719
Citicoline (mean ± SD)	-8,74 ± 3,70	-8,71 ± 4,17	0,957	78,50 ± 12,46	78,90 ± 13,16	0,687
DHA+Citicoline (mean ± SD)	-9,52 ± 4,36	-7,85 ± 4,36	0,001	75,15 ± 13,76	78,90 ± 14,82	0,008

COMPARISON BETWEEN PRE- AND POST-TREATMENT VISUAL FIELD INDEX SLOPES IN EACH GROUP

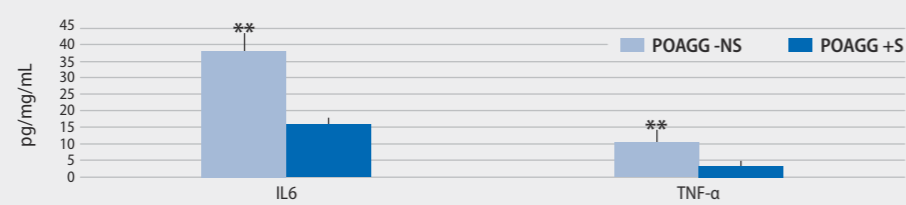
	MD Slopes (dB/Month)		p Value	VFI Slopes (%/Month)		p Value
	Pretreatment	Post-Treatment		Pretreatment	Post-Treatment	
Global (mean ± SD)	-0,0613 ± 0,1736	0,0867 ± 0,3092	0,005	-0,1107 ± 0,2781	0,1625 ± 0,8499	0,018
Vitamin C (mean ± SD)	-0,0502 ± 0,1459	0,055 ± 0,2895	0,350	-0,079 ± 0,3665	0,1425 ± 0,8992	0,485
DHA (mean ± SD)	-0,0135 ± 0,1116	0,0295 ± 0,1939	0,733	-0,0509 ± 0,1998	0,0150 ± 0,5971	0,532
Citicoline (mean ± SD)	-0,0624 ± 0,1307	0,1029 ± 0,4504	0,192	-0,1364 ± 0,3151	0,1733 ± 1,2439	0,371
DHA+Citicoline (mean ± SD)	-0,1041 ± 0,2471	0,1383 ± 0,2544	0,006	-0,1557 ± 0,2310	0,2780 ± 0,5661	0,006

MD, mean defect of visual field; dB, decibels; VFI, visual field index; SD, standard deviation; DHA, docosahexanoic acid.

Glaucoma¹

- n = 97 patients suffering from dry eye
- Group 1 – Mild-moderate dry eye (n = 30) ½ suppl. – ½ not suppl.
- Group 2 – Primary Open-Angle Glaucoma (POAG) (n = 31) ½ suppl. – ½ not suppl.
- Group 3 – Healthy controls (n = 36) ½ suppl. – ½ not suppl.
- 700 mg/day of Tridocosahexanoína-AOX®
- Duration = 3 months

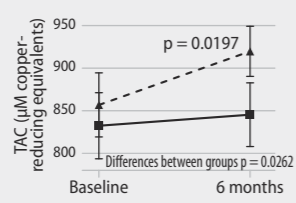
IL-6 AND TNF-α IN TEAR SAMPLES FROM THE GLAUCOMA GROUP



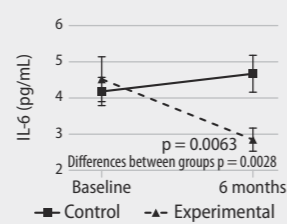
Glaucoma³

- n = 47 patients suffering from pseudoexfoliative glaucoma with controlled IOP
- Control group (n = 24) without supplement
- Experimental group (n = 23) 1050 mg/day of Tridocosahexanoína-AOX®
- Duration = 6 months

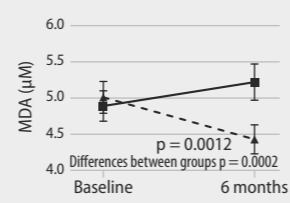
CHANGES IN TAC



CHANGES IN IL-6



DECREASE IN MDA



EYE INFLAMMATION



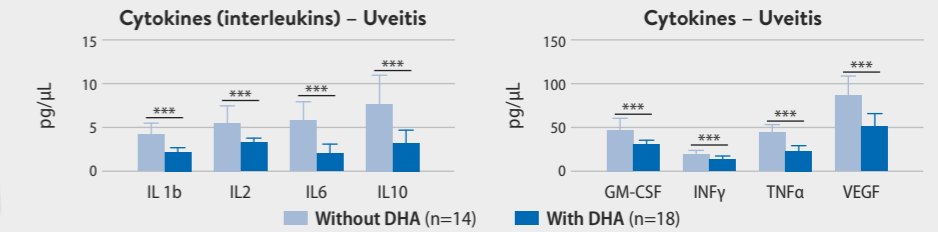
2 studies

- Reduced systemic inflammatory burden
- Antioxidant and anti-inflammatory effect to address the underlying pathophysiological mechanisms of keratoconus

Eye inflammation¹

- n = 62 patients
- Group 1 (n = 30) healthy volunteers ½ suppl. – ½ not suppl.
- Group 2 (n = 32) NIU in remission phase ½ suppl. – ½ not suppl.
- 500 mg/day of Tridocosahexanoína-AOX®
- Duration = 3 months

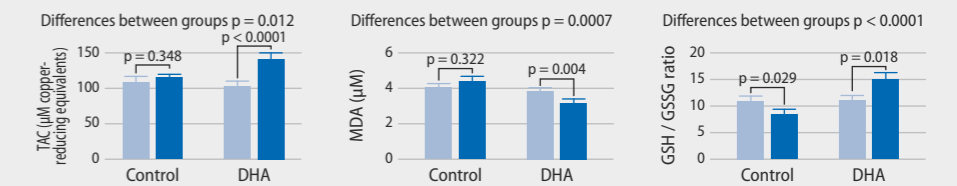
COMPARISON BETWEEN SUPPLEMENTED NIAU PATIENTS AND UNSUPPLEMENTED NIAU PATIENTS



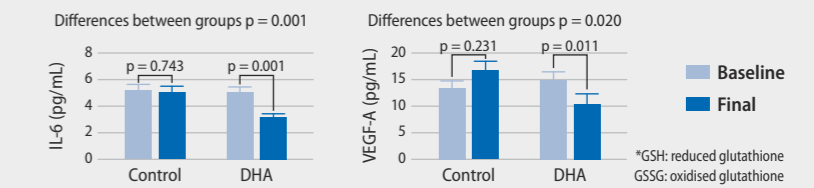
Eye inflammation²

- n = 34 patients with keratoconus
- Control group n = 15 without supplement
- Supplemented group n = 19 1000 mg/day of Tridocosahexanoína-AOX®
- Duration = 3 months

PLASMA ANTIOXIDANT PROTECTION



PLASMA INFLAMMATORY MARKERS



TOPICAL TREATMENT



1 study

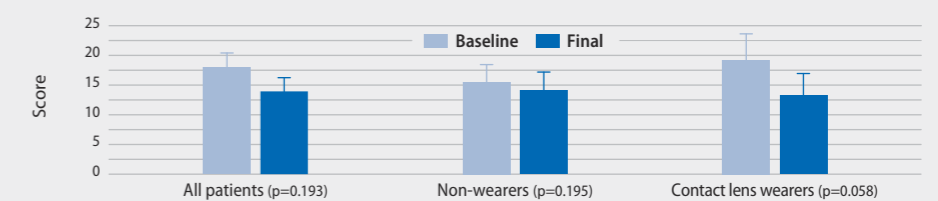
1 review article

- Clinically proven efficiency and safety of cream with Tridocosahexanoína-AOX®

Topical effect¹

- n = 60 healthy volunteers (30 contact lens wearers)
- 1 application of cream on the eyelid skin at night
- Duration = 2 weeks

OSDI – VALIDATION OF THE SYMPTOM QUESTIONNAIRE (Ocular Surface Disease Index)



References

1. Patent WO 2006120120 A1: Use of docosahexaenoic glycerides for the treatment of tumorous diseases.
2. Patent WO 2004050077 A1: Use of docosahexaenoic acid as active substance for the treatment of lipodystrophy .
3. Patent WO 2007/ 071733 A3: Use of DHA, EPA or DHA-derived EPA for treating a pathology associated with cellular oxidative damage.
4. Linda M Arterburn, et al; Distribution, interconversion, and dose response of ω-3 fatty acids in humans; *Am J Clin Nutr* 2006;83(suppl):1467S–76S.
5. List of authorised health claims for docosahexaenoic acid (DHA), as suggested by the EFSA (European Food Safety Authority). REGULATION (EU) no. 432/2012 and REGULATION (EU) no. 440/2011.
6. EC Regulation no. 1881/2006 (19 December 2006) setting maximum levels of contaminants in foodstuffs and amendment no. 1259/2011.
7. Carlos J Contreras; Modification of the oxidative damage in a group of cyclists after consuming docosahexaenoic acid at different doses; Doctoral Thesis, Catholic University of Murcia, 2014.
8. Domingo P, et al., Effects of docosahexanoic acid on metabolic and fat parameters in HIV-infected patients on cART: A randomized, double-blind, placebo-controlled study, *Clinical Nutrition* (2017), <http://dx.doi.org/10.1016/j.clnu.2017.05.032>.
9. P. Bogdanov, et al; Docosahexaenoic Acid Improves Endogen Antioxidant Defense in Arpe-19 Cells; *IOVS, ARVO Journals*; May 2008, Vol.49, 5932.

DRY EYE SYNDROME (DES)

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