The Forum For Integrative Medicine

April 3-6, 2025 AUSTIN, TX **Cracking The Code of Chronic Illness**

Utilizing the D6D Anti-Inflammatory Pathway to

Professor Brian Scott Peskin* Awarded 3 plant-based oil patents. Consults with numerous nutritional companies **Founder: Peskin Pharmaceuticals**

*Brian Peskin earned his Bachelor of Science degree in Electrical Engineering from the Massachusetts Institute of Technology — BSEE. He received an appointment as an Adjunct Professor at Texas Southern University in the Department of Pharmacy and Health Sciences (1998-1999). The former president of the University said of Brian's discoveries: "...His nutritional discoveries and practical applications through Life-Systems Engineering are unprecedented."

Plant-Based Seed Oil EFAs & Their Derivatives — The Key to Decreasing Patients' Chronic Inflammation

Healing Progression in Crushing Fall

86-Year-Old Patient -- Multiple Metabolic Pathways positively affected by plant-based Essential Fatty Acids (EFAs) and their long-chain metabolites, responsible for this "remarkable" healing progression via the D6D pathway.

"This 86-year-old patient experienced a significant bilateral orbitofrontal and nasal crushing injury in a fall. She incurred the anticipated bilateral upper and mid-facial swelling and ecchymosis but <u>surprisingly</u> had much less nasal injury than expected, including no fractured/comminuted nasal bones or orbital rim/floor fracture. Within 10 days, she demonstrated remarkable resolution of the facial edema and bruising and quickly thereafter eliminated the remainder over the ensuing weeks. So not only did it seem that her facial soft tissues and bones were more resilient and the bones also not brittle, but her overall recovery was unusually expedited and uncomplicated."

Geoffrey L. Robb, M.D., F.A.C.S., Board Certified ENT Professor / Past Chairman (1998-2013) Department of Plastic Surgery, The University of Texas MD Anderson Cancer Center, Houston, TX

Organic / Unprocessed Omega-6 Seed Oils = ANTI-**INFLAMMATION**

2 months to

heal



February 7 24 days after fall.



February 28 46 days after fall.



February 13

31 days after fall.





January 30

17 days after fall.

February 21 39 days after fall.





January 23 10 days after fall.

Plant-Based Seed Oil EFAs & Their Derivatives — The Key to Decreasing Patients' Chronic Inflammation

Expedited Severe Wound Healing: **Toe SAVED / Venous Ulcer Heal** With Organic / Unprocessed Omega-6 **Seed Oils**





Feb. 4, 2021

April 9, 2021

 57-year-old white male presents with ischemic eschar of the 2nd right digit and eschar of the anterior distal hallux with pain on palpation.

 Told by the vascular service that it would likely demarcate and he would lose the digit. I treated him with betadine and EZTREK[™] 5cc, 5-days-a-week. Healing time was 9 weeks.

Jan. 29, 2021 April 29, 2021 (Both images depict 2nd occurrence)

- of varicose veins and severe COPD. Patient has had two separate episodes with an open ulcer.
- compression, and it closed in 16 weeks.
- compression and 5cc of EZTREK[™] 5 days a week.

CASE #3



• 78-year-old male with a right ankle open venous ulcer. History

First treatment, ulcer was managed with foam and three-ply

This time (2nd occurrence) treated with foam, three ply

Diseases, Disorders & Impairment of the Δ-6 Desaturase Pathway Causing Chronic Inflammation

- Diabetes (Type 1 & Type 2) including Neuropathy
- Lipid-Enveloped Viruses (including COVID series)
- Dermatological Conditions (Eczema, etc.)
- □ Cardiovascular Disease (Soft plaque/ Calcified plaque, Hypertension, etc.)
- Inflammatory Bowel Disease
- □ Chronic Fatigue (including post-viral syndromes)
- □ Fatty Liver Disease, Including NAFLD
- Multiple Sclerosis (MS)
- Dementia / Alzheimer's
- □ Cancer

Respiratory Diseases With Chronic Inflammation

- Mesothelioma
- Idiopathic Pulmonary Fibrosis (IPF)
- Chronic Obstructive Pulmonary Disease (COPD)

Request Fully Referenced Paper: prof-peskin@peskinpharma.com

Inflammatory Markers:

 C-reactive protein #1 (cancer patient) – 12/06/2021 "....3 high-sensitivity CRP tests conducted since June [2021]. June value was 1.4, mid-September was 1.8, and late November was 0.7. The 0.7 value [61% decrease] represents approximately 3 ½ weeks of EZtrek[®] use."

- <u>C-reactive protein</u> #2 (CHF patient) 03/09/2022 "History of CHF, likely induced from Lyme, etc. Contributing is MTHFR. High-sensitivity CPR test went from 1.5 to 1.1. The 1.1 value [27%] *decrease*] represents 4 weeks of **EZtrek**[®] use."
 - Homocysteine 03/09/2022 (CHF patient #2 above) 03/09/2022 "Homocysteine decreased from 13 to 9 [**31%** decrease] after 4 weeks of **EZtrek**[®] use.

Pulmonology:

"Pulmonologist was guarded on any improvement, stating, 'Your case is grim.' She has labored breathing and required 24-hour oxygen. On Feb 28, 2022, she saw Pulmonologist and had a **CT and PFT evaluation of lungs** that showed obvious positive changes. (She has only been on EZtrek[®] for one month.) Her pulmonologist was pleasantly surprised."

> "She no longer requires oxygen and can now speak and move about without being winded. Nothing else changed other than adding Plantbased seed oil Medical Food EZtrek[®]."

Hypertensives / 1st Reported SIGNIFICANT CAC REVERSAL

- 03-20-2023: "20-yr hypertensive; Norvac and Hydrochlorothiazide. BP reduced to **138/90**. When a) EZtrek® was added, after 3 months BP reduced to 126/75."
- b) 03-29-2022 "Blood pressure consistently raising until approx. 178/93. Low salt diet stabilized BP @ 160/85. Patient still concerned.

Within **30 days of EZtrek[®] BP lowered near 126/69**, which is consistently measured. **Patient is** delighted."

□ (Unprecedented -- 2022) Coronary Artery Calcified hard plaque reduction (CAC) "Baseline non-contrast CT Agatston Score of 727 on 4/30/2021: Score reduced to 666 on 6/30/2022 [7 months taking EZtrek[®] This translates to an annualized calcified plaque reduction of 14.3%." [NEVER > 5% CAC reduction.]

Inflammation NOW known to be at FOREFRONT of Diseases

Cancer

2007: Inflammation FUELS Cancer^{*} "<u>Re-writing of the textbook</u> ..."

- Cancer researcher Robert Weinberg of MIT states [coined "oncogene."]: "The connection inflammation and cancer has moved to center stage in the research arena." The article continues...
- "...inflammation is the fuel that feeds it [the malignant cancer].

• "In this **rewriting of the textbook**... This new view implies that rooting out every last cancer cell in the body might not be necessary. Anti-inflammatory cancer therapy instead would prevent pre-malignant cells from turning fully cancerous or would impede an existing tumor from spreading to distant sites in the body. Cancer victims might then be able to survive."

between

Inflammation: **Dementia / Alzheimer's Example**

Medical **X**press

New leads on treating dementia and Alzheimer's

May 2 2018, by Robyn Mills



PET scan of a human brain with Alzheimer's disease. Credit: public domain

A new study by scientists in Australia and the US provides an explanation for why clinical trials of drugs targeting proteins in the brain that were thought to cause dementia and Alzheimer's have failed. The

2018: "Inflammation is a MAJOR CAUSE, not just a consequence....but only now is it identified as THE CAUSE."

"The new work turns previous thinking around."

Robert I Richards, Sarah A Robertson, Daniel L Kastner, "Neurodegenerative diseases have genetic hallmarks of autoinflammatory disease," Human Molecular Genetics, Volume 27, Issue R2, 01 August 2018, Pages R108–R118.

Theoretical Lipid PHYSIOLOGY

<u>Membrane</u> = Brick & Mortar a)

b) Eicosanoids = Cellular Messengers

Clinical Physiology 1st

Biochemistry 2nd

Epigenetic Solution Pathway Modulation / Cellular Terrain

Optimize <u>YOUR</u> Protocols



PEO: Parent Essential Oils (Essential Fatty Acids)

- Parent Essential Oils are Parent Omega-6 & -3 EFAs. processed / adulterated then poisonous.
- I coined this term to clearly delineate between **Parent & Derivative EFAs.**
 - Parent EFAs are Essential.

Derivative EFAs: EPA, DHA, AA, etc., are NOT EFAs because the body makes the derivatives (longer chain) from the 2 ESSENTIAL Parent Oils.

Food processors never use omega-3 fats, as these fats are far too oxygenreactive. Most Parent omega-3 in foods is not adulterated.

The problem lies *exclusively* in the processed, adulterated Parent omega-6 oils. This KEY issue is never properly addressed: "Processed vs Organic / **Unprocessed**."



Lipids are the #1 Modifiable Variable in Tissue Composition^{1,2}



1. Felton, CV, et al., "Relation of Plaque Lipid Composition and Morphology to the Stability of Human Aortic Plaques," Arteriosclerosis, Thrombosis, and Vascular Biology," vol. 17, no 7, 1997, pp. 1337-1345.

2. Wainwright, E, et al., "The Effects of Dietary n-3/n-6 Ratio on Brain Development in the Mouse: A Dose Response Study with Long-Chain n-3 Fatty Acids," *Lipids*, vol. 27, no. 2, pp. 98-103, 1992; Lands, WEM, et al., "Quantitative effects of dietary polyunsaturated fats on the composition of fatty acids in rat tissues," *Lipids*, vol. 25, no. 9, 1990, pp. 505-516.

LONGEVITY RADIANT HEALTH

Better Outcomes

Key FACT About Anti-inflammatory Δ -6 Desaturase (D6D)

Can an impaired Δ -6 Desaturase (D6D) metabolic pathway be cured? <u>NO</u>.

Once you are diagnosed with an inflammatory-based disease, your most powerful antiinflammatory metabolic **pathway** — Δ -6 Desaturase (D6D) — is permanently damaged, to a greater or lesser degree. Production of the body's most antiinflammatory — PGE_1 — will never be maximized.

Minimizing consumption of processed / adulterated plant seed-based oils will certainly help but will be insufficient for most patients. Medical Food EZtrek[®] is designed to help nutritionally compensate for impaired D6D pathways ON A DAILY BASIS.

The <u>Superiority</u> of Plant-Based Seed Oils...

- 100 TRILLION cells... Cell membrane is half protein / half lipid. Of every cell's lipid portion, 25% -33% are PEOs.¹
- Mitochondrion (cellular energy production) contain them —100s-1,000s in each cell (*cardiolipin* is **100%** Parent omega-6) — too.¹

KEY Insight: "...[S]ecretory cells [virtually all cells] are <u>hypersensitive to</u> changes of their membrane lipids induced by the diet [Cells sense adulterated] omega-6 oils and respond with stress and chronic inflammation]."² D6D metabolic pathway becomes highly impaired.

Today, <u>all patients suffer CHRONIC INFLAMMATION</u> from consuming PROCESSED / ADULTERATED Parent omega-6 cooking oils.

^{1.} Alberts, B., et al., Molecular Biology of the Cell (3rd edition), Garland Science, 1994, p 428; Murray, Robert K, et al., Harper's Illustrated Biochemistry (26th edition), McGraw-Hill, New York, 2003: p 97; Guyton, Arthur C & Hall, John E, Textbook of Medical Physiology (9th ed.), W.B. Saunders Co. 1996: 16, pp 861–862.

^{2.} Halbleib, K., et al., "Activation of the Unfolded Protein Response by Lipid Bilayer Stress," Molecular Cell, Vol. 67, Issue 4, pp 673-684.e8, August 17, 2017.

LA and ALA are the only 2 essential (body can't synthesize) fats.

- DHA from fish oil is NOT an EFA <u>NOT essential</u> body makes AS NEEDED
- EPA from fish oil is NOT an EFA <u>NOT essential</u> body makes AS NEEDED

21st Century Newsflash: < 1% (approx. 0.1%) of all PEOs are Converted into **Derivatives**^{1,2,3}

Contrary to popular belief, your body makes the derivatives AS NEEDED, such as DHA and EPA, with at least 99% staying in parent form.

^{1.} Salem N, Lin Y, Brenna JT, Pawlosky RJ. Alpha-linolenic acid conversion revisited. **PUFA Newsletter**, December **2003**.

Pawlosky RJ, Hibbeln JR, Novotny JA, Salem N Jr." Physiological compartmental analysis of alpha-linolenic acid metabolism in adult humans." J Lipid Res 2001;42:1257-65. 2. Govens PL, Spilker ME, Zock PL, Katan MB, Mensink RP. "Conversion of alpha-linolenic acid in humans is influenced by the absolute amounts of alpha-linolenic acid and linoleic 3. acid in the diet and not by their ratio." Am J Clin Nutr **2006**;84:44-53.

HEART DISEASE- #1 Killer

- 1) LDL-C is the TRANSPORTER of the Parent Omega Oils (esterified).
- 2) Lowering LDL-C was a good 1st cut, but the correct answer is to increase UNprocessed Parent omega-6 because 85% of an arterial clog is from **CONSUMING PROCESSED /Adultrated Omega-6 (already adulterated / oxidized** from food processing – NOT going "bad" in the body).¹
- 3) There is no SATURATED fat in an occluded (clogged) artery.¹

1 Felton, CV, et al., "Dietary polyunsaturated fatty acids and compositions of human aortic plaque," Lancet; "Identification and quantification of unique fatty acid and oxidative products in human atherosclerotic plaque using high-performance lipid chromatography," Annals of Biochemistry; "Structure elucidation of oxygenated lipids in human atherosclerotic lesions," Eicosanoids.

Major Newsflash 2009: <u>American Heart Association¹</u> Champions Omega-6 PUFAs to Counter Popular Nutrition Advice

A great deal of discussion in the world of nutrition has given omega-6 fatty acids a bad reputation, which, according to the American Heart Association, is unfounded. The debate came about because one of the components of omega-6 fatty acids, called *arachidonic acid*, is a "building block" for some inflammation-related molecules. This has led to concern that omega-6 consumption would lead to a greater risk of heart disease. This has been completely disproven.

""That reflects a rather naive understanding of the biochemistry,' says William S. Harris, Director of the Metabolism and Nutrition Research Center of the University of South Dakota Sanford School of Medicine and the nutritionist who led the science advisory committee that issued the report in Circulation." "'[O]mega-6 PUFAs [Derivatives] also have powerful anti-inflammatory properties that counteract any proinflammatory activity,' say the advisory authors. 'It's incorrect to view the omega-6 fatty acids as "proinflammatory."

^{1.} AHA Heartwire 2009, © 2009 Medscape, January 28, 2009 (Dallas, Texas), based on Journal of the American Heart Association. Ref.: AHA Science Advisory, Harris WS, et al., "Omega-6 fatty acids and risk for cardiovascular disease: a science advisory from the American Heart Association Nutrition Subcommittee of the Council on Nutrition, Physical Activity, and Metabolism; Council on Cardiovascular Nursing; and Council on Epidemiology and Prevention" downloaded from circ. ahajournals.org on January 29, 2009. Published in *Circulation*. 2009;119:902-907.

2017 Confirmation of Omega-6 Series: Serum Fatty Acid Analysis Gives the Truth Parent and Derivative Omega-6 is Beneficial, NOT HARMFUL / NOT Inflammatory

This newly reported analysis CONFIRMS OTHER STUDIES showing that both Parent omega-6 (LA) and <u>arachidonic acid (AA) are not inflammatory</u>—as measured by C-reactive protein (CRP), a strong, key marker of inflammation.

"Conclusions: Serum <u>n-6</u> PUFAs [AA, etc.] were <u>not associated with increased inflammation</u> in men. In contrast, the main n-6 PUFA linoleic acid [Parent omega-6] had a strong inverse association with the key inflammation marker CRP.

"Omega-6 fatty acids do not promote low-grade inflammation.

"The higher the serum linoleic acid [Parent omega-6] level, the lower the CRP."



^{*} Virtanen, JK, et al., "The associations of serum n-6 polyunsaturated fatty acids with serum C-reactive protein in men: the Kuopio Ischaemic Heart Disease Risk Factor Study," European Journal of Clinical Nutrition, online accessed November 18, 2017, https://doi.org/10.1038/s41430-017-0.

Parent Omega-6 Newsflash¹

• "(18:2) CL₄ [Parent omega-6] rescues [fixes the damage] the major remodeling in the cardiolipin lipidome **induced by long-term intake of DHA.** [Cardiolipin is in the inner mitochondrial membrane. Mitochondria are the cellular ENERGY SOURCES. Deficiency causes Chronic EXHAUSTION – the #1 complaint of Americans.]

• "...[I]t is **not the loss of linoleic acid alone** that drives the impairment in enzyme function since the Western diet alone did not impair enzyme activities. Instead, it was the replacement of linoleic acid with DHA that promoted the reduction in activities."

CONCLUSION: **2018**: **DHA** [Omega-3 DERIVATIVE] **RUINS** HUMAN Cardiac **Mitochondria** = Heart Failure Due to LACK of CELLULAR ENERGY:¹

Sullivan, E. Madison, et al., "Docosahexaenoic acid lowers cardiac mitochondrial enzyme activity by replacing linoleic acid in the phospholipidome," *Journal of Biological Chemistry*, **2018**, 293: 466-2018 Jan 12;293(2):466-483.

When these precious PEOs become oxidized (from food processing), oxygen transfer stops, and cardiovascular disease starts.

Outdated Medical Textbooks state the intima consists of a single layer (it is actually 15-20 layers), increasing potential for atherosclerosis – Prof. Subbotin) of endothelial cells containing significant LA, but no ALA.^{1,2} Consumed, processed (nonfunctional) LA deposited in arterial intimal cell membranes leads to *abnormal oxidation at the vascular injury site*, thus causing injurious inflammation. (Note: In this case, abnormal oxidation involves formation of a hydroperoxide from LA.)

• The culprit in our food supply is ubiquitous "processed foods." Masquerading as harmless, processed / adulterated oils — exclusively in Parent omega-6 oil — are the villains. ••

^{1.} Chapkin RS, Ziboh VA, Marcelo CL, Voorhees JJ. Metabolism of essential fatty acids by human epidermal enzyme preparations: evidence of chain elongation. *J Lipid Res* 1986: 27:945-954.

^{2.} Andersson A, Sjödin A, Hedman A, Olsson R, Vessby B. Fatty acid profile of skeletal muscle phospholipids in trained and untrained young men. Am J Physiol Endocrinol Metab 2000;279:E744-E751.

Lipids are the Variable in Tissue Composition We Make Great Use of This Medical Fact

- Tissue lipid composition is often LINEAR (proportional) to lipid intake Α. **percentages** (as are plasma, liver, and RBCs).¹
- The percentage of adulterated LA (Parent omega-6) in the tissue is in Β. proportion to the adulterated amount consumed.²

[As much as] 85% of arterial plaque occlusions are adulterated LA / C. oxidized derivatives are toxic (and inflammatory) to arterial cells³.

- 1. E. Wainwright, Y. S. Huang, et al., "The effects of dietary n-3/n-6 ratio on brain development in the mouse: a dose response study with long-chain n-3 fatty acids," Lipids, vol. 27, no. 2, pp. 98–103, 1992...
- 2. Lands, WE, et al., "Quantitative Effects of Dietary Polyunsaturated Fats [EFAs] on the Composition of Fatty Acids in Rat Tissues," Lipids, Vol. 25, No. 9, 1990, pages, 505-516.
- 3. C.V. Felton, et al., "Relation of Plague Lipid Composition and Morphology to the Stability of Human Aortic Plagues," Arteriosclerosis, Thrombosis, and Vascular Biology," Vol. 17, No. 7, 1997, pp. 1337-1345; C. V. Felton, D. Crook, M. J. Davies and M. F. Oliver, "Dietary Polyunsaturated Fatty Acids and Composition of Human Aortic Plaques," The Lancet, Vol. 344, No. 8931, 1994, pp. 1195-1196; H. Kühn, J. et al., "Structure Elucidation of Oxygenated Lipids in Human Atherosclerotic Lesions," Eicosanoids, Vol. 5, 1992, pp. 17-22; Peskin, B., "Why Fish Oil Fails to Prevent or Improve CVD: A 21st Century Analysis," Food and Nutrition Sciences, 2013, Vol. 4, No. 9A, 76-85.

Nature's Natural Statin: Omega-6 series

- **Prostacyclin** (PGI₂): Body's MOST potent natural blood thinner / platelet anticoagulant is made from arachidonic acid (AA).¹
- PGE₁: Body's most potent anti-inflammatory much stronger than Omega-3's PGE₃.²

Important Point: The intima (endothelial inner lining of arterial wall) EFA structure is entirely Parent omega-6 — with no Parent omega-3. Epithelial tissue and the skin contain no omega-3, either; regarding EFA content, it is exclusively Parent omega-6.

Bunting, S., Moncada, S., and Vane, J.R., "Prostaglandin-Thromboxane A2 Balance: Pathophysiological and Therapeutic Implications," British Medical Journal, (1993) Vol. 39, No. 3, 271-276.

Schmidt, M.A., Smart Fats: How Dietary Fats and Oils Affect Mental, Physical and Emotional Intelligence, 1997, pp 27-30.

Nobel Prize-winner Otto Warburg, MD/PhD: Seminal Cancer Discovery **#1 Physiologist of the 20th Century**



Otto Warburg, "On the Origin of Cancer Cells," Science, February 1956, Volume 123, Number 3191.

- The Oxygen/Cancer Connection: Cancer is always caused by long-<u>term</u> deprivation of oxygen (intermittent 35%) from chronic *inflammation.* This is the *prime* cause of cancer.* Heart disease (CVD) is caused by oxygen deprivation, too.
- All *secondary* causes of cancer, like asbestos, smoking, or other carcinogens, MUST lead back directly to the *prime* cause.
- Back then, Dr. Warburg didn't have the most efficient method to get vital oxygen to the cell... Fortunately, today, we do with Organic / fully functional Omega-6 based seed oils coming to the rescue.

Deficient Parent Omega-6 Causes Cancer Throughout the Body¹

- "...[R]estoration of a **normal fatty acid composition** should **enhance the supply of** Α. oxygen to the tissues and improve the general health of the patients...
- Β. "...<u>Interference with the movement of oxygen could then occur at any cell</u> **membrane** so that there could be a general reduction in the supply of cellular oxygen throughout the body..." [This is precisely why Cancer / Heart Disease may occur anywhere this condition occurs.]
- "...[M]any of their symptoms may result from essential fatty acid (Linoleic / LA) С. deficiency, leading to the decrease in the availability of cellular oxygen for respiration [*mitochondria's cardiolipin*—ALL Parent omega-6]...."

1. Campbell, IM, et al., "Abnormal fatty acid composition and impaired oxygen supply in cystic fibrosis patients, Pediatrics, 57:480-486, 1976.



Confirmation of the O_2 / Cancer Connection - ALWAYS in every case

- **A.** Published in 1993 and 1999, the cancer medical journal Radiotherapy and Oncology makes Dr. Warburg's seminal discovery clear. Intratumoral pO₂ predicts survival in advanced cancer of the uterine cervix.¹
- Also, in **1999**, the same cancer journal made Dr. Warburg's seminal fact clear again in the article Β. titled, "Oxygenation of head and neck cancer: changes during radiotherapy and impact on treatment outcome."²
- "Tumor oxygenation affects the prognosis of head and neck cancer independently of other C. known prognosis variables."
- "Tumor oxygenation predicts the likelihood of distant metastases [cancer spreading] in human soft D. tissue sarcoma (connective tissue-based cancer)."³
- Ε. "Tumor hypoxia [too little oxygen in the cell] adversely affects the prognosis of carcinoma of the head and neck."⁴

Knoop, Hockel, et al., "Intratumoral pO2 predicts survival in advanced cancer of the uterine cervix," Radiotherapy and Oncology 1993, Jan;26(1):45-50.

Brizel, D.M. et al., Radiotherapy and Oncology, 53, (1999): 113-117.

Brizel, D.M., et al., Cancer Research 1996:56-941-43. 3.

Brizel, D.M., et al., Int. J. Radiat. Oncol. Biol. Phys. 1997;38:285-290. 4.

2007: Inflammation Fuels Cancer¹ "Re-writing of the textbook..."

- Cancer researcher Robert Weinberg of MIT states: "The connection inflammation and cancer has moved to center stage in the research arena." The article continues...
- "...**inflammation is the fuel** that feeds it [the malignant cancer].
- "In this **rewriting of the textbook**... This new view implies that rooting out every last cancer cell in the body might not be necessary. Anti-inflammatory cancer therapy instead would prevent pre-malignant cells from turning fully cancerous or would **impede an existing tumor from spreading** to distant sites in the body. Cancer victims might then be able to survive."

1. Balkwill, F., et al., "Smoldering and Polarized Inflammation in the Initiation and Promotion of Malignant Disease," Cancer Cell, Vol. 7, No. 3, pages 211-217, March 2005; "Distinct Role of Macrophages in Different Tumor Microenvironments," Lewis, C. and Pollard, J., Cancer Research, Vol. 66, No. 2, pages 605-612; January 15, 2006; "Paradoxical Roles of the Immune System during Cancer Development," Visser, K., et al., Nature Reviews Cancer, Vol. 6, No.1, pages 24-37; January 2006; One Renegade Cell: How Cancer Begins, Robert A. Weinberg, Basic Books, New York, 1998, p. 146.

between

In The News: 2009

Nearly a Century Later, New Findings Support Warburg Theory of Cancer¹ (Note: Appropriately, this work was supported by the *National Cancer Institute* and the National Institute on Aging).

 "Major abnormalities in CL [cardiolipin] content or composition were found in all tumors. Hence, our findings in mouse brain tumors provide evidence linking abnormal CL to irreversible respiratory injury."

L.S.E. Analysis: What is cardiolipin (CL)? It is a **fat-based** complex phospholipid **found in all** mitochondrial membranes – intimately involved in maintaining mitochondrial functionality and membrane integrity. It is used for ATP synthesis and consists roughly of 20% lipids.² In mammals, the main substrate in CL is Parent omega-6 with virtually no Parent omega-3 or its derivatives.³

^{1.} Science Daily (January 14, 2009) and Kiebish MA, Han X, Cheng H, Chuang JH, Seyfried TN. Cardiolipin and electron transport chain abnormalities in mouse brain tumor mitochondria: lipidomic evidence supporting the Warburg theory of cancer. J Lipid Res 2008;49:2545-66.

Krebs JJ, Hauser H, Carafoli E. Asymmetric distribution of phospholipids in the inner membrane of beef heart mitochondria. J Biol Chem 1979;254:5308-16. 2.

Scottish Crop Research Institute (MRS Lipid Analysis Unit, Invergowrie, Dundee DD2 5DA, Scotland). Cardiolipin (diphosphatidylglycerol). Structure, occurrence, biology and 3. analysis. Retrieved January 20, 2009 from: http://www.lipidlibrary. co.uk/Lipids/dpg/index.htm.

Pharmaceutical Overdose:

Based on Physiology, Fish Oil Can't & Doesn't Work

Fish oil is (typically) *exclusively omega-3 derivatives* with enormous supra-physiologic overdose factors of plasma EPA and DHA. Therefore, prophylactic use has no basis in human physiology whatsoever. Just one (1) typical 1,000 mg capsule of fish / marine oil contains 100Xs the daily requirement of DHA. No success for CVD, Cancer, Vision, Alzheimer's, etc. –

Cochrane Reports --- the world leader in evaluating studies gave marine oils a dismal failure rating....

Fish / Marine Oil FAILURES: Extremely LITTLE IS NEEDED

New 21st Century Analysis: NIH researchers determined the amount of **DHA utilized in human brain tissue to be** a mere 3.8 mg \pm 1.7 mg / day. Therefore, brain tissue in 95% of all subjects, allowing for variation in brain size, would consume or naturally produce a mere 0.4 mg – 7.2 mg of DHA per day.* [Nearly nothing.]

•• COMPARE this amount with the doses of fish oil recommended and prescribed.••

• Fish oil contains almost no Parent Oils — often **20-500-times DAILY overdose of** omega-3 derivatives EPA/DHA [pharmacologic, physiologic, DAILY OVERDOSE].

* J. C. Umhau, W. Zhou, R. E. Carson, S. I. Rapoport, A. Polozova, J. Demar, et al., "Imaging Incorporation of Circulating Docosahexaenoic Acid [DHA] into the Human Brain Using Positron Emission Tomography," Journal of Lipid Research, Vol. 50, No. 7, 2009, pp. 1259–1268.

21st Century (2008) MAJOR NEWSFLASH: CONFIRMED—EFA **Derivatives Made "As Needed"**¹

"...[W]hich shows the effectiveness of ALA conversion [into DHA and EPA] and accretion into erythrocytes. The amounts of ALA [Parent omega-3] required to obtain these effects are amounts that are *easily achieved* in the general population by dietary modification [ADD] Parent omega-3]."

There is independent confirmation of conversion amounts, if anyone would care to look...²

•• Important Note: 2,600:1 times more ALA (Parent omega-3) than DHA (derivative of Parent omega-3) in the human body ••

^{1. &}quot;Flaxseed oil and fish-oil capsule consumption alters human red blood cell n-3 fatty acid composition: a multiple-dosing trial comparing 2 sources of n-3 fatty acid," American Journal of Clinical Nutrition, Vol. 88, No. 3, pp 801-809, September 2008.

Hussein, Nahed, et al., "Long-chain conversion of linoleic acid and alpha-linolenic acid in response to marked changes in their dietary intake in men," Journal of Lipid 2. **Research**, Volume 46, 2005, pp 269-280.

Fish oil accelerates aging

Fish oil increases oxidative stress and decreases lifespan.

- "<u>Conclusion: These findings suggest that intake of fish oil increases oxidative</u> stress, decreases cellular function, and causes organ dysfunction."*
- "<u>Docosahexaenoic acid</u> (22:6) -- **DHA**, which has six double bonds and consequently five bis-allylic hydrogens per chain, is **320 times more susceptible** to attack than the common mono-unsaturated oleic acid (18:1) -- like olive oil -which has "no" bis-allylic hydrogens in its chain.... Membrane lipid peroxidation should not be perceived solely as a 'damage to membranes' scenario but also as <u>a</u> significant endogenous source of damage $\frac{4}{10}$ other cellular macromolecules, such as proteins and DNA (including mutations)."**

[•] Tsuduki, K., et al., Long-term intake of fish oil increases oxidative stress and decreases lifespan in senescence-accelerated mice," Nutrition 27, (2011), pages 334–337. ** Hulbert, A.J., "Metabolism and Longevity: Is There a Role for Membrane Fatty Acids?" Integrative and Comparative Biology, Volume 50, Number 5, 808–817, 2010.

Fish oil increases oxidative stress

"Ten (10) grams of fish oil (18% EPA & 12% DHA) daily over 3 or 6 months increased **TBA** [a prime and significant **measure of lipid rancidity] from 6 to** 18.5... [3-fold increase].

"...omega-3 PUFAs [derivatives], which are *relatively* <u>unstable</u> compared with omega-6 fatty acids...

"Following **3 months of fish oil** supplementation, there was a **pronounced rise** in the total ω-3 fatty acid content of unirradiated skin. 'We confirmed the reported incorporation ω -3 PUFAs [derivatives] into *epidermal membrane* lipids after dietary fish oil supplementation."

^{*} Rhodes, Lesley, E., et al., "Dietary Fish–Oil Supplementation in Humans Reduces UVB-Erythemal [an abnormal red condition of the skin, resulting from capillary congestion] Sensitivity but Increases Epidermal Lipid Peroxidation," The Journal of Investigative Dermatology, 103:151–154, 1994.

Is Cod Liver Oil Better? <u>NO</u>. Cod Liver Oil Significantly Increases Risk of Melanoma¹

"A significant risk was found in women who used cod liver oil supplement. [W]e found a strong increased risk for the women using cod liver oil, a supplement rich in omega-3 fatty acids (EPA and DHA)." [There was approximately a **3xs greater incidence of melanoma (the most dangerous type of** skin cancer) in the cod liver oil users.]

"The increase is considered to be real and <u>not due to chance</u>."

"Mean time of follow-up was 12.4 years...." [Note: Sufficient time for an excellent analysis.] "The strengths of the study are the high number of participants selected in an unbiased manner, the high participation and response rate, the prospective design with *dietary data collected prior to onset of cancer*, and a complete follow-up with regard to incidence of cancer, deaths, and emigration. The complete follow-up is secured by the procedure established by the Cancer Registry, ensuring that all physicians, hospital departments, and histopathology laboratories in Norway are obliged to report malignant diseases to the Registry: as many as 98% of the cases were histologically [microscopic tissue analysis] verified." [Note: This guarantees superb tracking and confirmation of cancer cases.]

Fish Oil Has NO DIRECT Anti-Inflammation Benefits – Lowers Immune System Most notably, mimicking the immune-suppression properties of a steroid

• FAILURE 2000: Fish Oil decreases immune response*

• "...[S]tudies indicate that at the levels used, fish oil [omega-3 derivatives] decrease a wide range of immune cell responses (natural killer cell, cytotoxic T lymphocyte activities, lymphocyte proliferation and production of IL-2 and IFN-y (1,2))....

• "...Recent studies have indicated that <u>relatively low levels</u> of the long-chain omega-3 fatty acids (EPA or DHA)...<u>are sufficient</u> to bring about some of the suppressive effects..."

— WARNING: DHA & EPA, even in low doses, cause these immune-suppressive effects. —

* Calder, P., "Omega-3 Polyunsaturated Fatty Acids, Inflammation and Immunity," Institute of Human Nutrition, University of Southampton, Bassett Crescent End, Southampton, UK. Presented at: The International Society for the Study of Fatty Acids and Lipids (ISSFAL) 4th Congress, which met on June 4-9, 2000, in Tsukuba, Japan.

Patient CVD Screening to Detect STIFF ARTERIES

The World's Most Advanced Cardiovascular Measurement System in **Under 60 Seconds**

Has this happened to you or your loved ones?

- ... My cholesterol is **perfect**, but I had a Heart Attack.
- ... My blood pressure is perfect, but I had a Heart Attack.
- ... My labs are **perfect**, but I had a Heart Attack.
- ... I had no symptoms, but I had a Heart Attack. (30% of 1st Heart Attacks result in death)



Lipids in the Bloodstream

Percentages of Linoleic Acid (LA) & Alpha Linolenic Acid (ALA) in Plasma and Classes of Lipids				
Fatty Acid	Plasma % (Unesterified)	Plasma % Triglycerides	Plasma % Phospholipids	Plasma % Cholesterol Esters
LA (parent omega-6)	17	19.5	23	50
ALA (parent omega-3)	2	1.1	0.2	0.5
Parent omega-6: Parent omega-3 Ratio	8.5:1	17.5:1	115:1	100:1

- Approximately **70% of the cholesterol in the lipoproteins** of the plasma is in the form of cholesterol esters attached to ApoB.¹
- Of dietary cholesterol absorbed, 80%–90% is esterified with long-chain fatty acids in the intestinal mucosa.² The majority (about 55%-85%) of the *cholesteryl ester* component is LA (*Parent omega-6*).³
 - 85% of the material clogging an artery is adulterated / nonfunctional / oxidized Parent omega-6 by FOOD PROCESSING (See slides 17 & 22).
- Guyton A, Hall J. Textbook of Medical Physiology, 9th ed. Philadelphia, Pa:WB Saunders; 1996:872-873.

Sinclair HM. Essential fatty acids in perspective.1984;38C:245-260.

Bothem KM, Mayes PA. Cholesterol synthesis, transport, and excretion. In: Murray PK, Granner DK, Mayes PA, Rodwell VW, eds. Harper's Illustrated Biochemistry 27th ed. New York, N.Y.: McGraw-Hill; **2003**; 235.

Utilizing the D6D Anti-Inflammatory Pathway to Maximize <u>YOUR</u> Protocol



Info: 69-year-old male Cigar smoker, no exercise, social drinker, employed, caretaker for wife.

Results: Vessel state and blood circulation are excellent -- At LEAST 25 years biologically younger than chronological age.



pt using plant-based seed oil -- EZtrek® Note: Skin elasticity is excellent.

Given age (69) & cigar smoker, this is outstanding.

Pt using ORGANIC plantbased seed oil — EZtrek[®]—

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