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Hen egg yolk oil: A potential source of bioavailable lutein and zeaxanthin for skin and sun protection

Kumud Madan and Sanju Nanda*

Dept. of Pharm. Sciences, Maharshi Dayanand University, Rohtak, Haryana, India

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ABSTRACT

Ultraviolet radiation is the most harmful of electromagnetic spectrum causing deleterious effects on human skin which are responsible for severe dermatological conditions ranging from tanning, epidermal thickening, erythema, photoaging, immune suppression and skin cancers. The use of sunscreens has become a prerequisite to prevent skin from the detrimental effects of ultraviolet radiations. The adverse effects of synthetic agents is shifting the mindset of the consumer towards the use of natural sources due to their antioxidant properties. One such natural agent reported to be augmented with antioxidants is egg oil, extracted from the yolk of avian eggs. Egg yolk oil, obtained by cold extraction, is a source of phospholipids, triglycerides, cholesterol and similar compounds which resemble the cell membrane in its constitution. Its unique properties of wound healing, burn healing and anti-inflammation has already been established. Egg oil is therefore now being explored as a skin and sunprotectant mainly because it contains abundant lipids and enriched with antioxidant molecules such as *lutein* and *zeaxanthin*. This review article discusses the extraction methods, constituents, dermatological applications, commercial sources and patents signifying the tremendous potential of egg oil.

Keywords: sunprotection, natural, egg oil, lutein, zeaxanthin, photoaging, antioxidant.

INTRODUCTION

The electromagnetic spectrum ranges from short wavelength cosmic rays to high wavelength radio waves. Amongst these, ultraviolet (UV) radiation consists of three distinct bands with different wavelengths. Although UV-C is filtered by ozone layer, UV-A and UV-B are responsible for sunburn to human skin^[1]. Acute response to UV exposure includes tanning, epidermal thickening and erythema^[2]. Chronic exposure to UV causes [3] photoaging immunosuppression and carcinogenesis.

Photo-exposure induces series of molecular and cellular changes that trigger a fast and dynamic disorder in the skin^[3]. The various structural and functional changes include increase in thickness of epidermis, stratum corneum layers, massive increase in elastic fibres with increased lysozyme, cleavage of collagen and wrinkle formation^[4]. UVB radiations are responsible for damage to the lipid barrier whereas UVA is reported to enhance the effect of UVB on the epidermis^[5].

UV radiation causes oxidative damage which contributes to inflammation, gene mutation and immunosuppression. All these changes lead to carcinogenesis. The International Agency for Research on Cancer (IARC) has established UV rays as a human carcinogen^[6]. The form of exposure to ultraviolet radiation plays a significant role in degree and risk of cancer developed. Intermittent ultraviolet exposure causes cutaneous malignant melanoma and basal cell cancer while chronic exposure is responsible for squamous cell cancer. Melanin granules in the epidermal cells of the skin is said to possess an SPF of 4, but if the exposure is for a longer time, it may not be possible to protect the skin from photo damage. Therefore, the use of sunscreens has become a requisite to prevent skin from harmful ultraviolet radiations. Though sunprotection measures like umbrella, clothes, sunglasses have always been suggested but topical formulations like creams and lotions for extemporaneous and instant application are popular due to convenience and ease of application.

*Corresponding Author Address: Dr. Sanju Nanda, Dept. of Pharm. Sciences, Maharshi Dayanand University, Rohtak, Haryana, India; E-mail: sn_mdu@rediffmail.com

French chemist Eugène Schueller, founder of L'Oreal, is reportedly the first person to introduce sunscreen products commercially in the year 1936^[7]. Since then, there has been a considerable progress in the field of sunscreens and the market is flooded with sunscreens in various form like creams, lotions, gels, sticks, etc.

The performance and efficacy of these products is gauged by their SPF value. SPF or the "Sun Protection Factor" can be defined as the numerical ratio between the minimal erythemal dose (MED) of sunscreen-protected skin, applied in the amount of 2mg/cm² and the minimal erythemal dose (MED) of unprotected skin^[8]. Mathematically it can be expressed by the following equation:

SPF= MED Protected/MED Unprotected

Sunscreen agents are broadly classified as physical and chemical sunscreens. Physical or inorganic sunscreens act by reflecting or scattering UV rays while chemical or organic sunscreen absorbs them¹⁶ ¹. A list of sunscreen agents approved by USFDA is provided in Table 1. Nevertheless, all of them claim to be highly sun protective and safe. However, the studies carried out by various researchers have reported systemic and topical toxicities of these agents. Some of them are tabulated in Table 2.

The planet is filled with people of different skin phototypes ^[9] and hence there could be no ideal product which could satisfy all of them[Table 3]. However, the inclination towards natural products is gaining momentum not only due to the fact that they are from natural origin or less irritant but also because they provide holistic benefits.

Specifically, certain classes of compounds like flavonoids, carotenoids and terpenoids from natural sources are rich in anti oxidative protective network^[10] for example, aloe vera, ^[11]tomato,^[12] pomegranate,^[13]grapes,^[14] cucumber,^[11] green tea,^[15] almond,^[16] *etc.*

Antioxidants are responsible for reducing the damage caused by free radicals, thus avoiding damage at the cellular level. They also help to inhibit inflammation and offer protection against photo-damage and photocarcinogenesis. These antioxidant supplements can potentially boost the body's natural reserve and neutralize Reactive Oxygen Species (ROS) from intrinsic sources (e.g., cellular metabolism) and extrinsic factors (e.g., UV damage and environmental pollution). They either inhibit generation of ROS or quench the formed free radicals.[Figure 1] Besides this, the fixed and essential oils also contribute additional

;l properties like emolliency, occlusiveness and nourishment to the skin.

In case of chronic exposure to UV rays, oxidative stress is induced in the cells of epidermis and dermis due to the production of reactive oxygen species like '*singlet molecular oxygen*' and '*peroxy radicals*'. It is now established that carotenoids such as lutein and zeaxanthin react with these free radicals and nullify their effects^[17].

Natural oils and sunprotection: Fats and fatty acids present in fixed oils and essential oils have been used topically from time immemorial as emollients, protectants and occlusives. These oils also possess remarkable skin penetrating power. As compared to oils obtained from petrochemical industries, oils extracted from seeds of fruits and vegetables are light, low in viscosity and allows skin to breathe^[18]. Rich content of tocopherols, carotenoids and essential fatty acids in these oils highly valuable make them as natural sunprotectants. Oils from almond, coconut, cottonseed, olive, peanut, sesame, soyabean have been reported to possess a considerable SPF value too.

Shea butter, ^[19] jojoba oil, ^[20] borage oil,^[11] and fish oil^[21] have also been used in prevention of photoaging due to their antioxidant constituents.

The quest to add new but well documented sources of natural sunprotective agents has lead to the discovery of oil extracted from avian eggs (duck,fowl,hen,etc.). Being rich in bioavailable lutein and zeaxanthin, hen egg oil stands high in merit, especially in topical formulations meant to treat inflammation and photodamaged skin conditions. Hen eggs obtained from Gallus gallus domesticus are sources of protein, fats and micronutrients that play an important role in basic nutrition^[22]. It contains a variety of bioactive compounds that can influence pro- and antiinflammatory pathways^[23]. Egg white and yolk, contain numerous compounds with both activities, namely antioxidant ovalbumin, ovotransferrin, phosvitin, phospholipids, vitamin E, vitamin A, selenium, and carotenoids ^[22]. The presence of more phospholipids and saturated fatty acids make egg yolk a very good antioxidant probably due to the presence of many physiologically active components such as lutein, zeaxanthin lysozyme, sialic acid, etc. Bioavailability of egg carotenoids is considered to be superior as compared to those from green leafy vegetables due to their increased solubilization in yolk lipids. This makes egg a unique and important carrier of bioactive carotenoids as reported by Chung *et a*.^[24].

Hen Egg Yolk Oil: Source of Lutein and Zeaxanthin: Hen egg yolk oil also known as chicken egg oil, ovum oil or egg oil has been used in traditional cosmetics since time immemorial and is an established source of phospholipids, triglycerides and cholesterol. It is a potential source of lutein and zeaxanthin too whose antioxidant property is well known [22]. Both dihydroxycarotenoids with the ionone ring systems substituted at both 3 and 3' carbon^[25]. The presence of free hydroxyl group at each end of these molecules provides them unique biochemical properties [Figure 2]. They play an active role in the scavenging of two major ROS viz. singlet molecular oxygen and peroxy radicals and are effective deactivators of electronically excited sensitizer molecules responsible for the generation of radicals and singlet oxygen^[26].

Procurement of Egg oil: Egg oil is obtained from yolk of avian eggs and they have been of interest to researchers from time to time. Hen eggs are preferred over other avian eggs in view of their low cost and ready availability. The quality of egg can also be modified by dietary components fed to hen. Some of the methods of extraction of egg oil from egg yolk are described below:

Heating method: Weaver B Cooper patented a method of egg oil extraction for the first time in 1950. He heated the yolk portion of hard boiled eggs with a fat as a starter until a dark brown liquid of low viscosity was obtained^[27].

Solvent Extraction method: Since heating method lead to loss of thermo labile constituents present in egg oil, liquid- liquid extraction method was by Larsen and Frowning in 1980. The adopted crude oil fraction obtained was degummed, refined and bleached. They also recommended improvement of refinement techniques so that the extraction could be done at a commercial scale^[28]. Warren et al modified this procedure in 1988 by extracting the lipid components from spray dried egg yolks using a mixture of hexane: isopropanol and chloroform: methanol (2:1). They concluded that hexane removed less cholesterol and pigment than polar solvents ^[29]. The extraction process was futher improvised by Werner C Nawrocki in 1997 in which he claimed high purity egg oil suitable to be used for treating skin burns, including sunburn, and for skin regeneration ^[30]. Aleksandrs Kovalcuks et al further developed this process by using 2-propanol/hexane to give a better yield [31]. Simultaneously, a novel process was invented in 1991 by Jeong S Sim for the separation of different fractions of fresh liquid egg yolk such as protein, lecithin and neutral egg oil^[32].

Supercritical Fluid Extraction: Various fractions of egg yolk such as cholesterol and phospholipids have also been extracted and patented using supercritical fluid extraction method. The reported involved treating egg yolk with a method vegetable oil, homogenizing the mixture and then extracting the homogenate with supercritical carbon dioxide at an extraction pressure between 2,000 and 4,500 psig and a temperature between 31° C. and 45° $C^{[33]}$. This process proved to be faster, safer and economical. Amongst the various methods, solvent extraction method is the most widely used method. [Figure 3] Egg oil is available in market under brand names of OLEOVA, (VAV Life Sciences), charismon[©]. etc.[Table 5] A brief description of patents regarding extraction and use of egg oil is given [Table 6].

Skin and sun protective applications of egg oil: Egg oil being rich in cholesterol, triglycerides, carotenoids, tocopherols, phospholipids and other valuable components, has been exalted for its emollient, wound healing, anti-inflammatory, antiaging, antimicrobial and anti rachitic activity by various research groups.

Emollient action: Egg oil, being rich in cholesterol has an ability to nourish and moisturize the skin and hair. As the oil is protein-free, the cholesterol is readily absorbed into the skin and is suitable for those who are allergic to egg proteins. It has also been used to formulate cosmetic emulsions like hand creams, body creams and massage creams, *etc.* when added in a range of 0.3 to 0.4% by weight^[34].

Burns and wound healing: Burn injuries are one of the major causes of death and disability in the world and hence a local wound healing medication which can promote re-epithelialisation would always be appreciated. The egg oil seems excellent in healing process as proved by a research carried out by Rategar et al. Egg oil was applied topically burnt skin of the animal which showed to abundant re-epithelialization without tissue scar in comparison to the animals treated with silver sulphadiazine ^[35]. It has also been patented as a 'burn healer' by Carol J. Burg for the treatment of second and third degree burns to human skin^[36]. Another research study carried out by Hahn et al cites the wound healing potential of charismon[©], a branded egg oil^[37]. They studied the effects of egg oil on a variety of physiological reactions in cells. HaCaT-cells (immortalized human keratinocytes), human umbilical vein endothelial cells in culture (HUVEC), peripheral blood mononuclear lymphocytes (PBML) and a full thickness human skin model (FTSM) were used to study the cellular

responses. The main objective of their study was to see the influence of egg-oil on cell migration and proliferation, mitochondrial membrane potential, reactive oxygen (ROS) production and IL-8 production. According to them, egg oil promoted wound healing by re-epithelialisation and some other interlinked reactions. They also reported that egg oil containing toothpastes are available for the relief of gum inflammation and bleeding and egg oil containing ointments for healing of ulcers, scar formation and burn wound healing.

Analgesic and anti-inflammatory activity: Inflammation is purported to eliminate the initial cause of cell injury, clear out necrotic cells and tissues damaged from the original insult and the inflammatory process, and to initiate tissue repair. The cell injury can result from irritant or pathogen. Sunburn can also cause skin inflammation primarily by ROS which is responsible for cell death via DNA damage, necrosis and apoptosis. Egg oil modulated photosensitivity was studied by Hahn *et al* using full thickness human skin model (FTSM)^[37]. They observed that the cell section which was treated with egg oil had normal epidermal cells with usual nuclei even after UV irradiation as compared to FTSMs without egg oil application, which showed pycnotic nuclei. Pycnosis is an irreversible condensation of chromatin in the nucleus of a cell undergoing necrosis and apoptosis.

Egg oil is also reported to possess significant analgesic properties and has been recommended in inflammatory joint pain conditions. In a study conducted by Mahmoudi *et al* the oil extracted from duck egg and hen egg showed significant anti-inflammatory activity at 100-300 mg/kg as compared to controls (p < 0.001)^[38]. These oils demonstrated analgesic and anti-inflammatory activity, possibly due to the presence of *choline* since its presence reduced inflammatory markers by 20%. The presence of palmitoylethanolamide in phospholipid fraction of egg yolk can also be a contributing factor in reducing inflammation^[39].

Other indications: Psoriasis is a chronic inflammatory skin disorder that has a hereditary basis for susceptibility along with environmental factors such as trauma, infections $etc^{[40]}$. A patent filed in 1989 by Mohamed Abdessalam Yacobi, a qualified physician claimed the skin beneficial effects of a formulation containing egg oil along with sea salt, attar of roses and chalkatte in patients suffering from psoriasis ^[41]. He applied this formulation to the affected skin parts. In all the clinical studies carried out by him, the inventor noticed disappearance of hardened epidermis and

itching within 2 days of application and subsequent normalisation of the affected skin.

Egg oil is also reported to be helpful in skin aging (improvisation of skin texture and wrinkle formation delay) using a formulation on face, traditionally known as "crème du ciel" and containing 4.5% of this oil^[37]. This activity of egg oil can be attributed to the reduction of inflammatory processes and ROS production, as observed by various qualitative changes in skin parameters like moisture, surface lipids, elasticity, lipid peroxidation and photoprotection.

Rickets, a disease in which the bone tissue doesn't properly mineralize, leading to soft bones and skeletal deformities is associated with Vitamin D deficiency ^[42]. In a primeval study of 1934, an attempt to elucidate and compare the antirachitic efficacy of olive oil, egg oil and fatty acids of butter and lard was made by Stanisław Kazimierz Kon and Roland Gordon Booth^[43]. Though they claimed egg oil as probably the most concentrated source of vitamin D activity, any clinical data in this regard would be a ray of hope for vitamin D deficient patients.

RESULT AND DISCUSSION

Several organic and inorganic agents have been used to prevent the skin from deleterious effects of ultraviolet radiations of sun. These agents are also associated with various adverse effects which have shifted the paradigm to natural agents which are rich in antioxidants. Boosting the antioxidant system of the skin with the help of natural agents is a significant approach for reducing UV induced skin damage. One such spagyric agent discussed in this paper is egg oil which is derived from egg of hen.

Egg oil is a rich source of phospholipids, triglycerides, cholesterol and resembles the constituents of a biological membrane. It is also a good carrier for sunprotective agents where carotenoids such as lutein and zeaxanthin are stabilized with emollients. Additionally, the presence of phosvitin, and free aromatic amino acids make it a very good antioxidant. Being protein free, it is a safer product for the patients allergic to egg products.

The presence of cholesterol makes it a very good emollient when used in the concentration of 0.3-0.4%.It has tremendous applications in inflammation related skin conditions such as wound healing, burns, psoriasis *etc.* due its involvement in re epithelialisation. Studies by various researchers and their published results

provide a compelling explanation to unravel a newer, safer antioxidant rich compound from natural source.

Although clinical trials of egg oil are still going on by some producers of this oil, yet it can be used safely, being edible and natural in origin. An attempt was made by the authors to extract egg oil in the laboratory and determine its SPF^[44]. It can further be incorporated in various topical formulations and delivery systems such as liposomes and nanoparticles. The manufacturers, researchers and dermatologists should all come together and develop some commercially viable safe and effective topical products enriched with egg oil for the benefits of patients suffering from skin disorders.

Conflict of Interest: The authors have no conflict of interest.

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Sunscreen Agent	Permissible limit (%)
Organic Agents	
PABA	Upto 15%
Avobenzone	3%
Cinoxate	3%
Dioxybenzone.	Up to 3%
Homosalate.	up to 15%
Menthyl anthranilate.	up to 5 percent
Octocrylene.	up to 10 percent
Octyl methoxycinnamate.	up to 7.5 percent
Octyl salicylate.	up to 5 percent
Oxybenzone.	up to 6 percent
Padimate O.	up to 8 percent
Phenylbenzimidazole sulfonic acid.	up to 4 percent
Sulisobenzone.	up to 10 percent
Trolamine salicylate	up to 12 percent
Inorganic Agents.	
Zinc oxide	up to 25 percent.
Titanium Dioxide	up to 25 percent.

 Table 1: FDA Approved Sunscreen Agents^[44]

Table 2: Toxicity associated with sunscreen agents

S.No.	Toxicity	Reference
1.	Photodegradation of the sunscreen agent	[45]
2.	Endocrine disruptive properties of the chemical sunscreen agents	[46]
3	Clown like effect with higher concentration of physical sunscreens	[47]
4	Photogenotoxicity associated with nanosized physical sunscreen	[47]
5	Allergic contact dermatitis caused by PABA derivatives	[48]
6	Higher risk of melanoma associated with the use of sunscreens	[49]
7	Chemical oxidation and DNA damage caused by inorganic sunscreen agent	[50]
8	Increased ROS production upon penetration of organic sunscreens	[51]

Table 3: Fitzpatrick skin phototypes^[9]

Skin Type	Sunburn and tanning history
Ι	Burns easily, never tans
Π	Burns easily, tans minimally with difficulty
III	Burns moderately, tans moderately and uniformly
IV	Burns minimally, tans moderately and easily
V	Rarely burns, tans profusely
VI	Never burns tans profusely
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S.No.	Oil	Source	SPF	
1	Olive oil	Fruit of Olea europaea	7.5	
2	Coconut oil	Kernel of Cocos nucifera	7.1	
3	Castor oil	Seeds of Ricinus communis	5.6	
4	Almond Oil	Fruit of Prunus amygdalus,	4.6	
5	Mustard Oil	Mustard seeds	2.1	
6	Sesame oil	Sesame seeds	1.7	

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Table 4: SPF values of natural oils ^[18]	

Table 5: Egg oil Specifications

Synonyms: Egg Yolk Oil, Ovum Oil, Hen Egg Yolk Oil)	
CAS NO.: 8001-17-0,	
INCI Name: Egg oil	
BRAND NAMES: Charismon [©] , Oleova	
PHYSICOCHEMICAL PROPERTIES	COMPOSITION OF EGG OIL
	Moisture 1% Max
Appearance: Yellow coloured hazy liquid	Free Fatty Acids 1% Max
Consistency : Viscous Liquid, thickened semisolid at 25 ^o c	Triglycerides 92% Min
Odour:Mild characteristic odour of egg	Phospholipids 1% Max
Refractive Index:1.46148	Cholesterol 4% Max
Flash Point:>80 ⁰ c	Moisture 1.5% Max
Solubility in Water: Insoluble	Lutein and Zeaxanthin
Specific Gravity: 0.80-0.85.	

Table 6: List of Patents on egg oil extraction and applications

S.NO.	Patent No.	Publication year	Patentee	Claim as taken from respective patent	Reference
1.	US 2555731 A	1951	Weaver B cooper	Method of obtaining a new oil product from the yolks of eggs which is highly beneficial as a healing medicant for external use.	[27]
2.	US4219544A	1980	Carol J. Burg	Method of treatment of burns using topical application of egg oil ointment wherein egg oil is extracted by heat treatment of egg.	[36]
3.	US4219585	1980	Nancy I. Herring	An efficient simplified process for extracting the maximum amount of unadulterated, sterilized and medically pure egg oil from egg yolks by use only of heat.	[52]
4.	US 4364930 A	1982	Jacqueline Griat, et al	Composition in the form of stable emulsion for cosmetic ns pharmaceutical use composed of (1) mono- or di-alkyl carboxylates of α - methyl glucoside, or a mixture thereof and (2) mono- or di-alkyl carboxylates of α -methyl glucoside polyoxyethylenated with 10-30 moles of ethylene oxide, or a mixture thereof, the said alkyl moiety of each being linear or branched and having from 11- 21 carbon atoms, (ii) lecithin, (iii) egg yolk oil, and (iv) a polyacrylic type polymer.	[34]
5.	US 5028449	1991	Hatanaka Hiroji H	Egg oil from dried yolks, dietary supplement.	[52]

6.	WO 1991003946 A1	1991	Jeong S Sim	A novel process for the separation of fresh liquid egg yolk into a yolk protein fraction, neutral egg oil fraction and an egg lecithin fraction.	[32]
7.	WO199201271 9 A1	1992	Mohamed Abdessala m Yacobi	Method of treatment of psoriasis using an egg oil product wherein egg of chicken turtle duck etc have been used for oil extraction.	[41]
8.	US5670175 A	1997	Warner C. Nawrocki	A process for the preparation of an ultrapure egg oil which can be obtained from avian or reptilian egg yolk by solvent extraction method.	[30]







Figure 2: Structure of Carotenoids in egg oil



Figure 3: Solvent Extraction of egg oil

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