

Available online on 15.06.2024 at <http://ajprd.com>

Asian Journal of Pharmaceutical Research and Development

Open Access to Pharmaceutical and Medical Research

© 2013-24, publisher and licensee AJPRD, This is an Open Access article which permits unrestricted non-commercial use, provided the original work is properly cited

Open  Access

Review Article

Skin Aging and Herbal Anti-Aging Serums: A Comprehensive Review

Shintre Aniruddha Dileep* Naikawadi Nishigandha Dayanand, Dr. V.K. Redasani

YSPM'S, Yashoda Technical Campus, Faculty of Pharmacy, Wadhe, Satara

ABSTRACT

This review explores the intricate anatomy of the skin, giving a brief synopsis of its structure and role. It describes the various aspects of skin aging and the roles that both intrinsic and extrinsic factors play in this process of aging. Focusing on the topic of herbal medicines, the article investigates the scientifically proven anti-aging properties of a variety of herbs. It also provides information about the composition and varieties of serums, emphasizing important components such as hyaluronic acid, vitamin C, vitamin B3, and herbal extracts that are effective in preventing facial ageing. The aim of this review is to provide readers with a comprehensive understanding of skin aging and the wide range of anti-aging techniques accessible, especially in the context of herbal based face serum

Key words: Skin, Ageing, serums, anti-ageing, herbal extracts, Facial serum.

ARTICLE INFO: Received 10 Jan 2024; Review Complete 15 May 2024; Accepted 05 June 2024 ; Available online 15 June. 2024



Cite this article as:

Dileep SA, Dayanand NN, Redasani VK, Skin Aging and Herbal Anti-Aging Serums: A Comprehensive Review, Asian Journal of Pharmaceutical Research and Development. 2024; 12(3):208-213, DOI: <http://dx.doi.org/10.22270/ajprd.v12i3.1415>

*Address for Correspondence:

Shintre Aniruddha Dileep, YSPM'S, Yashoda Technical Campus, Faculty of Pharmacy, Wadhe, Satara

INTRODUCTION:

The skin serves as a barrier to protect the body's internal organs from the outside world^[1]. The epidermis, dermis, and subcutaneous tissue are the three layers that make up the skin^[2]. The outer layer, known as the epidermis, is made up of a stratified, squamous epithelium that is mostly made up of keratinocytes and dendritic cells, which include Merkel, Langerhans, and melanocytes^[3].

Skin structure and physiological function gradually deteriorate as a result of the complex, multifaceted phenomena known as aging, which is caused by both intrinsic and extrinsic parallel processes^[3]. A person's smooth, pale, drier, less elastic skin with fine wrinkles is the hallmark of intrinsic skin aging, which is only noticeable in old age^[5].

It is generally acknowledged that reactive oxygen species (ROS) and free radical reactions generate cumulative damage that leads to intrinsic aging^[6]. A number of conditions, including ionizing radiation, extreme physical and mental stress, alcohol consumption, poor nutrition, overindulgence in food, pollution from the environment, and exposure to ultraviolet radiation (UVR), can cause extrinsic skin damage^[7].

It is possible that topical skin therapies could slow down these changes in the skin and possibly even return it to its prior quality^[8]. Since ancient times, the herbal preparations have been utilized to treat a variety of skin conditions due to their many therapeutic qualities. A harmonious synergy of bioactive components found in herbal substances, as opposed to synthetic ones, work together to enhance skin health. A number of herbal products are receiving a lot of attention in the skincare sector due to their discovery of therapeutic potential^[9]. Among the constituents in the facial serum is a neuropeptide, which has been linked to improved barrier function and a reduction in the appearance of fine lines and wrinkles^[10].

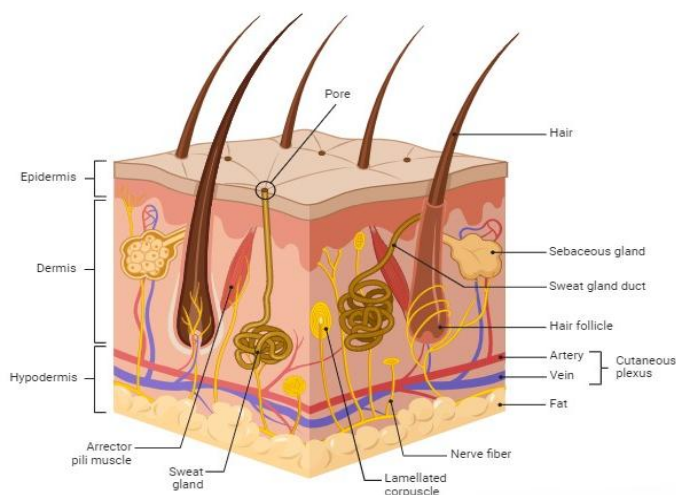
An essential factor in the aging of the skin is antioxidants. They are naturally occurring radical scavengers. Antioxidants are extremely beneficial for both preventing and improving aging and UV-damaged skin^[11]. previous researchers had developed an anti-aging product using natural ingredients. A cosmetic preparation, serum offers the benefit of having a high concentration of active ingredient^[12]. Plant-based active chemicals, such as flavonoids are commonly employed in topical preparations due to their antioxidant capabilities^[13].

The synthesis of anti-aging serum from plant extract has been studied in the past. Thus This review offers an update on skin

anatomy, a brief synopsis of skin aging that covers both intrinsic and extrinsic aging, and a list of various anti-aging herbs. It also explores an overview of serum, different kinds of serums used in skincare routines to prevent ageing, and essential components found in antiaging face serum.

Anatomy of skin:

In terms of both structure and function, the skin is a complicated organ. It assists by guarding against harmful germs, decreasing chemical penetration, and limiting losses of water and electrolytes. The epidermis, dermis, and hypodermis are the three layers that comprise the skin^[14].



Structure of skin

1. Epidermis:

The epidermis, the outermost layer of skin, varies in thickness from 0.5 mm for the eyelid to 1.5 mm for the palms and soles. It is made up primarily of melanocytes and keratinocytes, which combine to form a binary system, and is created by a stratified squamous epithelial layer.^[15]

2. Dermis :

The dermis is a thick layer of elastic, fibrous tissue that lies underneath the epidermis. The layer of connective tissue that sits between the subcutaneous and epidermis is called the dermis. The dermis is composed of ground substance, fibers, and cells since it is connective tissue. Fibroblasts are the cells that make collagen and elastin fibers. Up to 30% of the dermal

volume and 75% of the dry weight are made up of collagen.^{[15][16]}

3. Hypodermis:

The innermost, or subcutaneous, layer of skin is called the hypodermis. The hypodermis shields the skin, provides a reserve energy source, and permits mobility by gliding over underlying components. Adipocytes are the main constituents of the hypodermis, and they are arranged into lobules that are characterized by fibrous connective tissue^[15].

Overview on skin Aging

The main signs of aging skin include the appearance of wrinkles, uneven pigmentation, darkening, thinning, sagging, and roughening of the skin. Either intrinsic or extrinsic mechanisms could be responsible for this^[17]. There are numerous theories that attempt to explain the aging process, but the most likely ones center on DNA damage and the accompanying repair process, which cause epigenetic changes throughout the genome that cause cell senescence, loss of proper cell function, and genomic aberrations^[18].

Extrinsic aging is brought on by environmental factors like ultraviolet (UV) radiation and pollution and results in the production of reactive oxygen species. Intrinsic aging is linked to programmed aging and cellular senescence, which are caused by endogenous oxidative stress and cellular damage^[19].

Intrinsic skin aging

It is a chronologically-driven physiological transformation process. Essentially, intrinsic skin aging is a condition that only appears in old age and is symbolized by smooth, unblemished, pale, dry, less elastic skin that has fine wrinkles^[20]. Intrinsic aging is characterized by a decline in dermal fibroblast count, which lowers the synthesis of collagen and elastin, especially types I and III collagen, and results in skin thinning, wrinkles, and loss of elasticity.^[21] Skin aging is partly caused by increased expression of matrix metalloproteinases (MMPs), which break down collagen and elastin in the dermal skin layer. Antioxidants such as resveratrol and isoorientin can reduce its expression^[22]. Both intrinsic and extrinsic variables can contribute to skin aging.^[23]



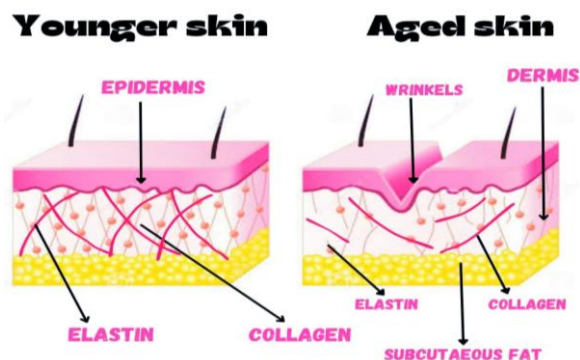
Intrinsic factor responsible for skin ageing

Extrinsic Skin aging

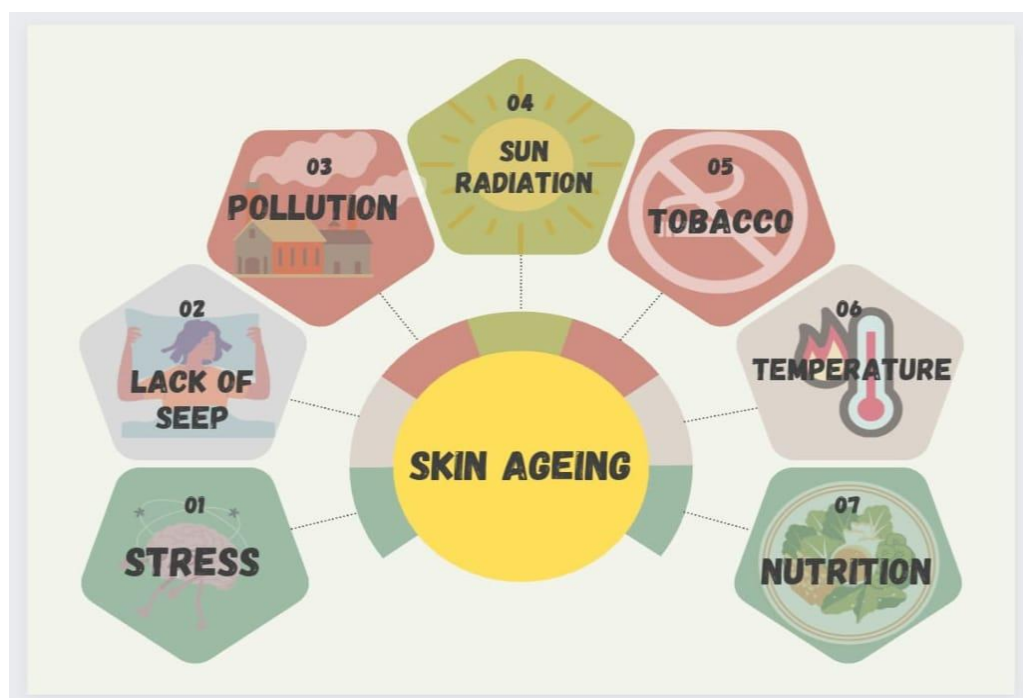
Low-grade chronic UVR exposure is responsible for over 80% of the aging process of facial skin, while exposure can also result in sunburn, tanning, inflammation, and damage to dermal connective tissue. Extrinsically aged skin, primarily from UVR rays, has sallow complexion with uneven pigmentation, rough texture, coarse wrinkles, and decreased skin elasticity^[20].

Accelerated skin aging can be caused independently by long-term exposure to UV radiation and cigarette smoking. ROS produced by UV exposure lead to harmful oxidative stress. Because O₂ can take electrons, it produces extremely reactive chemical compounds known as ROS^[24]. Reactive oxygen species (ROS) are produced when the skin is exposed to photoaging stimuli. These ROS subsequently stimulate dermal enzymes such as collagenase and elastase. These enzymes speed up premature skin aging by breaking down elastin and

collagen, respectively. Signs of this process include deep furrows or severe atrophy, wrinkles, freckles, sallowness, laxity, and a leathery appearance^[25].



Difference between young skin and aged skin



Extrinsic factors responsible for skin ageing

Antiaging Effect of Various Herbs:

In the pursuit of natural solutions for combating aging, traditional herbal medicine has emerged as a promising avenue. Herbs have long been revered for their potential to

promote longevity and vitality, and modern research is increasingly validating these claims. Additionally, there are several herbs have been identified from various selected reports and researches studies showcasing there potential Antiaging activity some of them are mentioned in below table.

Table 1: Several selected reports and research on anti-aging activities of Traditional herbs

Scientific Name	Common name	Part	Solvent	Phytochemicals	Activity	Ref
<i>Epilobium angustifolium L.</i>	Fireweed	stem, leaves and flowers	70% (v/v) ethanol	Phenoloic acid, Gallic acid,	The antioxidant, anti-ageing, anti-inflammatory	[26]
<i>Coffea arabica</i>	Coffee plant	leaves	Methanol	chlorogenic acid, caffeic acid, phenols	prevention of photoaging	[27]
<i>Azadirachta indica A.</i>	Neem	Leaves	50 % ethanol	Rutin	preventing wrinkle formation, Antiaging	[28]
<i>Hibiscus sabdariffa</i>	Rosella	calyx	ethanol	Hibiscus acid	Antiaging	[29]
<i>Ocimum sanctum Linn.</i>	Queen of herb	Whole plant	Hexane	propanoic acid, apigenin, cirsimaritin,	Antiaging antioxidant	[30]

				isothymusin, isothymonin, carnosic acid, rosmarinic acid		
<i>Glycine max</i>	Black Soyabean	-	70% Ethanol	Daidzein flavonoids, isoflavones.	Antioxidant, Antiaging.	[31]
<i>Syzygiumjambos L</i>	Jamun	Leaves	Distilled water	Malic acid, Citric acid Gallic acid Kaempferol Quercetin	Antiaging anti-biofilm, antioxidant	[32]
<i>Silybum marianum (L.)</i>	milk thistle	Fruit	Aqueou-s Ethanol	silybins A isosilybins A Taxifolin	Antiaging Antioxidant	[33]
<i>Centella asiatica</i>	Gotu kola	-	90% Ethanol	Flavonoids Eugenol triterpenoids	Antiaging	[34]
<i>Morus alba</i>	Mullberry	fruit	50% Ethanol	flavonoids phenolic acids and anthocyanins	Antiaging, Antioxidant	[35]
<i>Ginkgo biloba</i>	ginkgo	Leaves	95% ethanol	Bilobalide Ginkgolide quercetin and kaempferol	Antiaging	[36]
<i>Rosa damascena</i>	Rose	petals	70% Ethanol	Flavonoids, terpenoids, phenolics, tannins, and alkaloids	Antiaging, anti-hyaluronidase.	[37]
<i>Calendula officinalis</i>	Marigold	flower	Water/Ethanol extract	Quercetin, Flavonoids	Protect against oxidative stress, Antioxidant	[38]
<i>Mangifera indica L</i>	Mango	Leaves	70% methanol	1,3,5,6-tetrahydroxyxanthone-C-4-β-D-glucopyranoside	Antiaging, anti-collagenase, anti-elastase, anti-hyaluronidase	[39]
<i>Hylocereus polyrhizus</i>	red pitaya	Fruit Peel	82% ethanol	—	Antiaging	[40]
<i>Moringa olifera</i>	Moringa	Leaves	90% ethanol	Apigenin 8-C-glucoside, Quinic acid, 3-Caffeoylquinic acid.	Antiaging, Anti wrinkle	[41]
<i>Crocus sativus. L</i>	Saffron	Stigmas	80% Ethanol	—	Antioxidant, antiaging	[42]
<i>Camellia sinensis</i>	green tea	seeds	70% ethanol	—	Prevention of skin photoaging and wrinkle formation	[43]

Serums:

In order to be faster and more effective than cream treatments, serum, also known as concentrate, contains 10 times more biologically active ingredients. Having the capacity to reach the skin's deeper layers, serum absorbs quickly. More bioactive ingredients are found in serum. Because serum has a lower viscosity than other moisturizers, it has the advantage of being more comfortable to apply and spreading more easily over the skin's surface^[12].

Face serums are composed of minuscule molecules that facilitate rapid and deep skin penetration. Numerous beneficial nutrients and active ingredients, including ceramides, amino acids, and antioxidants, are abundant in serum. The facial serum has a number of components that are linked to a reduction in the visibility of fine lines.[44].

Antioxidant serums are a type of preparation that works to stop reactive oxygen species from oxidatively damaging tissues and cells. Wrinkles and fine lines can be prevented or minimized with the use of anti-aging serums^[45].

Different Types of serum:

Oil type: Among the most basic face serums are oil serums. These oils provide skin-metabolizable substances like polyphenols and vital fatty acids, as well as moisturizing and barrier-repairing properties. They also include high-performing oil-soluble extracts and fragrance mixes containing healthy aromatic compounds. Oil serums fortify and moisturize the skin's protective layers.

The gel serum: Gel serums provide the skin a momentary tightening and lifting sensation. They use formulations that are based on water, which makes it possible to add hydrophilic botanical extracts. Glycerites and powdered botanicals can be used to color or make transparent gels.

Emulsion type: It is appropriate for preparations with high levels of oily components and UV absorber because this type has a high emollient content. In situations when repellent nature is required, the w/o type is appropriate.

Transparent Or semi – transparent Lotion type: often has a higher humectant content than lotion. A humectant and a

water-soluble polymer can be chosen, and their combination can be changed, to alter the texture^[46].

Key Ingredients used in Antiaging face serum

Vitamin C: The most prevalent antioxidant in the skin is L-ascorbic acid, which is also the form of vitamin C that is physiologically active. Vitamin C is believed to have anti-aging properties and is necessary for the synthesis of collagen.^[47] It has been demonstrated that vitamin C guards against UV-induced immunosuppression and photoaging. By boosting collagen synthesis, maintaining collagen fiber stability, and reducing collagen breakdown, it also has an antiaging impact^[48]. Due to the lack of the enzyme L-glucono-gamma lactone oxidase, humans cannot produce vitamin C in vivo. Therefore, a balanced diet and enough nutritional intake of vitamin C are essential for human health. Several cosmetic products, including those for sun protection, skin whitening, and anti-aging, have been found to include L-ascorbic acid^[49].

Hyaluronic Acid: One of the most popular macromolecules for treating skin aging is hyaluronic acid^[50]. N-acetylglucosamine and D-glucuronic acid combine to form the disaccharide units that make up hyaluronic acid (HA), a polysaccharide that is a member of the glycosaminoglycan family. With its moisturizing and anti-aging properties, HA has several advantages over other chemicals used in skin regeneration [51]. Collagen and elastin fiber interfaces and their peripheries have been observed to contain HA, which helps to maintain the appropriate conformation of these two proteins. In both cosmetic and nutricosmetic products, HA has emerged as one of the most important components. HA is a component of almost all products with skin-protective, anti-aging, and moisturizing qualities^[52].

Vitamin B3: The amide form of water-soluble vitamin B3 (nicotinic acid, niacin) is called nicotinamide or niacinamide. For many years, nicotinamide has been used in cosmetics to brighten skin tones and delay the aging process of the skin.^[53] Topical niacinamide treatment enhances the surface structure, minimizes wrinkles, and prevents the development of skin cancer. Furthermore, nicotinamide has been shown to promote keratinocyte differentiation and boost the skin's synthesis of ceramides and collagen, which enhances the skin's barrier function and attractiveness^[54]. Strong anti-aging ingredients like niacinamide neutralize free radicals and reactive oxygen species (ROS), shielding skin cells from oxidative harm. In addition to increasing the activity of enzymes like superoxide dismutase and catalase, it modulates enzymes like NADPH oxidase and nitric oxide synthase to lower the formation of reactive oxygen species (ROS). By promoting skin integrity and cellular health, these activities aid in delaying the onset of aging^[55].

Herbal Extracts: It has been documented that a number of plant bioactive compounds interact with cellular signaling pathways that are either directly or indirectly responsible for the aging of the skin. Herbal products can be used topically to act as cosmetics for body care; bioactive molecules affect the skin's biological processes and supply the nutrients required for good skin health^[56]. Due to its considerable effect on skin aging, herbal cosmetics are becoming more and more popular nowadays, not just in Asian countries but also in other countries^[57]. Studies reveal that certain phytochemicals included in these herbs, such as flavonoids and polyphenols,

are the most desirable class of substances due to their beneficial effects on human health and anti-aging properties^[58]. Phenolic substances include ferulic acid, vanillic acid, caffeic acid, chlorogenic acid, and gallic acid. And flavonoids such as epicatechin, rutin, myricetin, naringenin, kaempferol, quercetin, and myricetin. tannins such as ellagic acid and stilbenes such as resveratrol^[59].

REFERENCES:

1. Gragnani A, Mac Cornick S, Chominski V, de Noronha SM, de Noronha SA, Ferreira LM. Review of major theories of skin aging. *Advances in Aging Research*. 2014 Aug 29;2014.
2. Kolarsick PA, Kolarsick MA, Goodwin C. Anatomy and physiology of the skin. *Journal of the Dermatology Nurses' Association*. 2011 Jul 1;3(4):203-13.
3. Gilaberte Y, Prieto-Torres L, Pastushenko I, Juarraz Á. Anatomy and Function of the Skin. In *Nanoscience in dermatology 2016 Jan 1* (pp. 1-14). Academic Press.
4. Farage MA, Miller KW, Elsner P, Maibach HI. Structural characteristics of the aging skin: a review. *Cutaneous and ocular toxicology*. 2007 Jan 1;26(4):343-57.
5. Tobin DJ. Introduction to skin aging. *Journal of tissue viability*. 2017 Feb 1;26(1):37-46.
6. Poljšak B, Dahmane RG, Godić A. Intrinsic skin aging: the role of oxidative stress. *Acta Dermatovenerol Alp Pannonica Adriat*. 2012 Jan 1;21(2):33-6.
7. Poljšak B, Dahmane R. Free radicals and extrinsic skin aging. *Dermatology research and practice*. 2012;2012.
8. West BJ, Alabi I, Deng S. A face serum containing bakuchiol, palmitoyl tripeptide-38, hydrolyzed hyaluronic acid and a polyherbal and vitamin blend improves skin quality in human volunteers and protects skin structure in vitro.
9. Sharma RR, Deep A, Abdullah ST. Herbal products as skincare therapeutic agents against ultraviolet radiation-induced skin disorders. *Journal of Ayurveda and Integrative Medicine*. 2022 Jan 1;13(1):100500.
10. McCall-Perez F, Stephens TJ, Herndon JH Jr. Efficacy and tolerability of a facial serum for fine lines, wrinkles, and photodamaged skin. *J Clin Aesthet Dermatol*. 2011 Jul;4(7):51-4. PMID: 21779421; PMCID: PMC3140905.
11. Uwa LM. The anti-aging efficacy of antioxidants. *Curr Trends Biomed Eng Biosci*. 2017;7:1-3.
12. Mursyid AM, Waris R. Formulation and evaluation of pharmaceutically stable serum of *D. indica* (Arbenan). *Universal Journal of Pharmaceutical Research*. 2021;6(1):38-42.
13. M Calderon-Montano J, Burgos-Morón E, Pérez-Guerrero C, López-Lázaro M. A review on the dietary flavonoid kaempferol. *Mini reviews in medicinal chemistry*. 2011 Apr 1;11(4):298-344.
14. Khavkin J, Ellis DA. Aging skin: histology, physiology, and pathology. *Facial Plastic Surgery Clinics*. 2011 May 1;19(2):229-34.
15. Gilaberte Y, Prieto-Torres L, Pastushenko I, Juarraz Á. Anatomy and Function of the Skin. In *Nanoscience in dermatology 2016 Jan 1* (pp. 1-14).
16. Venus M, Waterman J, McNab I. Basic physiology of the skin. *Surgery (Oxford)*. 2010 Oct 1;28(10):469-72.
17. Lephart ED. Skin aging and oxidative stress: Equol's anti-aging effects via biochemical and molecular mechanisms. *Ageing research reviews*. 2016 Nov 1;31:36-54.
18. Sinclair DA, Oberdoerffer P. The ageing epigenome: damaged beyond repair? *Ageing research reviews*. 2009 Jul 1;8(3):189-98.
19. Shin SH, Lee YH, Rho NK, Park KY. Skin aging from mechanisms to interventions: focusing on dermal aging. *Front Physiol*. 2023 May 10;14:1195272. doi: 10.3389/fphys.2023.1195272. PMID: 37234413; PMCID: PMC10206231.
20. Tobin DJ. Introduction to skin aging. *Journal of tissue viability*. 2017 Feb 1;26(1):37-46.
21. Fisher GJ, Kang S, Varani J, Bata-Csorgo Z, Wan Y, Datta S, Voorhees JJ. Mechanisms of photoaging and chronological skin aging. *Archives of dermatology*. 2002 Nov 1;138(11):1462-70.
22. Lephart ED. Skin aging and oxidative stress: Equol's anti-aging effects via biochemical and molecular mechanisms. *Ageing research reviews*. 2016 Nov 1;31:36-54.
23. Zheng H, Zhang M, Luo H, Li H. Isoorientin alleviates UVB-induced skin injury by regulating mitochondrial ROS and cellular autophagy. *Biochemical and biophysical research communications*. 2019 Jul 5;514(4):1133-9.

24. Juan CA, Pérez de la Lastra JM, Plou FJ, Pérez-Lebeña E. The chemistry of reactive oxygen species (ROS) revisited: outlining their role in biological macromolecules (DNA, lipids and proteins) and induced pathologies. *International journal of molecular sciences*. 2021 Apr 28;22(9):4642.
25. Jiratchayamaethasakul C, Ding Y, Hwang O, Im ST, Jang Y, Myung SW, Lee JM, Kim HS, Ko SC, Lee SH. In vitro screening of elastase, collagenase, hyaluronidase, and tyrosinase inhibitory and antioxidant activities of 22 halophyte plant extracts for novel cosmeceuticals. *Fisheries and Aquatic Sciences*. 2020 Dec;23:1-9.
26. Nowak A, Zagórska-Dziok M, Perużyńska M, Cybulska K, Kucharska E, Ossowicz-Rupniewska P, Piotrowska K, Duchnik W, Kucharski Ł, Sulikowski T, Drożdżik M. Assessment of the anti-inflammatory, antibacterial and anti-aging properties and possible use on the skin of hydrogels containing *Epilobium angustifolium* L. Extracts. *Frontiers in Pharmacology*. 2022 Jul 1;13:896706.
27. Chiang HM, Lin TJ, Chiu CY, Chang CW, Hsu KC, Fan PC, Wen KC. *Coffea arabica* extract and its constituents prevent photoaging by suppressing MMPs expression and MAP kinase pathway. *Food and Chemical Toxicology*. 2011 Jan 1;49(1):309-18.
28. Ngo HT, Hwang E, Seo SA, Park B, Sun ZW, Zhang M, Shin YK, Yi TH. Topical application of neem leaves prevents wrinkles formation in UVB-exposed hairless mice. *Journal of Photochemistry and Photobiology B: Biology*. 2017 Apr 1;169:161-70.
29. Wang D, Nagata M, Matsumoto M, Amen Y, Wang D, Shimizu K. Potential of *Hibiscus sabdariffa* L. and hibiscus acid to reverse skin aging. *Molecules*. 2022 Sep 17;27(18):6076.
30. Chaiyana W, Anuchapreeda S, Punyoyai C, Neimkhum W, Lee KH, Lin WC, Lue SC, Viernstein H, Mueller M. *Ocimum sanctum* Linn. as a natural source of skin anti-ageing compounds. *Industrial crops and products*. 2019 Jan 1;127:217-24.
31. Juliana C, Lister IN, Girsang E, Nasution AN, Widowati W. Antioxidant and elastase inhibitor from black soybean (*Glycine max* L.) and its compound (daidzein). *Journal of Biomedicine and Translational Research*. 2020 Apr 30;6(1):11-4.
32. Mahdi I, Imbimbo P, Ortaakarsu AB, Adhiambo Ochieng M, Ben Bakrim W, Drissi BE, Ibrahim MA, Abdelfattah MA, Mahmoud MF, Monti DM, Sobeh M. Chemical profiling and dermatological and anti-aging properties of *Syzygiumjambos* L.(Alston): evidence from molecular docking, molecular dynamics, and in vitro experiments. *Frontiers in Molecular Biosciences*. 2024 Jan 5;10:1331059.
33. Drouet S, Leclerc EA, Garros L, Tungmunthum D, Kabra A, Abbasi BH, Lainé É, Hano C. A green ultrasound-assisted extraction optimization of the natural antioxidant and anti-aging flavonolignans from milk thistle *Silybum marianum* (L.) Gaertn. fruits for cosmetic applications. *Antioxidants*. 2019 Aug 14;8(8):304.
34. Karsono AH, Tandrasasmita OM, Berlian G, Tjandrawinata RR. Potential antiaging effects of DLBS1649, a centella asiatica bioactive extract. *Journal of experimental pharmacology*. 2021 Aug 11:781-95.
35. Jiang DQ, Guo Y, Xu DH, Huang YS, Yuan K, Lv ZQ. Antioxidant and anti-fatigue effects of anthocyanins of mulberry juice purification (MJP) and mulberry marc purification (MMP) from different varieties mulberry fruit in China. *Food and chemical toxicology*. 2013 Sep 1;59:1-7.
36. Wang X, Gong X, Zhang H, Zhu W, Jiang Z, Shi Y, Li L. In vitro anti-aging activities of ginkgo biloba leaf extract and its chemical constituents. *Food Science and Technology*. 2020 Feb 3;40:476-82.
37. Dewi DY, Ginting CN, Chiuman L, Girsang E, Handayani RA, Widowati W. Potentials of rose (*Rosa damascena*) petals and receptacles extract as antioxidant and antihyaluronidase. *Pharmaciana*. 2020;10:343-52.
38. Alnuqaydan AM, Lenehan CE, Hughes RR, Sanderson BJ. Extracts from *Calendula officinalis* offer in vitro protection against H₂O₂ induced oxidative stress cell killing of human skin cells. *Phytotherapy Research*. 2015 Jan;29(1):120-4.
39. El-Nashar HA, El-Labbad EM, Al-Azzawi MA, Ashmawy NS. A new xanthone glycoside from *Mangifera indica* L.: Physicochemical properties and in vitro anti-skin aging activities. *Molecules*. 2022 Apr 19;27(9):2609.
40. Vijayakumar RA, Gani SS, Mokhtar N. Anti-elastase, anti-collagenase and antimicrobial activities of the underutilized red pitaya peel: An in vitro study for anti-aging applications. *Asian J. Pharm. Clin. Res*. 2017;10(8):251-5.
41. Xu Y, Chen G, Guo M. Potential anti-aging components from *Moringa oleifera* leaves explored by affinity ultrafiltration with multiple drug targets. *Frontiers in Nutrition*. 2022 May 10;9:854882.
42. Samarghandian S, Asadi-Samani M, Farkhondeh T, Bahmani M. Assessment the effect of saffron ethanolic extract (*Crocus sativus* L.) on oxidative damages in aged male rat liver. *Der Pharm Lett*. 2016;8(3):283-90.
43. Lim JY, Kim OK, Lee J, Lee MJ, Kang N, Hwang JK. Protective effect of the standardized green tea seed extract on UVB-induced skin photoaging in hairless mice. *Nutrition research and practice*. 2014 Aug;8(4):398.
44. McCall-Perez F, Stephens TJ, Herndon JH Jr. Efficacy and tolerability of a facial serum for fine lines, wrinkles, and photodamaged skin. *J Clin Aesthet Dermatol*. 2011 Jul;4(7):51-4. PMID: 21779421; PMCID: PMC3140905.
45. Kamble R, Gamare D, Yeole P, Pathan MA, Indulkar A, Kale MK, Juvatkar PV. FORMULATION AND EVALUATION OF HERBAL BASED ANTI-AGING FACE SERUM. *Foldscape & its Applications*. 2022;127.
46. Shejul TS, Kudale K. Facial Serum: Its Formulation, Usage, Special Ingredients, Various Types and Benefits. *Int J Pharm Res Appl*. 2023 Mar-Apr;8(2):680-692.
47. Rattanawitpong P, Wanitphakdeedecha R, Bumrungpert A, Maiprasert M. Anti-aging and brightening effects of a topical treatment containing vitamin C, vitamin E, and raspberry leaf cell culture extract: a split-face, randomized controlled trial. *Journal of cosmetic dermatology*. 2020 Mar;19(3):671-6.
48. Al-Niaimi F, Chiang NYZ. Topical Vitamin C and the Skin: Mechanisms of Action and Clinical Applications. *J Clin Aesthet Dermatol*. 2017 Jul;10(7):14-17. Epub 2017 Jul 1. PMID: 29104718; PMCID: PMC5605218.
49. Mumtaz S, Ali S, Tahir HM, Kazmi SA, Shakir HA, Mughal TA, Mumtaz S, Summer M, Farooq MA. Aging and its treatment with vitamin C: a comprehensive mechanistic review. *Molecular biology reports*. 2021 Dec 1:1-3.
50. Galvez-Martin P, Soto-Fernandez C, Romero-Rueda J, Cabañas J, Torrent A, Castells G, Martinez-Puig D. A novel hyaluronic acid matrix ingredient with regenerative, anti-aging and antioxidant capacity. *International journal of molecular sciences*. 2023 Mar 1;24(5):4774.
51. Juncan AM, Moisés DG, Santini A, Morgovan C, Rus LL, Vonica-Tincu AL, Loghin F. Advantages of hyaluronic acid and its combination with other bioactive ingredients in cosmeceuticals. *Molecules*. 2021 Jul 22;26(15):4429.
52. Bukhari SN, Roswandi NL, Waqas M, Habib H, Hussain F, Khan S, Sohail M, Ramli NA, Thu HE, Hussain Z. Hyaluronic acid, a promising skin rejuvenating biomedicine: A review of recent updates and pre-clinical and clinical investigations on cosmetic and nutraceutical effects. *International journal of biological macromolecules*. 2018 Dec 1;120:1682-95.
53. Boo YC. Mechanistic basis and clinical evidence for the applications of nicotinamide (niacinamide) to control skin aging and pigmentation. *Antioxidants*. 2021 Aug 21;10(8):1315.
54. Mohiuddin A. Skin aging and modern age anti—Aging strategies. *Int. J. Clin. Dermatol. Res*. 2019 Jul 23;7:209-40.
55. Marques C, Hadjab F, Porcello A, Lourenço K, Scaletta C, Abdel-Sayed P, Hirt-Burri N, Applegate LA, Laurent A. Mechanistic Insights into the Multiple Functions of Niacinamide: Therapeutic Implications and Cosmeceutical Applications in Functional Skincare Products. *Antioxidants*. 2024 Mar 30;13(4):425.
56. Mukherjee PK, Bahadur S, Chaudhary SK, Harwansh RK, Nema NK. Validation of medicinal herbs for skin aging. In *Evidence-Based Validation of Herbal Medicine* 2015 Jan 1 (pp. 119-147). Elsevier.
57. Mukherjee PK, Maity N, Nema NK, Sarkar BK. Bioactive compounds from natural resources against skin aging. *Phytomedicine*. 2011 Dec 15;19(1):64-73.
58. Xue F, Li X, Qin L, Liu X, Li C, Adhikari B. Anti-aging properties of phytoconstituents and phyto-nanoemulsions and their application in managing aging-related diseases. *Advanced Drug Delivery Reviews*. 2021 Sep 1;176:113886.
59. Dhalaria R, Verma R, Kumar D, Puri S, Tapwal A, Kumar V, Nepovimova E, Kuca K. Bioactive compounds of edible fruits with their anti-aging properties: A comprehensive review to prolong human life. *Antioxidants*. 2020 Nov 13;9(11):1123.