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Skin Health and Nutrition: Separating the Science from the Trends

TODAY'S AGENDA:

- Introduction & Housekeeping
- Speaker Introduction
- Presentation
- Q&A
- Closing





WEBINAR HOST:

Keith Hine, MS, RD VP of Healthcare, Sports & Professional Education Orgain, LLC

WEBINAR PRESENTER:

Nancy Collins, PhD, RDN, LD, NWCC, FAND Nutrition Wound Care Certified dietitian (NWCC) Light Bulb Health, Inc. www.drnancycollins.com

Skin Concerns Over the Lifespan: Wrinkles to Wounds







Objectives/ Disclosures



Disclosures:

Consultant, Speaker's Bureau – Medtrition, Abbott Nutrition, Nutricia This webinar is sponsored by Orgain

Skin – Our Body's Coat

- Skin is our largest organ
 - adults carry about 8 pounds and 22 square feet of it
- Makes us look presentable
- Without it, we would literally evaporate



Functions of Skin



- Waterproofing
- Insulation and temperature regulation
- Manufactures vitamin D
- Sensor packed with nerves
 - Pain and pleasure
- Elastic and permits movement
- Protects against sunlight

Merck Manual. The Structure and Function of Skin. Available at: https://www.merckmanuals.com/home/skindisorders/biology-of-the-skin/structure-and-function-of-the-skin

The Layers of Skin



Epidermis – protective layer of dead cells

Dermis – sensation, temperature regulation

Subcutaneous – cushioning, insulation

Microcirculation of the Skin

- Arterioles deliver oxygen, nutrients
- Venules carry away cellular waste
- External pressure occludes arterioles and venules
 - e.g., the kink in a garden hose



Liao F, Burns S, Jan YK. Skin blood flow dynamics and its role in pressure ulcers. *J Tissue Viability*. 2013;22(2):25-36.

Skin Microbiome

- Supports a diverse community of microorganisms that train and support the immune system
- Fends off pathogenic threats
- Extensive communication between bacteria, skin cells and immune cells
- When disturbed, skin is susceptible to immune hypersensitivity disorders, such as eczema and skin allergies, or interfere with healing in people with chronic wounds





Causes of Skin Aging

- Intrinsic = Time + Genetics
- Extrinsic = Everything else
 - Pollution
 - Smoking
 - Climate/hydration
 - UV exposure/radiation
 - Sleep
 - Exercise
 - Stress- catecholamines and cortisol
 - Skin Care topical, routines
 - Nutrition/ supplements



4 main causes of aging

Aging of Skin Tissue



Intact Collagen Normal Fibroblasts Size



ROS= Reactive Oxygen Species

ROS is dangerous oxygen molecules generated by UV rays and pollution.

ROS attack and react with stable skin cell molecules, causing cross-linking of collagen and elastin while lessening skin's ability to repair itself.

Quan T, Fisher GJ. Role of Age-Associated Alterations of the Dermal Extracellular Matrix Microenvironment in Human Skin Aging: A Mini-Review. Gerontology, 2015, 61:427–434.



Collagen

- From the Greek word kólla, meaning "glue" and -gen, meaning "producing"
- Main structural protein in the extracellular matrix (ECM) in the various connective tissues in the body
- The most abundant protein in mammals
- Makes up approximately 25% to 35% of the whole-body protein content
- The fibroblast is the most common cell that creates collagen





Collagen Chemistry

- Composed of a triple helix
 - Two identical chains and an additional chain that differs slightly in its chemical composition
- The amino acid composition of collagen is atypical for proteins
 - Every third amino acid is a glycine molecule, according to the formula GLY-X-Y
 - Proline and hydroxyproline makes up about 23% of amino acid content of collagen
 - Proline 📫 Hydroxyroline 📫 Collagen
 - this hydroxylation is vitamin
 - Co-factor for this hydroxylation is vitamin C





Albaugh VL, Mukherjee K, Barbul A. Proline Precursors and Collagen Synthesis: Biochemical Challenges of Nutrient Supplementation and Wound Healing. *J Nutr*. 2017;147(11):2011-2017. doi:10.3945/jn.117.256404

Organization of Collagen



Individual α-chains are woven into triple helices via a zipper mechanism.

Bundles of triple helices form fibrils and these fibrils are aggregated into larger fibers.

Reilly DM, Lozano J. Skin collagen through the lifestages: importance for skin health and beauty. *Plastic and Aesthetic Research*. 2021; 8: 2. <u>http://dx.doi.org/10.20517/2347-9264.2020.153</u>. Available at <u>https://parjournal.net/article/view/3863</u>.

Types of Collagen¹

Туре	Function
Туре І	 Skin, tendon, vasculature, organs, bone (main component of the organic part of bone) 70% of the collagen in the human skin is Type I²
Type II	Cartilage
Type III	Reticulate (main component of reticular fibers)
Type IV	Forms basal lamina, the epithelium-secreted layer of the basement membrane
Type V	Cell surfaces, hair, and placenta

1. Ricard-Blum, S. (2011). The Collagen Family. Cold Spring Harbor Perspectives in Biology. 3 (1): a004978.

doi:10.1101/cshperspect.a004978. PMC 3003457. PMID 21421911.

2. Rangaraj A, Harding K, Leaper D. Role of collagen in wound management. Wounds. 7(2) June 2011.

Collagen and Aging



The Big Question



Can consuming collagen supplements improve skin from wrinkles to wounds?

Protein Requirements

- RDA = 0.8 g/kg
- Based on needs for healthy people
- May be too low for older adults

Aside from aging, the top reason people don't have enough collagen is poor diet

Your body can't make collagen if it doesn't have the necessary elements

National Research Council (US) Subcommittee on the Tenth Edition of the Recommended Dietary Allowances. Recommended Dietary Allowances: 10th Edition. Washington (DC): National Academies Press (US); 1989. 6, Protein and Amino Acids. Available from: https://www.ncbi.nlm.nih.gov/books/NBK234922/

Recommended Protein Intake for Pressure Injuries

1.25 to 1.5 grams protein/kg body weight

....for an adult at risk of a pressure injury or with an existing pressure injury who is assessed to be at risk of malnutrition when compatible with goals of care, and reassess as condition changes...





European Pressure Ulcer Advisory Panel, National Pressure Injury Advisory Panel, Pan Pacific Pressure Injury Alliance. *Prevention and Treatment of Pressure Ulcers/Injuries*. Haesler E, ed. 3rd ed. EPUAP/NPIAP/PPPIA; 2019.

Recommended Dietary Sources of Collagen

Bone broth

• Draws collagen out of beef, chicken or fish bones

• Protein rich foods

- Beef
- Chicken
- Fish
- Beans
- Eggs
- Dairy products (milk, cheese)

• Vitamin C rich foods

- Citrus fruits (oranges, grapefruits)
- Red and green peppers
- Tomatoes
- Dark green, leafy vegetables (broccoli, kale, Brussels sprouts)
- Zinc and copper-rich foods
 - Meats
 - Shellfish
 - Nuts
 - Whole grains
 - Beans

Collagen Synthesis



Evaluating Collagen Supplements

- Type bovine, porcine, marine
- Source pesticides, contaminants, husbandry concerns
- Form powder, liquid
- Cost
- Taste and palatability
- Amino acid profile
- Bioavailability



Hydrolyzed Collagen

- Denatured collagen forms gelatin
- Enzymatic hydrolysis results in collagen hydrolysates (CH)
- CH: collagen broken down into small protein chains
 - collagen peptides
- CH is soluble in water at ambient temperature due to low molecular weight
 - This allows product development



Effects of hydrolyzed collagen supplementation on skin aging: a systematic review and meta-analysis



- Medline, Embase, Cochrane, LILACS (Latin American and Caribbean Health Sciences Literature), and Journal of Negative Results in BioMedicine databases
- Randomized, double-blind, and controlled trials that evaluated oral supplementation with hydrolyzed collagen as an intervention and reported at least one of the following outcomes: skin wrinkles, hydration, elasticity, and firmness
- 19 studies were selected
 - 1,125 participants
 - Aged between 20 and 70 years
 - 95% women

de Miranda RB, Weimer P, Rossi RC. Effects of hydrolyzed collagen supplementation on skin aging: a systematic review and meta-analysis. Int J Dermatol. 2021 Dec;60(12):1449-1461. doi: 10.1111/ijd.15518. Epub 2021 Mar 20. PMID: 33742704.

The Meta Results

- Favorable results of CH supplementation compared with placebo for
 - \checkmark skin hydration
 - ✓ elasticity
 - ✓ wrinkles
- Based on results, ingestion of hydrolyzed collagen for 90 days is effective in reducing skin aging



de Miranda RB, Weimer P, Rossi RC. Effects of hydrolyzed collagen supplementation on skin aging: a systematic review and meta-analysis. Int J Dermatol. 2021 Dec;60(12):1449-1461. doi: 10.1111/ijd.15518. Epub 2021 Mar 20. PMID: 33742704.



Digestion of Proteins



Collagen Di-Peptides Specific to Wound Healing

- Readily absorbed through the wall of small intestine
- PO is a low molecular weight fibroblast-initiating factor
- Enhances wound healing by stimulating the growth of p75NTRpositive fibroblasts
- Promotes hyaluronic acid synthesis, required for maintaining dermal integrity

Pro-Hyp prolyl-hydroxyproline Hyp-Gly Hydroxyprolyl-Glycine

Not easily degraded

- PO is twisted
- OG is stacked

Sato K, Asai TT, Jimi S. Collagen-Derived Di-Peptide, Prolylhydroxyproline (Pro-Hyp): A New Low Molecular Weight Growth-Initiating Factor for Specific Fibroblasts Associated With Wound Healing. Front. Cell Dev. Biol.2020. 8:548975.doi: 10.3389/fcell.2020.548975

WOUND HEALING



✓ Epithelial cells migrate from the wound margins



Rodrigues M, Kosaric N, Bonham CA, and Gurtner GC. Wound Healing: A Cellular Perspective Nov 26 2018 https://doi.org/10.1152/physrev.00067.2017

SCIENTIFIC REPORTS

Received: 2 February 2018 Accepted: 18 July 2018 Published online: 30 July 2018

OPEN Ingestion of bioactive collagen hydrolysates enhanced pressure ulcer healing in a randomized double-blind placebo-controlled clinical study

Fumihito Sugihara¹, Naoki Inoue¹ & Sriraam Venkateswarathirukumara²

We conducted a double blind, multi-centric, placebo-controlled, randomized trial to compare the Pressure Ulcer Scale for Healing (PUSH) and Pressure Sore Status Tool (PSST) scores and wound area measurements at 16 weeks of subjects with pressure ulcers who were given standard care plus one of two types of collagen hydrolysate (CH-a), which contained low levels of prolyl hydroxyproline (Pro-Hyp) and hydroxyprolyl given (Hyp-Gly), and CH-b, which contained high levels of Pro-Hyp and Hyp-Gly) with the placebo group. A total of 120 subjects with stage II or III pressure ulcers were entered into the trial and 112 subjects completed the study. The subjects were randomized to receive CH-a (n = 39), CH-b (n = 39), or a placebo (n = 42) twice daily (10g per day) for 16 weeks. The PUSH score, PSST score, and wound area of the CH-b group were significantly lower than the placebo group at week 16 (PUSH score, P < 0.001; PSST score, P < 0.01; wound area, P < 0.05). The PUSH score of the CH-a group was significantly lower than the placebo group at week 16 (PLSH ingestion helps healing of pressure ulcers as an add-on to the standard therapy.

Sugihara F, Inoue N & Venkateswarathirukumara S. Ingestion of bioactive collagen hydrolysates enhanced pressure ulcer healing in a randomized double-blind placebo-controlled clinical study. *Sci Rep* 8, 11403 (2018). https://doi.org/10.1038/s41598-018-29831-7

Collagen Dipeptides in Pressure Injury Healing

DESIGN

- 16 week double blind study
- Multi-center
- n = 120 enrolled
- n = 112 completed

INTERVENTION

- Three randomized study groups
 - 1. CH-a (n = 39) Low level of PO and OG
 - 2. CH-b (n = 39) High level of PO and OG
 - 3. Placebo (n = 42)
- Twice daily (10g per day) for 16 weeks

OUTCOME

- Measure
- 1. PUSH score
- 2. PSST score
- 3. Wound area

Results: Pressure Ulcer Scale for Healing (PUSH) Score



CH-a = Ordinary collagen hydrolysate

CH-b = collagen hydrolysate with high concentration of PO and OG

Sugihara F, Inoue N & Venkateswarathirukumara S. Ingestion of bioactive collagen hydrolysates enhanced pressure ulcer healing in a randomized double-blind placebo-controlled clinical study. *Sci Rep* 8, 11403 (2018). https://doi.org/10.1038/s41598-018-29831-7

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"These results suggest that although all CH are derived from similar raw materials, it might be possible to control the healing effects of CH on pressure ulcers by altering their dipeptide content..."

PO and OG on Skin Health



Ingestion of bioactive collagen hydrolysates enhance facial skin moisture and elasticity and reduce facial ageing signs in a randomise double-blind placebo-controlled clinical study

Naoki Inoue,^{a*} Fumihito Sugihara^a and Xuemin Wang^b

Abstract

BACKGROUND: Several human studies have demonstrated occurrence of two major collagen peptides, prolyl-hydroxypr (Pro-Hyp) and hydroxyprolyl-glycine (Hyp-Gly), in human peripheral blood. Some *in vitro* studies have demonstrated Pro-Hyp and Hyp-Gly exert chemotaxis on dermal fibroblasts and enhance cell proliferation. Additionally, Pro-Hyp enhi the production of hyaluronic acid by dermal fibroblasts. These findings suggest that the amounts of Pro-Hyp and Hypblood are important factors to show the efficacy of collagen hydrolysates on skin health.

RESULTS: We conducted a randomised double-blind placebo-controlled clinical trial of ingestion of two types of col hydrolysates, which are composed of different amounts of the bioactive dipeptides Pro-Hyp and Hyp-Gly, to investigate effects on the improvement of skin conditions. Improvement in skin conditions, such as skin moisture, elasticity, wrinkle roughness, were compared with a placebo group at baseline, and 4 and 8 weeks after the start of the trial. In additio safety of dietary supplementation with these peptides was evaluated by blood test. Collagen hydrolysate with a higher co of bioactive collagen peptides (H-CP) showed significant and more improvement than the collagen hydrolysate with a content of bioactive collagen peptides (L-CP) and the placebo, in facial skin moisture, elasticity (R2), wrinkles and roug compared with the placebo group. In addition, there were no adverse events during the trial.

CONCLUSION: This study demonstrated that the use of the collagen hydrolysate with a higher content of Pro-Hyp and H led to more improvement in facial skin conditions, including facial skin moisture, elasticity, wrinkles and roughness. © 2016 Society of Chemical Industry

1. It hash blosting contider this roughout

- Double blind placebo-controlled study
- Three groups (placebo, low content of bioactive collagen peptide, high content of bioactive collagen peptides)
- Assessed at baseline, week 4, and week 8
- n=85 females aged 35 55 yo

Results

Higher concentration of PO and OG improved moisture, elasticity, wrinkles, and roughness

Inoue N, Sugihara F, Wang X. Ingestion of bioactive collagen hydrolysates enhance facial skin moisture and elasticity and reduce facial ageing signs in a randomised double-blind placebo-controlled clinical study. J Sci Food Agric. 2016 Sep;96(12):4077-81. doi: 10.1002/jsfa.7606. Epub 2016 Feb 10. PMID: 26840887.

PO and OG Enhance Cell Proliferation

- Exerts chemotaxis on dermal fibroblasts
- Enhances the production of hyaluronic acid
 - From the Greek word hyalos, meaning glass
 - Associated With smoothness of the skin
- May improve barrier function to improve moisture
- May stimulate the synthesis of Type I collagen
- May be able to enhance uptake by increasing the concentration of bioavailable peptides in products



HYALURONIC ACID

Inoue N, Sugihara F, Wang X. Ingestion of bioactive collagen hydrolysates enhance facial skin moisture and elasticity and reduce facial ageing signs in a randomised double-blind placebo-controlled clinical study. J Sci Food Agric. 2016 Sep;96(12):4077-81. doi: 10.1002/jsfa.7606. Epub 2016 Feb 10. PMID: 26840887.

The Future Will Bring Even More Applications



Abeer MM, Trajkovic S, Brayden DJ. Measuring the oral bioavailability of protein hydrolysates derived from food sources: A critical review of current bioassays. Biomed Pharmacother. 2021 Dec;144:112275. doi: 10.1016/j.biopha.2021.112275. Epub 2021 Oct 7. PMID: 34628165.

Separating the Science from the Trends

- Low molecular weight collagen peptides support skin health from wrinkles to wounds
- \checkmark Decreases breakdown of collagen
- \checkmark Increases production of collagen
- \checkmark Reduces ROS in ECM
- \checkmark Reduces inflammation
- Use your clinical judgment in finding reputable products



Time for Questions, Thoughts, Comments



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