

The Potential Impact of Collagen Supplementation in Healthy Aging and Longevity: A Mini Review

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ABSTRACT

Collagen supplementation has garnered attention as a promising intervention to support healthy aging and longevity. The synthesis of collagen, the body's primary structural protein, decreases by approximately 1% per year after age 25, contributing to an aging phenotype characterized by skin wrinkling, joint degeneration, and impaired tissue integrity. This mini-review critically evaluates the potential benefits of collagen supplementation, exploring its multifaceted mechanisms of action that extend beyond simply providing amino acids. Specifically, exogenous collagen peptides stimulate dermal fibroblasts, increasing the synthesis of elastin and fibrillin, key collagen types. Additionally, recent findings in model organisms highlight specific collagen genes as vital mediators of longevity, suggesting a regulatory role in cellular senescence and oxidative stress pathways prevalent in aging. This review aims to provide a comprehensive overview of the effects of collagen supplementation in the areas of skin, musculoskeletal, and metabolic health by synthesizing contemporary clinical evidence and mechanistic information. Furthermore, addressing safety concerns and research limitations, it questions whether collagen peptides, beyond their superficial benefits, can be proven as a legitimate component of longevity medicine to influence fundamental biological aging processes.

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Introduction

Collagen, the most abundant structural protein in the human body, has emerged as an impressive nutritional intervention to support healthy aging and longevity. The progressive, age-related decline in endogenous collagen synthesis (estimated at approximately 1% per year after age 25) constitutes a key component of the aging phenotype, manifesting as skin wrinkling, joint degeneration, and decreased tissue integrity [1]. Consequently, oral collagen supplementation has garnered significant scientific and commercial interest as a strategy to counteract this decline and support multiple interconnected physiological systems. The primary aim of this intervention is not merely cosmetic; rather, it targets the essential extracellular matrix that supports organ structure, cellular signaling, and overall systemic resistance [2,3].

The mechanisms by which exogenous collagen peptides exert their effects are multifaceted and extend beyond simply providing amino acid building blocks. After ingestion and hydrolysis, bioactive peptides are absorbed and can stimulate dermal fibroblasts, increasing the synthesis of type I and III collagen, elastin, and fibrillin [4,5]. Furthermore, novel evidence from model organisms such as *Caenorhabditis elegans* positions specific collagen genes, such as COL7A1, as critical mediators of longevity, linking collagen metabolism directly to conserved genetic pathways that

govern lifespan [6]. This suggests that collagen's role is not only structural but also regulatory, influencing cellular senescence, oxidative stress response, and inflammatory pathways, which are defining features of aging [7,8].

Therefore, this mini-review synthesizes current evidence to critically evaluate the potential of collagen supplementation as a multifaceted intervention for healthy aging. This study will systematically review the clinical and mechanistic evidence supporting the benefits of collagen peptides on skin, musculoskeletal, and metabolic health, while also addressing their safety profile and the limitations of the current research environment. By integrating findings from randomized controlled trials, meta-analyses, and preliminary preclinical studies, this review aims to provide a comprehensive assessment of whether collagen peptides can play a legitimate role in longevity medicine, going beyond superficial applications by addressing the fundamental biological processes of aging.

Materials and Methods

This mini-review synthesizes data from a comprehensive selection of current literature evaluating the impact of collagen supplementation on aging and longevity. A systematic search was conducted across various scientific databases, including PubMed, ScienceDirect, and Google Scholar, focusing on studies published between 2000 and 2025. The review aimed to include randomized controlled trials (RCTs), meta-analyses,

and preclinical studies examining the physiological effects of oral collagen peptide administration on skin elasticity, joint health, and metabolic function. Key inclusion criteria included middle-aged and elderly participants, with particular emphasis on studies investigating the biological activity of collagen peptides after ingestion and evaluating their safety profiles. Furthermore, to provide a holistic understanding of collagen's role, it was deemed necessary to include research addressing the molecular mechanisms surrounding collagen metabolism and its impact on age-related pathways such as oxidative stress response and inflammatory mediators. A narrative synthesis method was used to examine the data and bring the findings together. The main focus was on how collagen supplements could be used in interventions aimed at helping people live healthier and longer lives.

Discussion

The therapeutic potential of collagen supplementation rests on a complex interplay of direct and indirect mechanisms that work together to mitigate age-related tissue degeneration. Firstly, hydrolyzed collagen peptides provide a rich source of glycine, proline, and hydroxyproline, critical substrates for *de novo* collagen synthesis by fibroblasts and osteoblasts [1,9]. However, its effects are not solely nutritional; specific bioactive sequences act as signaling molecules, directly upregulating the expression of extracellular matrix genes while simultaneously downregulating matrix metalloproteinases (MMPs), the enzymes responsible for collagen degradation [10-12]. This dual function—enhancing anabolism while suppressing catabolism—is central to its anti-aging efficacy. Beyond the skin, the musculoskeletal system benefits significantly from collagen supplementation. A growing body of evidence suggests that collagen peptides, particularly when combined with resistance training, can improve body composition and functional strength—a crucial factor in combating sarcopenia (age-related loss of muscle mass and function). For example, a randomized controlled trial demonstrated that collagen peptide supplementation in conjunction with resistance training increased muscle strength and lean mass more effectively than training alone in older sarcopenic men [23]. Similarly, benefits have been observed in premenopausal women, highlighting its broad scope of application [13]. For bone health, meta-analyses suggest that collagen peptides, particularly in synergy with calcium and vitamin D, can positively influence bone mineral density and bone metabolism biomarkers, offering a non-pharmacological strategy for osteoporosis prevention [9]. Perhaps the newest and most promising area of research concerns the systemic metabolic and potential lifespan-extending effects of collagen. Preclinical models offer compelling insights: studies in *C. elegans* have shown that cod collagen peptides can extend mean lifespan by modulating insulin/IGF-1 and AMPK signaling pathways [14]. In mammalian models, collagen supplementation has been shown to improve diet-induced obesity, enhance glucose tolerance, and positively modulate gut microbiota composition by increasing the Firmicutes/Bacteroidetes ratio, which is generally associated with a healthier metabolic phenotype [15,16]. Human trials corroborate these findings, demonstrating that certain collagen hydrolysates can improve postprandial glucose metabolism in normoglycemic and prediabetic individuals [17]. This supporting evidence suggests that collagen's effects may indeed be systemic and may influence key metabolic pathways linked to healthy lifespan.

Results

Clinical studies consistently demonstrate that oral collagen peptide supplementation provides statistically significant and clinically important improvements in various health areas. The strongest

evidence is available for dermatological applications. A meta-analysis of 19 randomized controlled trials (n=1,125) revealed that hydrolyzed collagen supplementation for approximately 90 days significantly improved skin hydration, elasticity, and wrinkle reduction compared to placebo [18,19]. These objective measures are supported by histological and molecular analyses; for example, bovine-derived bioactive peptides have been shown to stimulate human dermal fibroblasts *in vitro* to increase the production of type I collagen, elastin, and proteoglycans [5]. It also appears to benefit specific conditions, particularly cellulite; studies show significant reductions in skin ripples and dimpling after six months of supplementation [1]. Its benefits on the musculoskeletal system are equally well-documented. Clinical studies show that collagen peptide intake, when combined with exercise, can lead to significant increases in lean mass and muscle strength. Furthermore, research indicates a positive effect on bone health, with supplementation increasing bone mineral density, particularly in the lumbar spine and femoral neck, areas critical for osteoporotic fractures [9]. New metabolic data from human studies are promising; a randomized study in adults over 50 years of age found that low molecular weight collagen peptide supplementation reduced body fat mass compared to placebo [20]. Another study reported that a specific collagen hydrolysate improved postprandial glycemic control and may play a role in metabolic syndrome management.

The most important finding across all clinical trials is the extremely positive safety profile of collagen supplementation. Systematic reviews and individual trials over periods ranging from 4 weeks to 6 months consistently demonstrate that collagen peptides are well tolerated and that no serious side effects have been reported [18,21]. The most frequently reported effects are mild gastrointestinal sensations, rarely leading to discontinuation of treatment. This strong safety record, combined with measurable efficacy across multiple tissue systems, positions collagen supplementation as a low-risk, potentially high-return intervention within the framework of preventive health and longevity medicine.

Limitations

Despite promising evidence, current research on collagen supplementation is constrained by several methodological limitations that require careful interpretation and pave the way for more rigorous research. A major concern is that most intervention studies are relatively short-lived; typically, 8 to 12 weeks for skin studies and rarely exceeding 6 months for musculoskeletal outcomes [21,9]. This short-term focus leaves critical gaps in our understanding of the long-term sustainability of benefits, the potential for tachyphylaxis, and the ten-year safety profile of use, which is crucial for lifetime anti-aging interventions. Furthermore, while mid-term safety data are reassuring, comprehensive pharmacovigilance studies monitoring rare side effects over extended periods are lacking. Another significant limitation is the marked heterogeneity in study designs, which complicates cross-trial comparisons and meta-analytic synthesis. There is a lack of standardization regarding critical parameters such as optimal dosage (ranging from 2.5 g to 15 g daily), collagen source (bovine, porcine, marine, chicken), molecular weight profile of peptides, and co-administration with other nutrients such as vitamin C [22,23]. This variability makes it difficult to establish definitive clinical guidelines. In addition, many studies, particularly early dermatological studies, have been industry-funded; while this does not invalidate the results, it highlights the need for more independently funded, large-scale confirmatory studies to eliminate potential bias and strengthen the evidence base [19].

Finally, while the mechanistic understanding is still developing, it is only partially elucidated. Although absorption and fibroblast stimulation have been determined, the precise signaling pathways by which specific collagen peptides affect gene expression in skin, muscle, bone, and metabolic tissues need to be elucidated in more detail. Much of the mechanistic information is derived from *in vitro* models or animal studies (e.g., *C. elegans*, mice), and the direct transfer of these pathways to humans is not always clear [6,15]. Future research should prioritize human-centered mechanistic studies using omics technologies to identify response and resistance biomarkers, thereby paving the way for truly personalized collagen supplementation strategies.

Implications for Future Research

In order to transform collagen supplementation from a promising nutritional supplement into a robust component of geriatric and longevity medicine, future research must adopt a more rigorous, standardized, and mechanistic approach. First, there is an urgent need for large-scale, long-term (≥ 1 year), independently funded randomized controlled trials. These studies should investigate sustained efficacy for primary endpoints such as skin aging, bone density, and muscle mass, as well as “hard” clinical outcomes such as fracture incidence, progression to sarcopenia or disability, and long-term metabolic health indicators. The inclusion of comprehensive safety monitoring over these extended periods is equally critical to robust the risk-benefit profile. Second, the field requires a collaborative effort to address the current heterogeneity in supplementation protocols. Dose-finding studies are necessary to determine minimum effective doses for different health outcomes and populations. Research should also systematically compare the efficacy of collagen from different sources (marine, bovine, and recombinant), varying molecular weights, and diverse processing methods to identify superior formulations. Furthermore, the synergistic potential of collagen with other known anti-aging compounds such as hyaluronic acid, certain vitamins, minerals, or polyphenols warrants investigation to develop powerful multi-target formulations [10].

Thirdly, future research should deepen the mechanistic understanding through translational human studies. The use of techniques such as metabolomics to track the fate of ingested peptides, transcriptomics (via biopsy where ethical) to map gene expression changes in target tissues, and advanced imaging to measure real-time changes in the extracellular matrix *in vivo* would be invaluable. Simultaneously, research should focus on identifying genetic, phenotypic, or microbiome-based biomarkers that predict individual response. This precise nutritional approach will allow clinicians to personalize collagen supplementation, maximize efficacy, and ensure resources are directed to those who will benefit most, thereby revealing the true potential of collagen as a targeted intervention to increase healthy lifespan and resilience in aging populations [24,25].

Conclusion

In conclusion, cumulative evidence from preclinical models, clinical trials, and meta-analyses supports the proposition that collagen peptide supplementation is a safe and versatile intervention with the potential to modify various signs of aging. Its benefits extend beyond cosmetic improvements in skin hydration and elasticity, encompassing significant support for musculoskeletal integrity, as evidenced by increased muscle protein synthesis and bone mineral density. Perhaps most compellingly, emerging research points to systemic metabolic benefits, including improvements in body composition and glucose metabolism, with preliminary data from model organisms suggesting direct longevity-enhancing properties

through conserved genetic pathways. The novelty of this mini-review lies in its integrative synthesis, linking these diverse health benefits through a unified mechanistic framework focused on extracellular matrix support, anti-catabolic signaling, and systemic metabolic regulation. This positions collagen not merely as a cosmetic supplement, but as a fundamental nutritional strategy addressing the progressive decline in tissue structure and function associated with aging. Potentially positive health outcomes in older adults range from improved physical function and reduced fracture risk to better metabolic health and preserved dermal integrity, collectively contributing to a longer healthy lifespan and a better quality of life. However, translating this promise into concrete clinical practice requires overcoming current research limitations. Future studies should prioritize long-term efficacy and safety data, standardized optimal dosing, and a deeper understanding of individual response variability. By addressing these gaps, collagen supplementation can evolve from a popular health product to an evidence-based cornerstone of preventative gerontology. Consequently, as part of a holistic strategy encompassing diet, exercise, and other lifestyle modifications, collagen peptides are becoming a noteworthy candidate in the expanding spectrum of scientifically supported healthy aging interventions, offering a promising, low-risk pathway to increase endurance, slow age-related decline, and extend longevity.

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