

JPPIPA 9(11) (2023)

Jurnal Penelitian Pendidikan IPA

Journal of Research in Science Education



http://jppipa.unram.ac.id/index.php/jppipa/index

Bombyx Mori as a Potential Sunscreen to Prevent Photoaging

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Received: September 29, 2023 Revised: October 10, 2023 Accepted: November 25, 2023 Published: November 30, 2023

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DOI: 10.29303/jppipa.v9i11.5507

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Abstract: Excessive exposure to solar UV radiation is a major cause of skin photoaging, which is characterized by premature wrinkling, hyperpigmentation, and loss of skin elasticity. Traditional synthetic sunscreens are effective, but their potential side effects have led to a search for natural alternatives. Bombyx mori has attracted attention for its rich composition of bioactive compounds, such as silk proteins, sericin and fibroin, which have antioxidant and anti-inflammatory properties. These components are believed to contribute to the protection of skin cells against oxidative stress and inflammation from UV rays, thereby slowing down the aging process. This review systematically examines recent studies that have investigated the photoprotective effects of Bombyx mori-derived extracts, silk proteins, and other related substances. Findings suggest that these natural agents have the potential to reduce UV damage by enhancing skin defense mechanisms, promoting collagen synthesis, and reducing matrix metalloproteinase expression. In addition, the hypoallergenic properties of Bombyx mori compounds make them suitable for individuals with sensitive skin, thereby expanding their application in the skin care industry. However, while promising, research on Bombyx mori as a sunscreen agent is still in its early stages. Further studies are needed to elucidate the mechanisms underlying their photoprotective effects, assess their long-term safety, and optimize their formulations for effective delivery to the skin. This literature review contributes to a comprehensive understanding of the potential of Bombyx mori as an alternative to natural sunscreens and highlights the need for further research in harnessing its benefits to fight photoaging and maintain healthy skin.

Keywords: Bombyx mori; Photoaging; Photoprotective

Introduction

Increasing awareness of the detrimental effects of prolonged sun exposure on the skin has led to a growing agents demand for effective sunscreen (Puaratanaarunkon et al., 2022). Photoaging, characterized by premature aging of the skin due to ultraviolet (UV) radiation, has become a significant concern in dermatology and cosmetology (Grimes, 2015). Traditional synthetic sunscreens often have limitations, including potential side effects and inadequate protection against UV-A and UV-B rays (Azyyati Adzhani et al., 2022; Wiraguna et al., 2019). Consequently, there is an urgent need to explore alternative, natural sources for photoprotection (Jesus et al., 2022; Solano, 2020).

The silkworm Bombyx mori, known for its silk production, has attracted attention for its potential photoprotective properties (Kawamoto et al., 2019). Silk, a protein biomaterial, has interesting qualities which suggest that it can be used as a natural sunscreen agent. Studies have shown that Bombyx mori silk contains bioactive compounds with UV blocking and antioxidant abilities, which may contribute to the prevention of photoaging (Hidayat et al., 2022; K. Zhang et al., 2021). However, a comprehensive exploration of the scientific literature is essential to establish the viability of Bombyx

How to Cite:

Djunaidy, A., & Damayanti, P. A. A. (2023). Bombyx Mori as a Potential Sunscreen to Prevent Photoaging. *Jurnal Penelitian Pendidikan IPA*, 9(11), 1209–1220. https://doi.org/10.29303/jppipa.v9i11.5507

mori silk as an effective and safe sunscreen ingredient (Khosropanah, Zolbin, et al., 2022; X. Zhang et al., 2022).

The silk moth (Bombyx mori, literally mulberry caterpillaris an insect from the class Bombycidae (Ishiguro et al., 2021). It is called the silk moth because this animal is a producer of original silk which has high economic value (Xu et al., 2020; G. Zhao et al., 2020).

Photoaging, the process of premature skin aging caused by prolonged exposure to ultraviolet (UV) radiation from the sun, is a major concern in the fields of dermatology and cosmetic science. It is characterized by visible changes in the skin, including wrinkles, fine lines, pigmentation irregularities, and loss of skin elasticity (Andarina et al., 2017; Guan et al., 2021). These changes are primarily due to the detrimental effects of UV radiation on various cellular and molecular components of the skin. UV radiation, specifically UV-A and UV-B rays, penetrate the skin and interact with molecules such as DNA, proteins and lipids (Chen et al., 2021; Krutmann et al., 2021; Wang et al., 2019). This interaction triggers a series of harmful events, including the generation of reactive oxygen species (ROS) and activation of inflammatory pathways. This process causes DNA damage, oxidative stress, collagen degradation, and disruption of the skin's natural barrier function (Gromkowska-Kepka et al., 2021; Krutmann et al., 2021; McCabe et al., 2020).

This review will begin by presenting an overview of photoaging and its underlying mechanisms, highlighting the role of UV radiation and oxidative stress in skin damage. Next, it will explore the limitations of conventional synthetic sunscreens, emphasizing the need for alternative photoprotective agents. The discussion will then turn to the unique properties of Bombyx mori silk, including composition, structure, and the presence of bioactive molecules such as sericin and fibroin. These components are believed to contribute to silk's potential as a natural sunscreen.

This review will examine extensively the in vitro and in vivo studies that have investigated the UV blocking properties of Bombyx mori silk and its extracts. Various experimental methodologies and assessment techniques will be discussed, providing insight into silk's ability to attenuate UV damage to skin. Next, the potential antioxidant effect of Bombyx mori silk, which can enhance its photoprotective ability, will be analyzed in detail.

To ensure a comprehensive evaluation, the potential challenges and limitations associated with the use of Bombyx mori silk as a sunscreen agent will be discussed. This may include formulation considerations, stability issues, and possible allergic reactions. A comparison between Bombyx mori silk and existing synthetic and natural sunscreen agents will also be carried out to assess their competitive advantages and disadvantages. Photoaging, the premature aging of the skin caused by prolonged exposure to ultraviolet (UV) radiation, is a major concern in dermatology and cosmetology. The search for natural and effective photoprotective agents has led researchers to explore various compounds derived from plants, marine organisms, and insects. One interesting candidate is Bombyx mori, commonly known as the silkworm. This comprehensive literature review aims to provide an indepth analysis of existing research on Bombyx mori as a potential sunscreen agent to prevent photoaging.

Photoaging, the process of premature skin aging caused by prolonged exposure to ultraviolet (UV) radiation from the sun, is a major concern in the fields of dermatology and cosmetic science. It is characterized by visible changes in the skin, including wrinkles, fine lines, pigmentation irregularities, and loss of skin elasticity. These changes are primarily due to the detrimental effects of UV radiation on various cellular and molecular components of the skin. UV radiation, specifically UV-A and UV-B rays, penetrate the skin and interact with molecules such as DNA, proteins and lipids. This interaction triggers a series of harmful events, including the generation of reactive oxygen species (ROS) and activation of inflammatory pathways. This process causes DNA damage, oxidative stress, collagen degradation, and disruption of the skin's natural barrier function.

The skin's main defense against the damaging effects of UV radiation is the use of sunscreen. Sunscreens are formulated to provide a protective barrier that absorbs or reflects UV rays, reducing their penetration into the skin. By acting as a protective barrier, sunscreen helps reduce the harmful effects of UV radiation on the cellular components of the skin. These precautions are essential in maintaining healthy skin and a youthful appearance. The molecular mechanisms underlying the photoprotective effect of Bombyx mori compounds are complex and diverse. These compounds, including sericin and fibroin, interact with several key biomolecules to carry out preventive actions against UV damage. These interactions collectively contribute to the preservation of healthy skin and the reduction of photoaging.

Reactive oxygen species (ROS) are highly reactive molecules produced by exposure to UV radiation. They play a central role in oxidative stress, which causes cell damage, inflammation and premature aging. Bombyx mori compounds, in particular sericin, exhibit antioxidant properties that counteract the harmful effects of ROS. By cleansing and neutralizing ROS, sericin helps maintain redox balance within skin cells, reduces oxidative stress and minimizes damage caused by UV radiation.

Matrix metalloproteinases (MMPs) are enzymes responsible for the breakdown of components of the extracellular matrix, including collagen and elastin. UV radiation can increase MMP expression, causing degradation of this structural protein and contributing to skin aging. Bombyx mori compounds, such as fibroin, were found to modulate MMP activity. By inhibiting MMP expression and activity, fibroin helps maintain the integrity of the extracellular matrix, maintaining skin elasticity and structure even under UV exposure.

Pro-inflammatory cvtokines are signaling molecules that initiate and regulate inflammatory responses in the skin. UV radiation triggers the release pro-inflammatory cytokines, which of cause inflammation and tissue damage. Sericin, found in Bombyx mori extract, has anti-inflammatory properties. It inhibits the production of pro-inflammatory cytokines, such as interleukin-6 (IL-6) and tumor necrosis factor-alpha (TNF- α), thereby attenuating the inflammatory cascade induced by UV radiation. This anti-inflammatory effect contributes to the prevention of skin redness, irritation, and long-term damage associated with chronic inflammation.

Furthermore, Bombyx mori compounds have been shown to enhance the skin's natural repair mechanisms. They promote the expression of genes involved in wound healing and tissue regeneration, facilitating the recovery of UV-damaged skin. This repair process involves the stimulation of collagen synthesis, which supports the restoration of skin's structural integrity and resilience. The interactions between these molecular mechanisms contribute to the overall photoprotective effect of the Bombyx mori compound. By reducing oxidative stress, inhibiting MMPs, reducing inflammation, and promoting skin repair, these compounds collectively fortify the skin's defenses against UV damage. The multifunctional nature of the Bombyx mori compound underscores its potential as an effective agent in preventing photoaging and maintaining healthy skin in the face of chronic sun exposure (Khosropanah, Vaghasloo, et al., 2022; Safonova et al., 2021).

However, it is important to note that while promising, more research is needed to fully elucidate the intricacies of this molecular mechanism and validate its effectiveness in real-world scenarios. Deeper clinical trials and molecular studies will provide a more comprehensive understanding of how Bombyx mori compounds interact with these biomolecules and contribute to their photoprotective effects.

Comparative analyzes between Bombyx mori compounds and conventional sunscreen agents reveal

valuable insights into the potential advantages and disadvantages of using silkworm-derived compounds for photoprotection. This check covers critical factors, including broad-spectrum protection, stability, and long-term effects (Lujerdean et al., 2022; Su et al., 2019).

Reactive oxygen species (ROS) are highly reactive molecules produced by exposure to UV radiation. They play a central role in oxidative stress, which causes cell damage, inflammation and premature aging. Bombyx mori compounds, in particular sericin, exhibit antioxidant properties that counteract the harmful effects of ROS. By cleansing and neutralizing ROS, sericin helps maintain redox balance within skin cells, reduces oxidative stress and minimizes damage caused by UV radiation. In addition to sunscreen formulations, Bombyx mori compounds can be used in other skincare products aimed at preventing or reducing photoaging. Investigating their incorporation into moisturizers, serums, or even oral supplements may provide a holistic approach to skin health and photo protection. Understanding potential mechanisms of action beyond photoprotection, such as wound healing and collagen synthesis, will guide the development of such products (Ali et al., 2018; Podbevšek et al., 2022; Silva-Silva et al., 2022).

This research aims to explore existing research to examine the potential of Bombyx mori silk as a sunscreen agent to prevent photoaging. This research aims to contribute to the development of knowledge about natural photoprotective agents by exploring the potential of Bombyx mori silk as a sunscreen to prevent photoaging. By synthesizing existing research and critically evaluating its findings, this review intends to elucidate the viability of Bombyx mori silk as a novel approach to protect the skin from UV damage. This investigation has implications for the fields of dermatology, cosmetics, and natural product development, paving the way for further studies and potential applications of Bombyx mori silk in photoprotection.

Method

The methodological approach used in this research is Systematic Review (Sugiyono, 2019). This research focuses on randomized controlled trials published on the website from 2017 to 2022. Researchers will screen, extract the data, and cross-check the results. The method used is to use a literature review from searching, selecting, and reviewing the results of literature research that has been published in national and international electronic media. Search international journal articles using Science Direct, ProQuest, CINAH, Pubmed and Clinical Trials.gov. This research will be conducted by collecting and evaluating various relevant literature sources to identify previous research that has been conducted on this topic. By summarizing and synthesizing the findings from the existing literature, the ultimate goal is to gain a comprehensive understanding of the effect of passion fruit seed content on sun protection and its impact on the skin aging process. The overall results of this research method will be set forth in the form of a systematic and structured literature review report, including introduction, methodology, findings, analysis, conclusions, as well as a bibliography that refers to the sources that have been analyzed (Bunawan et al., 2014; Creswel, 2016).

Results and Discussion

Result

Sixteen articles describe Bombyx mori as a potential sunscreen for preventing photoaging. Bombyx was

studied by experimental method and literature review. Samples vary from molecular to bedside investigations.

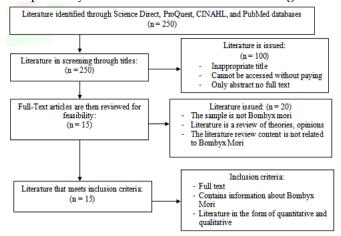


Figure 1. Literature review process flowchart

| Table 1. Bombyx Mori anal Author & year of publication | Research methodology | Results |
|--|---------------------------------|--|
| Belal et al. (2023) | Experimental Study | The results showed that ZnO-NP at concentrations of 50 and |
| Delai et al. (2023) | Experimental Study | $100 \ \mu\text{g/ml}$ significantly reduced the total hemocyte count |
| | | (THC) and the different haemocyte count (DHC) except for the |
| | | oenocyte count which increased significantly. |
| Li et al. (2020) | Experimental Study | The results showed that exposure to TiO2 NPs did not |
| 21 00 011 (2020) | 2.1. permernan eraday | significantly change the dominant species of gut microbiota, but |
| | | did change the abundance of individual species of gut |
| | | microbiota. |
| Mir et al. (2020) | In vivo experimental studies | Our study also shows that B. mori has the potential to serve as |
| | | an effective alternative animal model for biosafety, |
| | | environmental monitoring, and NP screening, especially for |
| | | evaluating its interaction with the invertebrate immune system. |
| Fometu et al. (2022) | Experimental Study | This study highlights a different approach in evaluating the |
| | | biological effects of TiO 2 NPs using the silkworm as a model |
| | | and assessing their impact on the intrinsic properties of silk, |
| | | which will be effective in biotechnological applications. |
| Fang et al. (2022) | RCTs | The results demonstrated that TiO2 NPs exhibited a dose- |
| | | dependent effect on relative cell survival, with significant toxic |
| | | effects observed above 20 mg/L. |
| Muhammad et al. (2022) | Experimental Study | This integrated study reveals the unwanted negative effects of |
| | | CuO and ZnO NPs on silkworms and highlights the potential to |
| | | affect all living things due to intensive and possible mishandling |
| | | of nanomaterials. |
| Xu et al. (2020) | | The results of catalytic activity studies confirmed that antioxidant |
| | | enzymes (SOD, CAT, GSH-PX) in midgut cells were expressed in |
| | | response to ZnO NPs. Expression of genes (Dronc and Caspase- |
| | | 1) related to apoptosis is increased, whereas the Trt gene is |
| | | down-regulated. |
| Zhao et al. (2020) | In vitro experimental studies | The results showed that nano-titanium dioxide pretreatment |
| | | could inhibit the proliferation of BmCPV in the midgut of |
| | | silkworms, activate the JAK/STAT and PI3K-AKT immune |
| | | signaling pathways, and increase the expression of key immune |
| | | genes, thereby enhancing silkworm immunity. and increase silkworm resistance to BmCPV. |
| J. Li et al (2020) | In vitro, ex vivo, and pre-post | These results provide new clues for studying the molecular |
| J. El Ct di (2020) | intervention clinical trials | mechanisms of insect resistance to high temperatures. |
| | intervention childer thats | incentation of insect resistance to mgn temperatures. |

| Author & year of publication | Research methodology | Results |
|------------------------------|------------------------------|--|
| Wang et al. (2023) | In vivo experimental studies | These results provide evidence for multiple DR anti-aging |
| 0 () | | mechanisms at the metabolic level and a new reference for the |
| | | future development of DR-simulated drugs or foods. |
| Tu et al. (2022) | Article review | The results showed that GABA increased the fecundity of adult |
| | | silkworms. And in silkworm larvae on day 3 to 5 instar, GABA |
| | | significantly reduced trehalose content in hemolymph, |
| | | triglyceride and glycogen levels in body fat, while markedly |
| | | increasing NAD+/NADH levels in body fat. |
| Yeerong et al. (2021) | Experimental Study | The anti-aging activity was investigated by determining the |
| | | collagenase and elastase inhibitory activity using a |
| | | spectrophotometric assay. |
| Ishiguro et al. (2021) | In vivo experimental studies | This finding of in vitro glia-neuron interactions led us to |
| | | evaluate the effect of oral administration of the peptide on brain |
| | | function and hair aging in accelerated aging (SAMP8) mice. |
| P. Zhao et al. (2018) | In vivo experimental studies | Our results show that BmApoD1 is essential for the metabolic |
| | | adaptation of B. mori to environmental challenges. |
| Qu et al. (2023) | RCTs | Our results show that most of the lipids in the host midgut |
| | | decreased when silkworms were infected with Nb. Reduction |
| | | or supplementation with PC may be a strategy to suppress or |
| | | encourage microsporidia replication. |

Discussion

Photoaging, the premature aging of the skin caused by prolonged exposure to ultraviolet (UV) radiation, is a major concern in dermatology and cosmetology. The search for natural and effective photoprotective agents has led researchers to explore various compounds derived from plants, marine organisms, and insects. One interesting candidate is Bombyx mori, commonly known as the silkworm. This comprehensive literature review aims to provide an in-depth analysis of existing research on Bombyx mori as a potential sunscreen agent to prevent photoaging.

The silk moth (Bombyx mori, literally mulberry caterpillar is an insect from the class Bombycidae. It is called the silk moth because this animal is a producer of original silk which has high economic value. The form of the caterpillar (larvae) which is often called the silkworm is better known to people than the adult form (imago) of this animal. These animals are often raised for their silk. Silkworms only eat leaves mulberry (Morus alba). It originates from northern China (Kawamoto et al., 2019).

Photoaging, the process of premature skin aging caused by prolonged exposure to ultraviolet (UV) radiation from the sun, is a major concern in the fields of dermatology and cosmetic science. It is characterized by visible changes in the skin, including wrinkles, fine lines, pigmentation irregularities, and loss of skin elasticity. These changes are primarily due to the detrimental effects of UV radiation on various cellular and molecular components of the skin. UV radiation, specifically UV-A and UV-B rays, penetrate the skin and interact with molecules such as DNA, proteins and lipids. This interaction triggers a series of harmful events, including the generation of reactive oxygen species (ROS) and activation of inflammatory pathways. This process causes DNA damage, oxidative stress, collagen degradation, and disruption of the skin's natural barrier function (Kumar et al., 2019).

The skin's main defense against the damaging effects of UV radiation is the use of sunscreen. Sunscreens are formulated to provide a protective barrier that absorbs or reflects UV rays, reducing their penetration into the skin. By acting as a protective barrier, sunscreen helps reduce the harmful effects of UV radiation on the cellular components of the skin. These precautions are essential in maintaining healthy skin and a youthful appearance (Brunetti et al., 2018; Guidi et al., 2016).

The molecular mechanisms underlying the photoprotective effect of Bombyx mori compounds are complex and diverse. These compounds, including sericin and fibroin, interact with several key biomolecules to carry out preventive actions against UV damage. These interactions collectively contribute to the preservation of healthy skin and the reduction of photoaging (Agustina et al., 2022).

Reactive oxygen species (ROS) are highly reactive molecules produced by exposure to UV radiation. They play a central role in oxidative stress, which causes cell damage, inflammation and premature aging. Bombyx mori compounds, in particular sericin, exhibit antioxidant properties that counteract the harmful effects of ROS. By cleansing and neutralizing ROS, sericin helps maintain redox balance within skin cells, reduces oxidative stress and minimizes damage caused by UV radiation (Manosroi et al., 2010). Matrix metalloproteinases (MMPs) are enzymes responsible for the breakdown of components of the extracellular matrix, including collagen and elastin. UV radiation can increase MMP expression, causing degradation of this structural protein and contributing to skin aging (Cabral-Pacheco et al., 2020; Laronha, Carpinteiro, et al., 2020). Bombyx mori compounds, such as fibroin, were found to modulate MMP activity. By inhibiting MMP expression and activity, fibroin helps maintain the integrity of the extracellular matrix, maintaining skin elasticity and structure even under UV exposure (Checchi et al., 2020; Niland et al., 2022).

Pro-inflammatory cvtokines are signaling molecules that initiate and regulate inflammatory responses in the skin (Laronha & Caldeira, 2020; Maybee et al., 2022). UV radiation triggers the release of proinflammatory cytokines, which cause inflammation and tissue damage. Sericin, found in Bombyx mori extract, has anti-inflammatory properties (Jordan et al., 2021; Pedro et al., 2021). It inhibits the production of proinflammatory cytokines, such as interleukin-6 (IL-6) and necrosis factor-alpha $(TNF-\alpha),$ thereby tumor attenuating the inflammatory cascade induced by UV radiation. This anti-inflammatory effect contributes to the prevention of skin redness, irritation, and long-term damage associated with chronic inflammation (Jordan et al., 2021; Nilsson, 2021).

Furthermore, Bombyx mori compounds have been shown to enhance the skin's natural repair mechanisms. They promote the expression of genes involved in wound healing and tissue regeneration, facilitating the recovery of UV-damaged skin. This repair process involves the stimulation of collagen synthesis, which supports the restoration of skin's structural integrity and resilience.

The interactions between these molecular mechanisms contribute to the overall photoprotective effect of the Bombyx mori compound. By reducing inhibiting MMPs, oxidative stress, reducing inflammation, and promoting skin repair, these compounds collectively fortify the skin's defenses against UV damage. The multifunctional nature of the Bombyx mori compound underscores its potential as an effective agent in preventing photoaging and maintaining healthy skin in the face of chronic sun exposure.

However, it is important to note that while promising, more research is needed to fully elucidate the intricacies of this molecular mechanism and validate its effectiveness in real-world scenarios. Deeper clinical trials and molecular studies will provide a more comprehensive understanding of how Bombyx mori compounds interact with these biomolecules and contribute to their photoprotective effects. Comparative analyzes between Bombyx mori compounds and conventional sunscreen agents reveal valuable insights into the potential advantages and disadvantages of using silkworm-derived compounds for photoprotection. This check covers critical factors, including broad-spectrum protection, stability, and long-term effects.

Skin aging can be influenced by intrinsic and extrinsic factors. Intrinsic factor is a process that occurs in the human body over time. While extrinsic factors are things that come from the environment, such as exposure to sunlight, air pollution, etc. These extrinsic factors can cause premature aging of the skin, and more specifically extrinsic factors of sun exposure can cause photoaging. Basically, the body has its own mechanism for filtering sunlight, namely with the pigment melanin. But continuous exposure to sunlight can cause damage to the skin. Consistent use of things that are protective against sunlight (photoprotective) is important to prevent photoaging, including to avoid the dangers of ultraviolet light. Photoprotection is very important for people who frequently carry out outdoor activities, one of which is a student. In which a student is required to always actively participate in various kinds of activities besides lecture activities in class, for example participating in social service events that require students to go directly into the field, as well as other activities that are usually carried out in the context of community service. Therefore, the use of photoprotection in students is highly recommended.

Sunscreen is a skin care product that combines several ingredients that can protect the skin from UV rays. Sunscreen is effective as primary photoprotection. To achieve maximum results, you should use sunscreen every day even in cloudy weather with a thickness of 2 mg/cm2 on the skin surface or the equivalent of one teaspoon for the face area only. Then it must be applied at least 15 minutes before exposure to sunlight and reapplied every 2 hours when doing outdoor activities. A broad spectrum sunscreen with an SPF of at least 30 is recommended for daily use.

Traditional sunscreen agents, usually formulated with chemical or physical filters, have long been the cornerstone of sun protection. These agents work by absorbing, reflecting, or scattering UV radiation to prevent it from penetrating the skin. They offer a variety of SPF (sun protection factor) values, allowing consumers to choose the level of protection that suits their needs. However, certain chemical filters, such as oxybenzone and octinoxate, have raised concerns because of their potential to cause skin irritation, allergic reactions, and environmental damage, especially in coral reef ecosystems.

In contrast, compounds derived from Bombyx mori, particularly sericin and fibroin, offer a more natural alternative to photoprotection. They are known for their antioxidant and anti-inflammatory properties, which can potentially counteract the harmful effects of radiation, including oxidative stress UV and inflammation. These compounds also have the advantage of being biocompatible with the skin, reducing the likelihood of adverse reactions. Increasing awareness of the detrimental effects of prolonged sun exposure on the skin has led to a growing demand for effective sunscreen agents. Photoaging, characterized by premature aging of the skin due to ultraviolet (UV) radiation, has become a significant concern in dermatology and cosmetology. Traditional synthetic sunscreens often have limitations, including potential side effects and inadequate protection against UV-A and UV-B rays. Consequently, there is an urgent need to explore alternative, natural sources for photoprotection.

The silkworm Bombyx mori, known for its silk production, has attracted attention for its potential photoprotective properties. Silk, a protein biomaterial, has interesting qualities which suggest that it can be used as a natural sunscreen agent. Studies have shown that Bombyx mori silk contains bioactive compounds with UV blocking and antioxidant abilities, which may contribute to the prevention of photoaging. However, a comprehensive exploration of the scientific literature is essential to establish the viability of Bombyx mori silk as an effective and safe sunscreen ingredient.

Broad spectrum protection is a major consideration in sunscreens, as it ensures protection against both UVA and UVB radiation. Conventional sunscreen agents often require a combination of filters to achieve broad spectrum protection. Bombyx mori compounds, while not broad spectrum protective, can potentially be combined with other ingredients to create formulations that offer comprehensive defense against different wavelengths of UV radiation.

Stability is an important aspect of sunscreen formulation. UV radiation and environmental factors can reduce the effectiveness of the active ingredients over time. Traditional sunscreen agents may require stabilizers or encapsulation technologies to maintain their efficacy. Bombyx mori compounds, which are derived from natural sources, can pose stability challenges, especially under prolonged sun exposure or varying environmental conditions. Developing an effective stabilization method can be a significant consideration in utilizing this compound.

Long-term effects are critical, as prolonged sun exposure over many years can cause cumulative skin damage. Several conventional sunscreen agents have been criticized for their potential endocrine disrupting effects, allergenicity, or the formation of reactive oxygen species following exposure to UV light. Bombyx mori compounds, with their potent antioxidant properties, may provide an advantage in minimizing the long-term damage caused by UV radiation. However, rigorous long-term safety and efficacy studies will be essential to establish its suitability for long-term use. In conclusion, the comparative analysis underscores that while Bombyx mori compounds offer a natural and potentially biocompatible alternative for photoprotection, there are several considerations to be aware of compared to traditional sunscreen agents. These considerations include their potential to offer broad-spectrum protection, stability in formulations, and their long-term effects on skin health. Exploration of synergistic combinations and innovative formulations can unlock the full potential of the Bombyx mori compound as an effective and safe photoprotective agent against the backdrop of existing sunscreen options.

This comprehensive review of Bombyx mori as a potential sunscreen to prevent photoaging not only synthesizes existing knowledge but also highlights areas where further investigation is needed. Looking ahead, there are several promising directions for future research and potential applications that could enhance the understanding and utilization of Bombyx mori compounds for photoprotection.

One of the most critical avenues for advancing this field is the commencement of well-designed clinical trials. While in vitro and in vivo studies have provided valuable insights into the photoprotective effects of Bombyx mori compounds, translation of these findings into real-world conditions is critical. Rigorous clinical trials involving human participants of various skin types and phototypes will provide conclusive evidence of the efficacy and safety of Bombyx mori-based sunscreens. These trials should involve controlled UV radiation exposure and thorough evaluation of photoprotective outcomes such as erythema, pigmentation, wrinkle formation, and overall skin health.

Moreover, exploring the potential synergistic effect of Bombyx mori compounds with existing photoprotective agents presents an interesting avenue for future research. Combining the unique properties of Bombyx mori with existing sunscreen ingredients can result in better protection against UV damage. Research in this direction should focus on understanding potential interactions, formulation compatibility, and optimal ratios to achieve synergistic effects while maintaining stability and safety.

In addition to sunscreen formulations, Bombyx mori compounds can be used in other skincare products aimed at preventing or reducing photoaging. Investigating their incorporation into moisturizers, serums, or even oral supplements may provide a holistic approach to skin health and photo protection. Understanding potential mechanisms of action beyond photoprotection, such as wound healing and collagen synthesis, will guide the development of such products.

In addition, exploring the feasibility of sustainably sourcing and producing Bombyx mori compounds on a larger scale is an important consideration. The review recognized the ecological significance of sericulture, and future research may focus on optimizing silkworm cultivation methods to ensure a consistent and environmentally friendly supply of Bombyx mori derived compounds. As the regulatory landscape evolves, establishing standardized testing protocols and safety assessments for Bombyx mori-based sunscreens and skincare products will become critical. Regulators need solid data on efficacy, safety and potential allergic reactions to ensure consumer protection. Addressing these requirements will facilitate the commercialization and widespread use of Bombyx mori compounds in the skin care industry.

In conclusion, the potential of Bombyx mori as a natural photoprotective agent to prevent photoaging is very promising, however further research is essential to validate its efficacy, safety and applicability. Welldesigned clinical trials, exploration of synergistic combinations, consideration of sustainable sourcing, and adherence to regulatory standards are important components for advancing this field. By setting these future directions, researchers and industry professionals can contribute to the development of innovative and effective strategies to combat the damaging effects of UV radiation on the skin.

In conclusion, the potential of Bombyx mori as a natural photoprotective agent to prevent photoaging is very promising, however further research is essential to validate its efficacy, safety and applicability. Welldesigned clinical trials, exploration of synergistic combinations, consideration of sustainable sourcing, and adherence to regulatory standards are important components for advancing this field. By setting these future directions, researchers and industry professionals can contribute to the development of innovative and effective strategies to combat the damaging effects of UV radiation on the skin. In conclusion, this literature review aims to contribute to the development of knowledge about natural photoprotective agents by exploring the potential of Bombyx mori silk as a sunscreen to prevent photoaging. By synthesizing existing research and critically evaluating its findings, this review intends to elucidate the viability of Bombyx mori silk as a novel approach to protect the skin from UV damage. This investigation has implications for the fields of dermatology, cosmetics, and natural product development, paving the way for further studies and potential applications of Bombyx mori silk in photoprotection.

Conclusion

In conclusion, this comprehensive literature review emphasizes the potential of Bombyx mori as a natural sunscreen agent to prevent photoaging. Its antioxidant, anti-inflammatory, collagen-preserving, and UVscattering properties demonstrate its effectiveness in countering the harmful effects of UV radiation on the skin. However, translating these findings into practical skincare applications requires rigorous scientific investigation and formulation development. Bombyx mori holds promise as an innovative pathway in the ongoing quest for an effective and safe photoprotective strategy. While the literature highlights the promising photoprotective properties of Bombyx mori, several challenges need to be overcome before its practical application as a sunscreen. This includes standardizing extraction methods. determining optimal concentrations, assessing potential allergenicity, and evaluating long-term safety. More research is needed to bridge the gap between laboratory findings and realworld sunscreen products.

Acknowledgments

The authors would like to thanks to Jambi University for give occasion for this research.

Author Contributions

Investigation, A.D and P.A.A.D; formal analysis, A.D and P.A.A.D; investigation, A.D and P.A.A.D; resources A.D and P.A.A.D; data curation, A.D and P.A.A.D; writing—original draft preparation, A.D and P.A.A.D; writing—review and editing, A.D and P.A.A.D; visualization A.D and P.A.A.D; supervision, A.D and P.A.A.D; project administration A.D and P.A.A.D; funding acquisition, A.D and P.A.A.D. All authors have read and agreed to the published version of the manuscript.

Funding

This research is fully supported by the author's funds without any external funding sources.

Conflicts of Interest

We certify that there is no conflict of interest with any financial, personal and other relationships with other peoples or organization related to the material discussed in the manuscript.

References

Agustina, F., Fadilah, F., Pangkahila, W., Putra Wiraguna, A. A. G., & Mahendra Dewi, I. G. A. S. (2022). Study of Sericin Sequences from Bombyx mori as Antiaging through ROS with Molecular Simulation and DPPH Evaluation. *Pharmacognosy Journal*, 14(5).

https://doi.org/10.5530/pj.2022.14.146

- Ali, T., Rehman, S. U., Shah, F. A., & Kim, M. O. (2018). Acute dose of melatonin via Nrf2 dependently prevents acute ethanol-induced neurotoxicity in the developing rodent brain. *Journal of Neuroinflammation*, 15(1). https://doi.org/10.1186/s12974-018-1157-x
- Andarina, R., & Djauhari, T. (2017). Antioksidan Dalam Dermatologi. *Jurnal Kedokteran Dan Kesehatan*, 4(1). Retrieved from https://core.ac.uk/download/pdf/267823473.pd f
- Azyyati Adzhani, Fitrianti Darusman, & Ratih Aryani. (2022). Kajian Efek Radiasi Ultraviolet terhadap Kulit. *Bandung Conference Series: Pharmacy*, 2(2). https://doi.org/10.29313/bcsp.v2i2.3551
- Belal, R., & Gad, A. (2023). Zinc oxide nanoparticles induce oxidative stress, genotoxicity, and apoptosis in the hemocytes of Bombyx mori larvae. *Scientific Reports*, 13(1). https://doi.org/10.1038/s41598-023-30444-y
- Brunetti, C., Fini, A., Sebastiani, F., Gori, A., & Tattini, M. (2018). Modulation of phytohormone signaling: A primary function of flavonoids in plantenvironment interactions. *Frontiers in Plant Science*, 9. https://doi.org/10.3389/fpls.2018.01042
- Bunawan, N. C., Rastegar, A., White, K. P., & Wang, N. E. (2014). Djenkolism: Case report and literature review. *International Medical Case Reports Journal*, 7(1). https://doi.org/10.2147/IMCRJ.S58379
- Cabral-Pacheco, G. A., Garza-Veloz, I., Rosa, C. C. D. La, Ramirez-Acuña, J. M., Perez-Romero, B. A., Guerrero-Rodriguez, J. F., Martinez-Avila, N., & Martinez-Fierro, M. L. (2020). The roles of matrix metalloproteinases and their inhibitors in human diseases. *International Journal of Molecular Sciences*, 21(24). https://doi.org/10.3390/ijms21249739
- Checchi, V., Maravic, T., Bellini, P., Generali, L., Consolo, U., Breschi, L., & Mazzoni, A. (2020). The role of matrix metalloproteinases in periodontal disease. *International Journal of Environmental Research and Public Health*, 17(14). https://doi.org/10.3390/ijerph17144923
- Chen, X., Yang, C., & Jiang, G. (2021). Research progress on skin photoaging and oxidative stress. *Postepy Dermatologii i Alergologii, 38*(6). https://doi.org/10.5114/ada.2021.112275
- Creswel, J. W. (2016). Research Design –Pendekatan Metode Kualitatif, Kuantitatif, dan Campuran. Yogyakarta: Pustaka Pelajar
- Fang, Y., Dai, M., Ye, W., Li, F., Sun, H., Wei, J., & Li, B.

(2022). Damaging effects of TiO2 nanoparticles on the ovarian cells of Bombyx mori. *Biological Trace Element Research*, 200(4), 1883–1891. https://doi.org/10.1007/s12011-021-02760-9

- Fometu, S. S., Ma, Q., Wang, J. J., Guo, J., Ma, L., & Wu, G. (2022). Biological Effect Evaluation of Different Sized Titanium Dioxide Nanoparticles Using Bombyx mori (Silkworm) as a Model Animal. *Biological Trace Element Research*, 200(12), 5260– 5272. https://doi.org/10.1007/s12011-021-03086-2
- Grimes, D. R. (2015). Ultraviolet radiation therapy and UVR dose models. *Medical Physics*, 42(1). https://doi.org/10.1118/1.4903963
- Gromkowska-Kępka, K. J., Puścion-Jakubik, A., Markiewicz-Żukowska, R., & Socha, K. (2021). The impact of ultraviolet radiation on skin photoagingreview of in vitro studies. *Journal of Cosmetic Dermatology*, 20(11). https://doi.org/10.1111/jocd.14033
- Guan, L. L., Lim, H. W., & Mohammad, T. F. (2021). Sunscreens and Photoaging: A Review of Current Literature. In *American Journal of Clinical Dermatology*, 22(6). https://doi.org/10.1007/s40257-021-00632-5
- Guidi, L., Brunetti, C., Fini, A., Agati, G., Ferrini, F., Gori, A., & Tattini, M. (2016). UV radiation promotes flavonoid biosynthesis, while negatively affecting the biosynthesis and the de-epoxidation of xanthophylls: Consequence for photoprotection? *Environmental and Experimental Botany*, 127. https://doi.org/10.1016/j.envexpbot.2016.03.002
- Hidayat, A. F., Aryani, R., & Hannisah, I. (2022).
 Preparation of Moisturizing Lotion Containing Silkworm (Bombyx mori L.) Sericin Nanoparticles. *Indonesian Journal of Pharmaceutical Science and Technology*, 9(1).
 https://doi.org/10.24198/ijpst.v1i1.30486

Ishiguro, S., Shinada, T., Wu, Z., Karimazawa, M., Uchidate, M., Nishimura, E., Yasuno, Y., Ebata, M., Sillapakong, P., Ishiguro, H., Ebata, N., Ni, J., Jiang, M., Goryo, M., Otsu, K., Harada, H., & Suzuki, K. (2021). A novel cyclic peptide (Naturido) modulates glia-neuron interactions in vitro and reverses ageing-related deficits in senescenceaccelerated mice. *PLoS ONE*, 16. https://doi.org/10.1371/journal.pone.0245235

- Jesus, A., Sousa, E., Cruz, M. T., Cidade, H., Lobo, J. M. S., & Almeida, I. F. (2022). UV Filters: Challenges and Prospects. *Pharmaceuticals*, 15(3). https://doi.org/10.3390/ph15030263
- Jordan, P. M., Andreas, N., Groth, M., Wegner, P., Weber, F., Jäger, U., Küchler, C., Werz, O., Serfling, E., Kamradt, T., Dudeck, A., & Drube, S. (2021). ATP/IL-33-triggered hyperactivation of mast cells

results in an amplified production of proinflammatory cytokines and eicosanoids. *Immunology*, 164(3).

https://doi.org/10.1111/imm.13386

- Kawamoto, M., Jouraku, A., Toyoda, A., Yokoi, K., Minakuchi, Y., Katsuma, S., Fujiyama, A., Kiuchi, T., Yamamoto, K., & Shimada, T. (2019). Highquality genome assembly of the silkworm, Bombyx mori. *Insect Biochemistry and Molecular Biology*, 107. https://doi.org/10.1016/j.ibmb.2019.02.002
- Khosropanah, M. H., Vaghasloo, M. A., Shakibaei, M., Mueller, A. L., Kajbafzadeh, A. M., Amani, L., Haririan, I., Azimzadeh, A., Hassannejad, Z., & Zolbin, M. M. (2022). Biomedical applications of silkworm (Bombyx Mori) proteins in regenerative medicine (a narrative review). *Journal of Tissue Engineering and Regenerative Medicine*, 16(2). https://doi.org/10.1002/term.3267
- Khosropanah, M. H., Zolbin, M. M., Kajbafzadeh, A. M., Amani, L., Harririan, I., Azimzadeh, A., Nejatian, T., Vaghsloo, M. A., & Hassannejad, Z. (2022). Evaluation and Comparison of the Effects of Mature Silkworm (Bombyx mori) and Silkworm Pupae Extracts on Schwann Cell Proliferation and Axon Growth: An In Vitro Study. *Iranian Journal of Pharmaceutical* Research, 21(1). https://doi.org/10.5812/ijpr-133552
- Krutmann, J., Schalka, S., Watson, R. E. B., Wei, L., & Morita, A. (2021). Daily photoprotection to prevent photoaging. In *Photodermatology Photoimmunology and Photomedicine*, 37(6). https://doi.org/10.1111/phpp.12688
- Kumar, J. P., & Mandal, B. B. (2019). Inhibitory role of silk cocoon extract against elastase, hyaluronidase and UV radiation-induced matrix metalloproteinase expression in human dermal fibroblasts and keratinocytes. *Photochemical and Photobiological Sciences*, 18(5). https://doi.org/10.1039/c8pp00524a
- Laronha, H., & Caldeira, J. (2020). Structure and Function of Human Matrix Metalloproteinases. *Cells*, 9(5). https://doi.org/10.3390/cells9051076
- Laronha, H., Carpinteiro, I., Portugal, J., Azul, A., Polido, M., Petrova, K. T., Salema-Oom, M., & Caldeira, J. (2020). Challenges in matrix metalloproteinases inhibition. *Biomolecules*, 10(5). https://doi.org/10.3390/biom10050717
- Li, J., Lu, Z., Mao, T., Li, M., Wang, H., Qu, J., Chen, J., Fang, Y., Li, F., & Li, B. (2020). Identification of the nucleotide exchange factor BmGrpE and its role under high-temperature stress in silkworm, Bombyx mori. *Archives of Insect Biochemistry and Physiology*, 104(1).

https://doi.org/10.1002/arch.21664

Li, M., Li, F., Lu, Z., Fang, Y., Qu, J., Mao, T., Wang, H., Chen, J., & Li, B. (2020). Effects of TiO2 nanoparticles on intestinal microbial composition of silkworm, Bombyx mori. *Science of the Total Environment*, 704.

https://doi.org/10.1016/j.scitotenv.2019.135273

- Lujerdean, C., Baci, G. M., Cucu, A. A., & Dezmirean, D. S. (2022). The Contribution of Silk Fibroin in Biomedical Engineering. *Insects*, 13(3). https://doi.org/10.3390/insects13030286
- Manosroi, A., Boonpisuttinant, K., Winitchai, S., Manosroi, W., & Manosroi, J. (2010). Free radical scavenging and tyrosinase inhibition activity of oils and sericin extracted from Thai native silkworms (Bombyx mori). *Pharmaceutical Biology*, 48(8).

https://doi.org/10.3109/13880200903300212

- Maybee, D. V., Ink, N. L., & Ali, M. A. M. (2022). Novel Roles of MT1-MMP and MMP-2: Beyond the Extracellular Milieu. In *International Journal of Molecular Sciences*, 23(17). https://doi.org/10.3390/ijms23179513
- McCabe, M. C., Hill, R. C., Calderone, K., Cui, Y., Yan, Y., Quan, T., Fisher, G. J., & Hansen, K. C. (2020). Alterations in extracellular matrix composition during aging and photoaging of the skin. *Matrix Biology Plus, 8.* https://doi.org/10.1016/j.mbplus.2020.100041
- Mir, A. H., Qamar, A., Qadir, I., Naqvi, A. H., & Begum, R. (2020). Accumulation and trafficking of zinc oxide nanoparticles in an invertebrate model, Bombyx mori, with insights on their effects on immuno-competent cells. *Scientific Reports*, 10(1). https://doi.org/10.1038/s41598-020-58526-1
- Muhammad, A., He, J., Yu, T., Sun, C., Shi, D., Jiang, Y., Xianyu, Y., & Shao, Y. (2022). Dietary exposure of copper and zinc oxides nanoparticles affect the fitness, enzyme activity, and microbial community of the model insect, silkworm Bombyx mori. *Science of the Total Environment, 813.* https://doi.org/10.1016/j.scitotenv.2021.152608
- Niland, S., Riscanevo, A. X., & Eble, J. A. (2022). Matrix metalloproteinases shape the tumor microenvironment in cancer progression. In *International Journal of Molecular Sciences*, 23(1). https://doi.org/10.3390/ijms23010146
- Nilsson, B. O. (2021). Mechanisms involved in regulation of periodontal ligament cell production of proinflammatory cytokines: Implications in periodontitis. *Journal of Periodontal Research*, 56(2). https://doi.org/10.1111/jre.12823
- Pedro, A. R. V., Lima, T., Fróis-Martins, R., Leal, B., Ramos, I. C., Martins, E. G., Cabrita, A. R. J., Fonseca, A. J. M., Maia, M. R. G., Vilanova, M., &

Correia, A. (2021). Dectin-1-Mediated Production of Pro-Inflammatory Cytokines Induced by Yeast β -Glucans in Bovine Monocytes. *Frontiers in Immunology*, 12.

- https://doi.org/10.3389/fimmu.2021.689879
- Podbevšek, D., Ledoux, G., & Dular, M. (2022). Investigation of hydrodynamic cavitation induced reactive oxygen species production in microchannels via chemiluminescent luminol oxidation reactions. *Water Research*, 220. https://doi.org/10.1016/j.watres.2022.118628
- Puaratanaarunkon, T., & Asawanonda, P. (2022). A Randomized, Double Blinded, Split-Face Study of the Efficacy of Using a Broad Spectrum Sunscreen with Anti-Inflammatory Agent to Reduce Post Hyperpigmentation Inflammatory After Picosecond Laser. Clinical, Cosmetic and Investigational Dermatology, 15. https://doi.org/10.2147/CCID.S355329
- Qu, R., Zhang, Y., Ma, Y., Zhou, X., Sun, L., Jiang, C., Zhang, Z., & Fu, W. (2023). Role of the Gut Microbiota and Its Metabolites in Tumorigenesis or Development of Colorectal Cancer. *Advanced Science.* https://doi.org/10.1002/advs.202205563
- Safonova, L., Bobrova, M., Efimov, A., Davydova, L., Tenchurin, T., Bogush, V., Agapova, O., & Agapov, I. (2021). Silk fibroin/spidroin electrospun scaffolds for full-thickness skin wound healing in rats. *Pharmaceutics*, 13(10). https://doi.org/10.3390/pharmaceutics13101704
- Silva-Silva, J. V., Moragas-Tellis, C. J., Chagas, M. S. S., Souza, P. V. R., Moreira, D. L., Hardoim, D. J., Taniwaki, N. N., Costa, V. F. A., Bertho, A. L., Brondani, D., Zapp, E., De Oliveira, A. S., Calabrese, K. S., Behrens, M. D., & Almeida-Souza, F. (2022). Carajurin Induces Apoptosis in Leishmania amazonensis Promastigotes through Reactive Oxygen Species Production and Mitochondrial Dysfunction. *Pharmaceuticals*, 15(3). https://doi.org/10.3390/ph15030331
- Solano, F. (2020). Photoprotection and skin pigmentation: Melanin-related molecules and some other new agents obtained from natural sources. *Molecules*, 25(7). https://doi.org/10.3390/molecules25071537
- Su, D., Ding, S., Shi, W., Huang, X., & Jiang, L. (2019).
 Bombyx mori silk-based materials with implication in skin repair: Sericin versus regenerated silk fibroin. *Journal of Biomaterials Applications*, 34(1).

https://doi.org/10.1177/0885328219844978

- Sugiyono. (2019). *Metode Penelitian Kuantitatif, Kualitatif, dan R&D* (1st ed.). Penerbit Alfabeta.
- Tu, J., Jin, Y., Zhuo, J., Cao, X., Liu, G., Du, H., Liu, L.,

Wang, J., & Xiao, H. (2022). Exogenous GABA improves the antioxidant and anti-aging ability of silkworm (Bombyx mori). *Food Chemistry*, 383. https://doi.org/10.1016/j.foodchem.2022.132400

- Wang, M., Charareh, P., Lei, X., & Zhong, J. L. (2019). Autophagy: Multiple Mechanisms to Protect Skin from Ultraviolet Radiation-Driven Photoaging. *Oxidative Medicine and Cellular Longevity*. https://doi.org/10.1155/2019/8135985
- Wang, M., Shen, Y., Tan, Z., Yasen, A., Fan, B., & Shen, X. (2023). Metabolomics analysis of dietary restriction results in a longer lifespan due to alters of amino acid levels in larval hemolymph of Bombyx mori. *Scientific Reports*, 13(1). https://doi.org/10.1038/s41598-023-34132-9
- Wiraguna, A. A. G. P., Dianasari, R., & Pangkahila, W. (2019). The Topical Skin Application of Purple Corn Extract (Zea mays) Inhibited the Increase in MMP-1 Levels and Decreased Collagen in Wistar rats (Rattus norvegicus) Exposed to UV-B Rays. *Biomedical and Pharmacology Journal*, 12(1). https://doi.org/10.13005/bpj/1641
- Xu, Y., Wang, W., Ma, L., Cui, X., Lynch, I., & Wu, G. (2020). Acute toxicity of Zinc Oxide nanoparticles to silkworm (Bombyx mori L.). *Chemosphere*, 259. https://doi.org/10.1016/j.chemosphere.2020.1274 81
- Yeerong, K., Srivab, S., Somwongin, S., Punyovai, C., Chantawannakul, Ρ., Anuchapreeda, S., Prommaban, A., & Chaiyana, W. (2021). Skin potential irritation and antioxidant, anticollagenase, and anti-elastase activities of edible insect extracts. Scientific Reports, 11(1). https://doi.org/10.1038/s41598-021-02382-0
- Zhang, K., Shen, L., Wang, X., Yang, H., Zhang, X., Pan, G., Li, C., Ji, H., Abbas, M. N., Li, C., & Cui, H. (2021). Scavenger receptor C regulates antimicrobial peptide expression by activating toll signaling in silkworm, Bombyx mori. *International Journal of Biological Macromolecules*, 191. https://doi.org/10.1016/j.ijbiomac.2021.09.084
- Zhang, X., Zhu, X., Zhang, Y., Wu, Z., Fan, S., & Zhang, L. (2022). Comparative Transcriptome Analysis Identifies Key Defense Genes and Mechanisms in Mulberry (Morus alba) Leaves against Silkworms (Bombyx mori). *International Journal of Molecular Sciences*, 23(21). https://doi.org/10.3390/ijms232113519
- Zhao, G., Zhang, X., Cheng, J., Huang, X., Qian, H., Li, G., & Xu, A. (2020). Effect of Titanium Dioxide Nanoparticles on the Resistance of Silkworm to Cytoplasmic Polyhedrosis Virus in Bombyx mori. *Biological Trace Element Research*, 196(1), 290–296. https://doi.org/10.1007/s12011-019-01901-5

Zhao, P., Xia, F., Jiang, L., Guo, H., Xu, G., Sun, Q., Wang, B., Wang, Y., Lu, Z., & Xia, Q. (2018). Enhanced antiviral immunity against Bombyx mori cytoplasmic polyhedrosis virus via overexpression of peptidoglycan recognition protein S2 in transgenic silkworms. *Developmental and Comparative Immunology*, 87, 84–89. https://doi.org/10.1016/j.dci.2018.05.021