

**BIOACTIVE CONSTITUENTS OF SOPHORA MOLLIS AND THEIR
POTENTIAL ROLE IN CLINICAL PHARMACY AND PHYSIOTHERAPY-
BASED MANAGEMENT OF INFLAMMATORY CONDITIONS**

¹Muzammil Khalid, ²Tahir Jamshaid, ³Muhammad Naeem Qaisar, ⁴Ayesha Ali,
⁵Noor us Saba Mansoor (Ph.D.), ⁶Kamran Ayyub, ⁷Muhammad Kamran Khan,
⁸Joher Shujaat, ⁹Ummara Ijaz, ¹⁰Muhammad Usman Sarwar,
¹¹Muhammad Shoaib Sabir, ¹²Askarifard Jahromi Reza, ¹³Idress Hamad Attitalla,
¹⁴Muhammad Shafique, ¹⁵Imran Riaz Malik, ¹⁶Tahir Hafeez,
¹⁷Sahibzada Kazim Mehmood

¹Faculty of Pharmaceutical Sciences, Government College University Faisalabad (GCUF)

²Department of Pharmaceutical Chemistry, Faculty of Pharmaceutical Sciences, Government
College University Faisalabad (GCUF)

³College of Pharmacy, University of Sargodha

⁴Lahore Institute of Professional Studies, Lahore

⁵Assistant Professor, Department of Pharmacy Practice, Faculty of Pharmacy, Lahore Institute of
Professional Studies, Lahore

⁶The University of Faisalabad, Faisalabad

⁷Oxford College of Medical Sciences, Mandi Bahauddin

⁸Oxford College of Medical Sciences, Mandi Bahauddin

⁹Oxford College of Medical Sciences, Mandi Bahauddin

¹⁰Department of Eastern Medicine, Government College University Faisalabad, Pakistan

¹¹Department of Eastern Medicine, Government College University Faisalabad, Pakistan

¹²Dalian Medical University, China

¹³Omar Al-Mukhtar University, Faculty of Science, Department of Microbiology, Libya

¹⁴Institute of Microbiology, Faculty of Life Sciences, Government College University Faisalabad,
Pakistan

¹⁵Department of Biotechnology, Faculty of Sciences, University of Sargodha

¹⁶Tahir Holistic Healing and Research Institute, Mandi Bahauddin, Pakistan

¹⁷Faculty of Eastern Medicine, Hamdard University, Karachi

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Abstract

Pain, disability, and poor quality of life are often caused by inflammatory diseases, such as arthritis, injuries to the musculoskeletal system, and chronic inflammatory diseases. Natural products are being considered increasingly as adjunctive therapeutic agents due to their range of bioactive compounds and acceptable safety profiles. Sophora mollis is an alkaloid-rich, flavonoid-rich, phenolic rich and saponin-rich medicinal herb with promising anti-inflammatory, antioxidant and immunomodulatory effects in laboratory-based studies. These bioactive properties work through regulating key pathways involved in inflammation, inhibiting production of pro-inflammatory cytokines, reducing oxidative stress and altering cellular signaling systems involved in tissue inflammation and repair. With respect to clinical pharmacy, these pharmacologic activities suggest the possibility of developing Sophora mollis-based formulations as adjunctive therapies to traditional anti-inflammatory medications which might enhance therapeutic outcomes while decreasing an individual's likelihood of experiencing adverse drug reactions. Considering physiotherapy, the use of phytotherapeutic agents as part of a rehabilitation program may provide additional ways of managing pain related to inflammatory musculoskeletal injuries, enhancing the rate of tissue repair, and facilitating functional recovery in individuals with inflammatory musculoskeletal conditions. This literature reviews the phytochemical composition, pharmacologic mechanisms and clinical applications of Sophora mollis as an evidence-based therapeutic option for use in both clinical pharmacy and physiotherapy practice.

Introduction

The body reacts to infection, physical trauma, toxic chemicals, or other damaging stimuli by producing a natural defence mechanism known as inflammation. It is a multi-step response of the immune system, involving activation of immune cells and release of inflammatory mediators and multiple signalling pathways that work together to eliminate the invading pathogen and begin the healing of wounded tissue. Acute inflammation is beneficial and usually settles down once the threat is resolved, but the development of chronic inflammation can worsen the progression of many types of diseases (including rheumatoid arthritis, osteoarthritis, inflammatory bowel disease, atherosclerosis and cardiovascular disease, diabetes and neurodegenerative diseases) and significantly interfere with a patient's quality of life due to chronic pain; limited mobility; damage to tissues; and dysfunction. Conventional pharmacological therapies for the treatment of inflammation include non-steroidal anti-inflammatory drugs (NSAIDs) and corticosteroid drugs; however, long-term use of these drugs can lead to unwanted side effects including bleeding from the stomach

and/or intestines; kidney dysfunction; increased risk of cardiac complications; and immune suppression. Consequently, researchers are increasingly focused on using medicinal plants that may serve as safer, more sustainable therapeutic alternatives or as complementary treatments to conventional pharmacotherapy. Bioactive compounds found in many medicinal plants can modulate inflammatory pathways with minimal to no side effects. Some examples of these plants include: (Abd-Alla et al., 2021)

The Fabaceae family includes *Sophora mollis*, which has been utilized in many traditional indigenous healing systems to treat diseases that cause inflammation, such as skin disorders, infections caused by microorganisms, and pain. Research has demonstrated that *Sophora mollis* contains high amounts of biologically active phytochemicals such as flavonoids, quinolizidine alkaloids, phenolic compounds, saponins, tannins, and triterpenoids. These compounds have been shown to exhibit strong antioxidant and anti-inflammatory activities by scavenging reactive oxygen species (ROS), mitigating oxidative stress, and inhibiting the action of inflammatory

mediators that lead to tissue injury. Flavonoids inhibit cyclooxygenase (COX) and lipoxygenase (LOX) enzymes that are responsible for producing prostaglandins and leukotrienes, whereas alkaloids modulate the immune response through downregulation of pro-inflammatory cytokines (such as TNF- α , IL-1 β and IL-6). Collectively, this combination of actions provides a reduction in inflammation, greater cellular protection, and enhanced tissue repair. Continuous improvements in phytochemical analysis and molecular pharmacology provide increasing amounts of scientific support for the clinical effectiveness of *Sophora mollis*. The increasing popularity of botanically based medicines globally has prompted researchers to look for new/novel molecular entities (active compounds) from medicinal plants which can provide safer alternatives to synthetic pharmaceuticals. Therefore, *Sophora mollis* is an excellent candidate for providing a new/novel approach to the treatment of chronic inflammatory disorders. (Aly et al., 2020)

The growth of clinical pharmacy has gone beyond simply dispensing medications to now be an integral part of optimizing medication therapy, enhancing the safety of patients, and encouraging evidence-based health care practices. The rise of herbal remedies around the world has created a new responsibility for clinical pharmacists to assess the effectiveness and safety as well as quality, and the possible drug/herb interaction of the many medicinal plants used today. *Sophora mollis* presents a tremendous therapeutic potential, given its multiple pharmacological properties as bioactive ingredients, and their ability to compliment traditional anti-inflammatory medications. Plant-derived products may improve treatment outcomes by decreasing the amount of inflammatory cytokines and oxidative stress when consumed. As a result, these products may allow health care providers to delay or reduce the dosage of synthetic medications leading to few adverse events associated with pharmaceuticals. One of the primary responsibilities of a clinical pharmacist is to ensure that herbal remedies are being rationally used by providing education and advice about how to use herbal remedies, including the appropriate dosage, contraindications, toxicity, and potential

interactions with prescription medications. In addition, research continues to be undertaken by pharmaceutical scientists to develop improved dosage forms, enhanced delivery systems for herbal products, and the creation of standardized and novel formulations, all of which may enhance the bioavailability and therapeutic efficacy of the active constituents of *Sophora mollis*. Ongoing investigative research supports integrating herbal medicines into contemporary pharmacy and individualized therapeutic care. Therefore, understanding the pharmacological properties of *Sophora mollis* is critical to expanding its role in evidence-based pharmacy and multidisciplinary management of inflammatory disease. (Ullah et al., 2021)

Physiotherapy is a major part of an overall health care program for patients with muscular and inflammatory conditions. The goal of physiotherapy is to restore movement, relieve pain, increase joint mobility, and improve quality of life through effective physical rehabilitation. Therapeutic exercises, manual therapy, electrotherapy, stretching and strengthening programs, and educating the patient can all be used as physiotherapeutic treatments. However, persistent inflammation can delay tissue healing and extend the duration of pain, which reduces the success of the rehabilitation program. In recent years, there has been interest in using complementary therapies that include herbal medicine as adjunct therapies for physiotherapy because of the anti-inflammatory and antioxidative effect of many compounds found in plants. *Sophora mollis* is one such plant that presents a potential alternative therapy for physiotherapy because of its high levels of flavonoids, alkaloids, and phenolic compounds, all of which can inhibit substage inflammatory mediators, decrease oxidative damage, and promote tissue repair. Combined use of physiotherapy and phytotherapy has the potential to enhance functional recovery, alleviate pain, improve muscle function, and promote quicker healing. Such a combined approach may help in reducing the need for long-term pharmacotherapy, thus decreasing adverse effects associated with medications. (Boozari et al., 2019)

Bioactive agents derived from the plants within the genus *Sophora* possess anti-inflammatory properties as seen in newer studies using pharmacological methods to investigate how these bioactive agents function and how they affect the human body. The bioactive agents will modify key cellular signaling pathways (i.e., NF- κ B, MAPK, PI3K/Akt, and JAK/STAT) that are important mediators of excessive levels of pro-inflammatory cytokines, chemokines, prostaglandins, and reactive oxygen species, which ultimately lead to increased tissue damage and development of long-term disease processes. At the same time, the pharmacologically active phytochemicals that are found in *S. mollis* are able to inhibit these same cellular signaling pathways resulting in a decrease in the production of these same pro-inflammatory cytokines, chemokines, prostaglandins, and reactive oxygen species, thereby preventing the production of tissue damage due to oxidative stress. In addition, flavonoids and other phenolic compounds have significant antioxidant properties that can enhance our body's own innate antioxidant defenses by increasing the activities of major antioxidant enzymes such as superoxide dismutase, catalase, and glutathione peroxidase. The combined actions of these compounds are also helpful in tissue repair processes through reduction of cell death via apoptosis, and support the recovery process after tissues have been subjected to inflammatory injuries. In addition, new developments in pharmaceutical biotechnology, nanotechnology, and targeted drug delivery systems are generating more interest in the use of standardized formulations made from *S. mollis* extracts to increase therapeutic efficacy and improve patient compliance. Clinical applications will continue to be developed in the future to allow for better translation of positive laboratory data into clinical practice. (Krishna et al., 2012)

There is great potential for *Sophora mollis* as a medicine. However, many factors play a role in limiting its use in clinical settings. Most of the available research has been conducted on laboratory animals or in vitro, and there are few human clinical trials to support these findings. In addition to a lack of studies in humans, there is also considerable variability in the composition of

pharmacologically active phytochemicals found in the whole plant, depending on where the plant is grown, when the plant is harvested, how it is extracted, and how it is processed. The variation seen in the current body of the literature regarding the therapeutic efficacy of *Sophora mollis* due to differences in phytochemical composition is an example of how lack of standardization of dosage, toxicity, pharmacokinetic properties, and herb-drug interactions hampers the use of *Sophora mollis* as an alternative to synthetic medications in practice. Future research should concentrate on isolating the most potent phytochemical constituents of *Sophora mollis*, developing standardized pharmaceutical formulations for clinical use, and conducting randomized controlled studies on the efficacy and safety of the whole plant and its constituents to determine the most effective long-term therapeutic applications. The integration of evidence-based recommendations for including *Sophora mollis* in multidisciplinary health care will depend on collaboration among pharmacologists, clinical pharmacists, physiotherapists, toxicologists, and molecular biologists. If these collaborations can successfully achieve their goals, it is reasonable to expect the use of such partnerships to increase the safety, effectiveness, and personalization of the management of inflammatory diseases and reduce the dependence on synthetic products. As a result, *Sophora mollis* holds a tremendous amount of promise for future use as a medicinal plant in clinical pharmacy and physiotherapy, provided its efficacy can be substantiated through scientific research. (Fariás Cea & Iturriaga Vásquez, 2022)

Inflammation and Its Clinical Significance

Inflammation refers to the immune system's protective response triggered by injury, infection, or exposure to harmful stimuli and is comprised of a coordinated response involving immune cells, inflammatory mediators, and blood vessel changes to help clear away damaged tissue and begin the healing process. While it is necessary for acute inflammation to function normally as a defense mechanism in the human body, chronic and unregulated inflammation can damage the body and aid in creating several chronic diseases. Chronic inflammation has been linked to conditions like rheumatoid arthritis, osteoarthritis,

cardiovascular disease, metabolic disorders, and neurodegenerative diseases. Chronic inflammation biologically destroys tissue by continually releasing cytokines (small proteins released by cells), chemokines (small proteins that attract immune cells to sites of infection), and reactive oxygen species (toxic compounds containing oxygen) and is associated with the development of pain, decreased mobility, and decreased physiologic function. Managing inflammation in clinical practice is important to slow the progression of disease and improve the patient's quality of life. Conventional anti-inflammatory medications are often prescribed, but they can have negative side effects if taken long term, increasing the interest in finding natural therapies that are less harmful. Medicinal plants containing bioactive compounds are being evaluated for their ability to modulate inflammatory pathways and oxidative stress. (Pahwa et al., 2023)

A key factor contributing to many diseases is chronic inflammation. Chronic inflammation results from long-term changes in both the immune system's ability to control inflammation and the tissue function. Chronic inflammation does not arise immediately after exposure to a tissue-damaging substance and can continue for months or years; during this time period, chronic inflammation can cause damage to otherwise normal and healthy tissues. The diseases associated with chronic inflammatory activity can often be linked with the tissue-damaging effects of inflammatory mediators on the body's structural tissues. Two examples of chronic inflammatory diseases are rheumatoid arthritis (RA) and osteoarthritis (OA) (which have been linked with chronic inflammatory conditions) and both RA and OA are characterized by chronic inflammatory mediators resulting in damage to the joint structures, joint pain, joint stiffness, and decreased range of motion. In addition, chronic inflammation has been implicated in multiple metabolic disease processes where the effects of inflammatory mediators negatively impact the normal cellular activity associated with metabolism. Nevertheless, multiple cytokines (e.g., tumor necrosis factor-alpha [TNF- α], interleukins, and other signaling molecules) have been found to

stimulate an ongoing inflammatory response that contributes to the progression of disease and tissue destruction, while the generation of excessive amounts of reactive oxygen species (ROS) within tissues can worsen tissue damage associated with chronic inflammation and delay or impede the recovery of tissues damaged as a result of chronic inflammation. Due to the negative impact of chronic inflammation, it is clear that there is a need for the targeted treatment of inflammatory diseases in both clinical medicine and rehabilitation therapy. To effectively treat chronic inflammatory disease, both the inflammatory symptoms and the restoration of the normal functioning of the affected tissues must be addressed. As research studies demonstrate that many naturally occurring phytochemicals may provide positive anti-inflammatory properties by acting on the various inflammatory pathways and may afford significant protective effects against the chronic inflammation associated with multiple diseases. (Coussens & Werb, 2002)

Various molecular pathways regulate the body's immune system response and tissue repair. A few of the significant inflammatory signaling pathways in the body include nuclear factor kappa B (NF- κ B), mitogen-activated protein kinases (MAPK), and cyclooxygenase (COX) pathways. Each of these pathways regulates the production of inflammatory mediators (prostaglandins, cytokines, and enzymes involved in immune activation). Generally speaking, when functioning normally, these signaling pathways provide protection to the body and aid in healing the body from injuries. Nonetheless, the excessive stimulation of these pathways can lead to prolonged inflammation and contribute to the development of inflammatory diseases. The majority of conventional anti-inflammatories are examples of inhibitor drugs acting upon one or more of these pathways to achieve their effects of reducing pain and swelling; however, most of these drugs will eventually create unwanted side effects when taken over a long period. For this reason, there has been an increasing interest among researchers to explore natural alternative therapies derived from medicinal plants. There are many classifications of medicinal plant materials that may be beneficial in

regulating inflammatory signaling and reducing oxidative cellular damage, including but not limited to flavonoids, alkaloids, and phenolic compounds found in many herbs and spices. Understanding the various inflammatory molecular signaling mechanisms is crucial to establishing a scientific basis for evaluating medicinal plant-derived therapies. It is necessary to collate such knowledge to better understand the development of new interventions that can be incorporated into the clinical pharmacy practice and physiotherapy-based management of patients with inflammatory disorders. (Lawrence, 2009)

Treatment of inflammatory conditions often uses medicines, such as NSAIDs, corticosteroids and immunomodulators. Although effective in decreasing inflammation, controlling pain and assisting patients with their functional ability, the prolonged use of these drugs may have significant adverse side effects such as irritation of the gastrointestinal tract, problems with the kidneys, increased cardiovascular risk and/or altered immune function. As a result, researchers are increasingly interested in identifying safe and effective alternatives to traditional forms of therapy. Plants are known to be a source of biologically active components that may have anti-inflammatory, antioxidant and/or protectant properties. Phytochemicals derived from plants can exert their effects by a variety of different means, including inhibition of inflammatory enzyme activity, modulation of cytokine release and/or reduction of oxidative stress. With the growing interest in these materials due to their vast array of different chemical compounds and their potential to lead to the development of new therapeutic agents, it is imperative that scientists thoroughly evaluate their safety, appropriate dosages, quality and efficacy before clinical use. Combining evidence-based natural medicinal therapies into conventional medicine may provide valuable methods to manage inflammatory conditions and improve outcomes for patients receiving care for these conditions. (Newman & Cragg, 2020)

Physiotherapy has a vital role to play when it comes to the management of pain, function and repair. The presence of inflammation from injuries or diseases that affect muscles, joints or connective

tissue may cause stiffness, weakness, reduced range of motion and subsequently a decrease in performance. Physiotherapy treatment options such as therapeutic exercises, manual therapy, stretching, and strengthening programs, plus rehabilitation techniques, can all help restore normal function and improve quality of life. However, when inflammation persists too long or is too severe, healing may be delayed and the effectiveness of rehabilitation programmes may be limited. Therefore, it is important to control the inflammatory response as part of physiotherapy management plans. Therefore, combining rehabilitation with anti-inflammatory management strategies may result in more rapid recovery through decreased pain perception and increased rate of tissue repair. Recently, there has been an increased interest in using natural products that possess anti-inflammatory properties as adjuncts to standard physiotherapy treatment, which could improve clinical outcomes. Compounds found in medicinal herbs may modulate inflammation and oxidative stress; therefore they may help enhance the body's ability to heal itself. Research into the co-application of phytotherapy with physiotherapy may provide new therapeutic options for managing the inflammatory components of musculoskeletal conditions in a safe and effective manner. (Kisner et al., 2018)

A growing awareness of inflammation has led to the emergence of interdisciplinary strategies that include clinical pharmacy, pharmacology, physiotherapy and complementary therapies. Chronic diseases often result from inflammation; therefore, the identification of the most effective and safe therapeutic alternatives is a high priority to health care providers. Many medicinal plants contain many bioactive components that could affect the inflammation response mechanism and aid in managing the condition. Bioactive components found in plants, such as flavonoids, alkaloids, phenolic compounds, etc., have demonstrated antioxidant and immunomodulatory activities and have the potential to minimize injury caused by inflammation. An example of a plant being studied is *Sophora mollis*, which is being investigated for its ability to modulate inflammation via various mechanisms. Clinical

pharmacy research may include developing new formulations using enrolled patients and integrating physiotherapy with clinical pharmacy for improved pain management and functional recovery. Even though experimental evidence is promising, the literature lacks evidence of clinical trials to demonstrate the safety, efficacy, proper dosage, and long-term effects of botanical products. It is essential to conduct evidence-based research to validate traditional medicinal evidence supporting health care practices. The management of inflammatory-related conditions in the future may be achieved through a combination of conventional therapies and scientifically-based natural medicines for better patient care. (Atanasov et al., 2021)

Overview of *Sophora mollis*

Sophora mollis is one of the medicinal plants of the Fabaceae family that has long been used for its medicinal benefits through many systems of traditional herbal medicine. This plant is found throughout the world and has traditionally been used to treat many types of inflammatory, infectious and painful conditions, as well as other health related issues. Various parts of the *Sophora mollis* (e.g., leaves, roots, etc.) contain a wide range of different phytochemicals responsible for these medical benefits, such as alkaloids, flavonoids, phenolic compounds, terpenoids and a number of other secondary metabolites. These phytochemicals have been shown to have a variety of pharmacological effects such as antioxidant, anti-inflammatory, antimicrobial, and protective. *Sophora mollis*' medical properties may also primarily be due to its ability to modulate and control oxidative stress and the inflammatory response, which play a significant role in the development of many diseases. The recognition of *Sophora mollis* as one of the many sources of bioactive compounds available from nature has led to increased interest from the scientific community in conducting research on this plant as a potential source of drug discovery for pharmaceutical industry advancement. Ultimately, this type of research may help determine and develop ways in which to utilize *Sophora mollis* for clinical pharmacy and physiotherapy in the treatment of inflammatory disorders. (Ullah et al., 2021)

Sophora mollis has gained increasing attention among researchers because of its wide variety of chemically composed components and numerous biological effects. The various second metabolites contained in this plant play essential roles in protecting cells from oxidative damage and controlling the immune response. The alkaloids found in different *Sophora* species have therapeutic properties; the flavonoids and phenolic compounds primarily exert antioxidant and anti-inflammatory effects. The phytochemicals found in *Sophora mollis* are believed to affect the medium of different cellular pathways associated with inflammation through the reduction of inflammatory mediator production and support tissue repair processes. Due to its traditional use as an herbal remedy for chronic inflammatory illness, investigations into the potential health-related uses of *Sophora mollis* within conventional medicine are ongoing, particularly as a natural alternative and/or adjunct therapy for patients with chronic inflammatory diseases. In addition, as new methods for analysis and characterization of the phytochemical components of different *Sophora* species have emerged, greater understanding of how these active components interact with various therapeutic effects has also developed. However, there are significant factors affecting the biological activity of *Sophora mollis*, including changes in growth conditions (such as soil types), methods of extraction, and chemical profiles. Therefore, extensive pharmacologic and clinical study is needed to establish standard preparation protocols for use in healthcare and to ensure an adequate degree of safety and efficacy. (Abd-Alla et al., 2021) *Sophora mollis* has significant potential as a source of medicine; it may help identify new therapeutic substances for treating a wide range of conditions. The active compounds found in *Sophora mollis* possess numerous bioactive properties that can help combat disorders caused by inflammation. The antioxidant effects of the Phytochemicals can neutralize oxidative stress on the body, while their anti-inflammatory action can help reduce excessive immune reaction. Both of these qualities makes *Sophora mollis* as applicable in clinical pharmacy and physiotherapy approaches of managing inflammation while promoting the restoration of

functional capacity of patients. Antioxidant and anti-inflammatory substances derived from plants such as *Sophora mollis* may be the basis for improved and safer products in clinical pharmacy. Additionally, the potential anti-inflammatory properties of *Sophora mollis* may support rehabilitation in physiotherapy through the reduction of pain and promotion of tissue repair when used in a therapeutic manner. While the potential for the benefits of using *Sophora mollis* are encouraging, more studies are needed to further evaluate the safety, efficacy, and bioavailability of this medicine. In order to provide evidence-based health care for those individuals presenting with conditions caused by inflammation, additional research to define the molecular mechanisms and conduct clinical trials of *Sophora mollis* will be necessary to demonstrate the value of *Sophora mollis* as therapeutic agent. (Aly et al., 2020)

Sophora mollis has been extensively researched for its diverse range of pharmacological activities attributed to its extensive and complex array of phytochemical constituents. The phytochemicals present in *Sophora mollis* may have the ability to interact with all biological targets involved in inflammation, oxidative stress, and protection from cellular injury. Flavonoid and phenolic compounds are of particular interest because they have demonstrated beneficial effects on reducing oxidative insult and are effective at regulating inflammatory signalling pathways. These effects have the potential to reduce tissue injury and contribute to the maintenance of normal function. Numerous studies conducted on *Sophora* species have also demonstrated significant biological effects in the areas of antimicrobial, antioxidant, and immunomodulation. It is therefore useful to evaluate the therapeutic suitability of *Sophora mollis* as a potential source of therapeutic agents for modern drug discovery efforts. The identification of active phytochemicals from this species may also allow the production of natural products with superior safety profiles. However, the wide range of phytochemical variations that result from environmental conditions and extraction methods necessitates the need for standardisation. In addition, further scientific research on *Sophora*

mollis is essential to develop a greater understanding of its biological actions and to explore the potential utilization of *Sophora mollis* in healthcare, particularly the management of inflammatory diseases. (Boozari et al., 2019)

Sophora mollis is important therapeutically as it has a traditional use for medicinal purposes and the recent acceptance of such uses scientifically. Throughout the world, members of the *Sophora* genus have been used historically in various cultures to treat various diseases, including infections, pain, inflammation, etc. Pharmacological studies conducted today have proven some of these historical uses of *Sophora* peoples by showing that they contain bioactive compounds that produce the medicinal activity attributed to these plants. Alkaloids and flavonoids have been found to be among the bioactive compounds present in *Sophora mollis* and form a basis for investigating the plant's role in reducing inflammatory and oxidative stress. By influencing the immune response, these compounds may facilitate or enhance the body's ability to recover from injury by controlling the release of inflammatory substances and promoting the repair process. Therefore, *Sophora mollis* may provide an adjunctive treatment alternative to conventional medical therapies. Pharmacists who work with herbal formulations must understand the chemical and biological characteristics of *Sophora mollis* to ensure that they create safe products. Physiotherapy management may also benefit from the anti-inflammatory activity associated with *Sophora mollis* to achieve better rehabilitation results through improved pain control and enhanced healing of tissues. More clinical studies are necessary to ascertain the effectiveness of *Sophora mollis* and to develop appropriate guidelines for using the plant for therapeutic purposes. (Krishna et al., 2012)

While *Sophora mollis* presents significant potential for therapeutic use, several hurdles must be overcome before its general use in clinical settings. Human studies are limited, there is a lack of standardized extraction methods, and there is limited information regarding both the long-term safety of *Sophora mollis* and the presence of active compounds in it. The effectiveness of herbal

medicines will rely on the ability to accurately identify active compounds, determine appropriate dosages and maintain quality control during preparation of herbal medicines. Future studies should be directed toward understanding the pharmacokinetic properties, toxicity and clinical effectiveness of *Sophora mollis* by performing thoroughly designed pre-clinical and clinical studies. The use of advanced pharmaceutical techniques such as improved extraction methods and targeted delivery systems may also improve the efficacy of the bioactive constituents of *Sophora mollis*. Future collaborations between pharmacists, pharmacologists, physiotherapists and researchers can further establish evidence-based application of *Sophora mollis*. Further, the growing interest in natural therapies makes *Sophora mollis* a prime candidate for using to provide complementary therapies for inflammatory diseases. Further investigation into the potential of the various biological activities of *Sophora mollis* will enhance our understanding of its potential role as an intervention within contemporary health care and inclusion into multi-disciplinary interventions. (Fariás Cea & Iturriaga Vásquez, 2022)

Bioactive Constituents of *Sophora mollis*

Sophora mollis has a wide variety of bioactive ingredients contributing to its medicinal and pharmacological value. These phytochemicals include many different types which can be organised into groups such as alkaloids, flavonoids, etc. The alkaloids present in *Sophora* species are some of the most relevant compounds to their biological properties and include properties such as anti-inflammation, antimicrobial, and antioxidants. The flavonoids and phenolic compounds present in *Sophora mollis* have an important role in the protection of cells from damage caused by oxidative stress. They do this by neutralizing reactive oxygen species. In addition to their direct cell protection, they also modulate inflammation by modulating the production of Cytokines and other mediators involved in Immune reactions. The combination of all of these bioactive compounds may lead to the synergistic effects that will strengthen the overall biological activity of *Sophora mollis*. The chemical profile of *Sophora mollis* should be understood to assess its

therapeutic properties and to develop standardised formulations for use in healthcare, specifically with regard to the treatment of inflammatory disorders. (Ahmad et al., 2021)

Sophora mollis contains alkaloids, which are the primary components responsible for its medicinal effects. Quinolizidine alkaloids have shown various effects on bioactivity including modulating immune responses and regulating reactivity of inflammation. They may modulate signalling pathways related to inflammation and oxidative processes. The antioxidant capacity of *Sophora mollis* is due in part to its flavonoids, which include flavonols and flavones. Additionally, phenolic compounds are critical to their ability to provide cellular protection from oxidative damage. Chronic inflammatory diseases often occur as a result of oxidative stress; thus, the anti-oxidant properties of the various constituents in the plant will likely reduce tissue injury. The combination of the three classes of compounds enhances the overall therapeutic potential of *Sophora mollis*. Because of these properties, the plant has been investigated for possible use as a complementary strategy to treating conditions related to inflammation. Additional studies are warranted to identify the specific active molecules and their mechanism of actions. (Zhang et al., 2020)

The key point to note about the anti-inflammatory properties of *Sophora mollis* is that flavonoid and phenolic constituents play a significant role in this function due to their ability to inhibit the production or activity of pro-inflammatory mediators by inhibiting different enzymes and the molecular pathways related to inflammation. By acting as inhibitors of COX and LOX, flavonoids and phenolic compounds have the ability to decrease the amount of inflammatory mediators such as prostaglandins and leukotrienes generated from their respective pathways. In addition to these mechanisms of action, they may also modulate other important signaling pathways, such as NF- κ B, which regulate the expression of pro-inflammatory genes. By decreasing pro-inflammatory signaling, flavonoids and phenolic compounds will reduce tissue damage and promote recovery. They may also provide an additional protective mechanism via their antioxidant properties, which will assist in

maintaining cellular homeostasis by protecting against oxidative injury. Thus, these properties make *Sophora mollis* a potential source of natural products for pharmaceutical application. The natural presence of these bioactive compounds supports their use as potential therapeutic targets in the development of chronic inflammatory disease and improving the health of individuals using natural therapies. (Kumar et al., 2022)

Sophora mollis possesses many classes of compounds, including alkaloids, flavonoids, saponins, and terpenoids, all of which would have a diverse range of biological activities. Terpenoid compounds are known to be both protective and to have regulatory roles; saponins exhibit numerous biological properties, including antioxidant, anti-microbial and immunomodulating activities. The presence of these two groups of bioactive compounds results in a complex phytochemical profile that may account for the numerous traditional and experimental uses attributed to *Sophora mollis* throughout history. These two classes of compounds may also combine together to provide enhanced anti-inflammatory responses, promote healing of damaged tissues and improve the overall health of the immune system. In clinical pharmacy, knowledge about these bioactive constituents is essential to evaluating the safety and efficacy of plant-based products and potential for drug interactions prior to developing a product. In physiotherapy based management, selected compounds from *Sophora mollis* with anti-inflammatory and antioxidant properties have the potential to assist in the rehabilitation of injuries by reducing inflammatory responses, promoting recovery and modulating pain associated with musculoskeletal disorders. Extraction methodologies and environmental variables can significantly affect the concentration of these compounds, so it will be necessary to standardize formulations for therapeutic use. (Sharma et al., 2021)

The therapeutic effect of presages is the interaction between the plant's bioactive components and the biological systems. The majority of plant-based compounds do not operate via solely one use; on the contrary, they work collectively across multiple

pathways involved with inflammatory responses, immune reactions and cellular protection functions in organisms. *Sophora mollis* has demonstrated that it possesses an array of complex pharmacological properties which may include its alkaloids, flavonoids and phenolic components; these combinations of constituents may work together for the reduction of inflammatory responses; enhancement of antioxidant defense activity; as well as providing tissue protection against damage. Thus, through this multi-targeted activity, the potential uses of *Sophora mollis* in assisting with the management of complex inflammatory disorders is significantly enhanced. Recent developments within the field of phytochemical analysis have provided researchers with the technology to isolate and characterise active compounds derived from *Sophora mollis* as well as assist researchers in developing an understanding of their pharmacological properties. These data are also vital when transitioning from historical medicinal knowledge to practice within an evidence-based healthcare framework. Although the biological effects of *Sophora mollis* seem to indicate there may be benefit from future research examining the individual contributions of its plant's constituents to determine their clinical relevance, necessitating that extensive studies must be conducted on the area surrounding *Sophora mollis* before definitive conclusions can be definitively drawn. (Singh et al., 2023)

Identification and evaluation of bioactive constituents from *Sophora mollis* will enable future development of natural therapeutic products. The plant's promising phytochemical profile, while highlighting its potential medicinal value, presents challenges with standardization, determination of dose, evaluation of toxicity, and validation of clinical efficacy. Variations in environmental conditions during growth, method of harvesting plant material, and extraction technique could affect the quantity of active constituents and their associated biological activity. Therefore, more advanced research methodologies will be needed to verify that *Sophora mollis* products have consistent quality and efficacy. Future research efforts should focus on clinical

trials (pharmacokinetics and pharmacodynamics), molecular analysis (biochemical assays), and development of new formulations to maximize therapeutic benefits. Phytochemistry, clinical pharmacy, and rehabilitation sciences may provide a basis for establishing the role of *Sophora mollis* in contemporary health care. Bioactive constituents derived from *Sophora mollis* will be an important source for developing anti-inflammatory agents and can serve as an adjunct to currently available therapies for managing inflammatory conditions in a safe manner. (Rahman et al., 2019)

Anti-inflammatory and Antioxidant Mechanisms

Oxidative Stress & Inflammatory Process Biologically Related, Inflammation Oxidative Stress & Their Involvement in the Development of Many Types of Illnesses. Immune Cells Are Responsible for Regulating the Body's Immune Response Against Harmful Stimuli Through Immunological Signalling Molecules & Process Activations (Inflammation); When This Immune Response Has Been Overly Activated, There Continually Produced Cytokines, Enzymes & Reactive Oxygen Species (Creating) Damage to Tissues. *Sophora mollis* Is A Type Of Plant That Is Been Used As A Source Of Anti-Inflammatory Agents For The Purposes of Reducing Osteoporosis, Reducing Pain Associated With Arthritis, Supporting The Body's Immune System (General), & Preventing Damage From High Levels of Oxidative Stress. *Sophora mollis* Has Many Bioactive Components (Flavonoids, Alkaloids, & Phenolics) That May Help Regulate The Production of Cytokines (Inflammation) & Support Cellular Protection (Antioxidants). *Sophora mollis* Antioxidants May Neutralise Free Radicals, Reduce Oxidative Stress, & Protect The Body's Cells. The Essentially Antioxidant Action May Help To Maintain Balance Between Reactive Oxygen Species (Oxidative Damage) & Natural Defences (Neutralisation). Also, Anti-Inflammatory Actions Are Connected With How Important Mediators That Promote Pain, Swelling, & Tissue Damage Are Regulated. All Combined (Antioxidants & Anti-Inflammatories Together) May Contribute To The Therapeutic Potential of *Sophora mollis*, As It Relates To Referral to Clinical Pharmacy & Physiotherapy-Based Care

Management Approaches By Understanding These Processes. (Ali et al., 2020)

The ability of *Sophora mollis* to modulate important inflammatory signaling pathways appears to be its main mechanism by which it exerts anti-inflammatory effects. The bioactive compounds found in *Sophora mollis* may prevent the activation of the transcription factor nuclear factor-kappa B (NF- κ B), which is one of the most important transcription factors that regulates the expression of genes that participate in inflammation. NF- κ B is activated when present in the cell in excess, and when it is activated excessively, it also stimulates the production of pro-inflammatory cytokines such as tumor necrosis factor-alpha (TNF- α), interleukin-1 β (IL-1 β) and interleukin-6 (IL-6) and therefore contributes to chronic inflammation. Consequently, phytochemical compounds from *Sophora mollis* may inhibit the release of these inflammatory mediators and return the immune system to homeostasis. Additionally, *Sophora mollis* may modulate the activity of the enzymes cyclooxygenase (COX) and lipoxygenase (LOX), which are involved in producing inflammatory mediators such as prostaglandins (PGs) and leukotrienes (LTs). By modulating these pathways, *Sophora mollis* may lessen the deleterious effects of inflammation on tissues, and currently support the healing process. Collectively, these actions illustrate that *Sophora mollis* possesses great potential to serve as a natural source of anti-inflammatory agents that could enhance current methods of treating inflammation. (Patel et al., 2021)

Antioxidant action from *Sophora mollis* is due to the phenolic and flavonoid content being potent free radical regulators. The cellular processes that generate oxidising agents also produce reactive oxygen species creating free radicals, that can contribute oxidative stress which leads to damage to proteins, lipids and DNA. Chronic inflammation develops due to oxidative stress causing increased cellular injury, resulting in chronic inflammatory disorder via inflammatory signaling pathways. Antioxidants contained within *Sophora mollis* may assist to prevent oxidative damage by neutralising reactive oxygen species, and/or increasing the activity of the body's own

antioxidant enzymes such as: superoxide dismutase, catalase and glutathione peroxidase. The antioxidants' protective effects maintain the cell's structural integrity, while providing benefits for healing injured tissues. The anti-inflammatory properties of these compounds are also indicative of their therapeutic relevance for managing disease caused by oxidative imbalance, as well as creating oxidative stress. Therefore, *Sophora mollis*'s antioxidant potential provides strong therapeutic relevance for the management of conditions associated with oxidative stress and chronic inflammatory processes. (Mehta et al., 2022)

The combined effect of *Sophora mollis*' antioxidant and anti-inflammatory mechanisms enhances its overall biological efficacy. Oxidative stress and inflammation create a continuous loop in many inflammatory diseases; each process stimulates the other. Elevated oxidative stress causes the activation of inflammatory signaling, and the resulting inflammation increases the generation of reactive oxygen. The bioactive constituents of *Sophora mollis* could break this cycle through the reduction of oxidative damage and the ability to modulate the release of inflammatory mediators. *Sophora mollis*' varied phytochemical constituents (flavonoids and other phytochemicals) provide a dual mechanism of protection against oxidative damage; they also serve to regulate the immune response. These dual actions could help repair tissues, decrease cell damage, and promote recovery after inflammatory states. Due to this multi-targeting effect, *Sophora mollis* is an excellent candidate for future pharmaceutical investigation. The use of these mechanisms will help clinical pharmacists develop plant-based formulations for patients, while physiotherapists will use these mechanisms to promote recovery and enhance the function of patients during rehabilitation from inflammation and oxidative stress. (Hassan et al., 2019)

Molecular studies indicate that plant compounds possessing antioxidant and anti-inflammatory properties can regulate multiple cellular signalling pathways impacted by disease progression. *Sophora mollis* constituents are capable of regulating several pathways such as MAPK, to regulate various immune pathways or immune responses and

oxidative balance. By changing these pathways, the constituents may reduce hyper-inflammation and increase the ability of cells to defend against free radicals or oxidative stressors. This effect is important for chronic disease processes whereby long-term inflammation leads to continued degeneration of tissue. The protective role of *Sophora mollis* will assist in maintaining normal physiological function through a decrease in oxidative damage. While experimental data support these mechanisms, additional research is necessary to establish the precise molecular targets and therapeutic implications of each individual component of *Sophora mollis*. A clearer understanding of these modes of action will aid in developing evidence-based therapeutic strategies utilizing *Sophora mollis* for use as a complementary therapy for management of inflammatory disease processes. (Verma et al., 2023)

Sophora mollis's anti-inflammatory and antioxidant effect gives it potential to be used by health professionals today and provides research validation. This plant can affect multiple biochemical pathways due to its variety of bioactive compounds, including inhibition of inflammatory mediators, alleviation of oxidative damage, and enhancement of cellular defense mechanisms. Utilized with existing medical therapies and rehabilitative techniques, *Sophora mollis* may provide improved results for those with inflammatory diseases. However, before *Sophora mollis* becomes widely used in a therapy regimen, operational challenges, such as standardization, dosage adjustment, safety assessments, and limited availability of clinical data, must be resolved. Future studies should focus on randomized clinical trials, as well as advanced pharmacological investigations to validate its safety and efficacy and develop consistent treatment protocols. The combination of natural products with conventional pharmaceutical and physical therapy might lead to improved approaches to manage inflammatory diseases more safely and effectively. *Sophora mollis* shows promise as a twofold medical resource for controlling inflammation and supporting tissue healing. (Khan et al., 2021)

Challenges, Limitations, and Future Perspectives

Despite the growing attention from scientists focusing on *Sophora mollis* many obstacles and constraints are due to its translation from traditional use to science-based proof of clinical use. While the plant's use has been shown to exhibit some encouraging results in anti-inflammatory, antioxidant and medicinal qualities most trials have either taken place in laboratories or on animals. Limited research has been conducted on humans; therefore the plant's definitive outcome, safety and quantity required for therapeutic use remain unknown. Other important impediments are found in the diversity of phytochemicals found in plants based on geographic location, environmental conditions, farming practices and methods of extraction. This diversity may have an impact on the amount of active chemicals present and create inconsistent biological activity resulting in different outcomes when used in medicine. To ensure quality, reproducibility and reliable results all plant extracts need to be standardized. Additionally, the mixture of bioactive compounds within the plant makes it difficult to determine what compounds are contributing to specific pharmacological properties of the plant. Additional studies utilizing advanced analytical and molecular methods are necessary to facilitate the development of *Sophora mollis* as a scientifically documented medicinal resource. (Raza et al., 2020)

The absence of complete data regarding the pharmacokinetics, toxicity, and long-term safety of *Sophora mollis* significantly limits its therapeutic development. Because natural products often have a variety of chemical constituents that can react with each other and/or with prescription or over-the-counter medications, the study of possible herb-drug interactions is particularly critical in clinical pharmacy practice, where patients frequently combine herbal remedies with their prescribed medications. However, because of limited information on the absorption, distribution, metabolism, and elimination of active compounds from *Sophora mollis*, its clinical application is limited. Moreover, preparation methods may alter both the stability and bioavailability of important phytochemicals found in *Sophora mollis*.

Therefore, advanced formulation techniques and controlled delivery systems may improve the effectiveness of products containing *Sophora mollis* as active ingredients. Conducting toxicological studies to determine the safe therapeutic range for *Sophora mollis* is also necessary to help prevent possible adverse effects. By addressing these issues, the transformation of traditional knowledge into valid therapeutic strategies will occur. Future research should focus on conducting comprehensive pharmacological evaluations; developing standardized products; and completing formal clinical trials to determine the actual therapeutic efficacy of *Sophora mollis*. (Iqbal et al., 2021)

Sophora mollis has a lot of Future opportunities in clinical pharmacy and physiotherapy for the treatment of disorders, with multiplicity of bioactive constituent and in many instances bioactivity. Clinical pharmacy may see the development of a natural anti-inflammatory formulation for use with other conventional treatments. Antioxidant and immunomodulatory characteristics could aid in reducing the possibility for complications associated with inflammation and improving patient quality of life. Controlling inflammation is also a key area in physiotherapy to reduce pain, increase range of motion, and assist in tissue repair. A combination therapeutic option incorporating phyto-therapeutics into rehabilitation programs could be an all-encompassing option to treat inflammatory conditions of the musculoskeletal system. However, to be clinically successful, both scientific evidence and interprofessional collaboration will need to take place. Pharmacologists, Pharmacists, Physiotherapists and researchers need to work collaboratively to determine the therapeutic benefits and safety of *Sophora mollis* as a treatment option. Future research will be needed to investigate whether *Sophora mollis* can produce positive outcomes in conjunction with conventional therapies to improve recovery and quality of life for patients with chronic inflammatory disorders. (Shah et al., 2022)

Separate parts of future research studies using *Sophora mollis* can be utilised to identify which of the plant's active constituents have therapeutic

potential as well as to clarify their mode of action at a molecular level. Metabolomics (a modern scientific method of identifying metabolite levels), molecular docking and using advanced phytochemical analysis will be essential to discovering these important compounds and determining their mode of action against biological targets. By conducting research using metabolomics, molecular docking and phytochemical analyses, scientists can also learn how the plant regulates oxidative stress, inflammation and the immune response by examining how the active compounds interact with the human body. Future researchers should also evaluate how different extraction methodologies affect the concentration and activity of the bioactive compounds contained in the plant's extracts. The use of improved extraction methodologies and new pharmaceutical technologies may enhance the stability and efficacy of plant-derived products. Establishing a method of creating standardised extracts, which have consistent chemical characteristics, will be critical to ensuring that these products can be used effectively in clinical trials in the future. Combining traditional medicinal knowledge with modern scientific methodologies will ultimately provide researchers with tools to achieve safer and more effective therapeutic modalities. Additionally, the development and application of these proposed methodologies could increase the likelihood of establishing evidence-based healthcare practices in the management of chronic inflammatory diseases using *Sophora mollis*. (Naseer et al., 2019)

One of the key future uses of *Sophora mollis* relates to appropriate regulation, quality control and evidence of clinical effectiveness. Evaluation of herbal medicine is critical to assure that they are as pure, safe and effective as possible before being recommended for widespread use; however, since no standardised method exists for production of such products, there could be significant variation in the quality of products produced and their effectiveness. Regulatory guidelines are needed to set out acceptable limits for active compounds, contaminants and dosages. Furthermore, health care providers must have access to accurate information to ensure that they can provide their

patients with accurate recommendations regarding the use of herbal products. Increasingly, health care professionals are becoming more knowledgeable about how to use herbal products appropriately so as to avoid misuse and improve health outcomes. future development of new therapies based on *Sophora mollis* will depend upon providing consistently reliable production techniques, as well as generating more clinical research evidence about them. The future combination of natural compounds with contemporary pharmaceutical technology may also facilitate personalising individual therapy and multidisciplinary approaches towards treating inflammatory diseases. (Gupta et al., 2023)

In conclusion, *Sophora mollis* is an important medicinal plant that has great potential to be developed therapeutically in the future. It exhibits a wide variety of phytochemical compounds and exhibits many activities such as anti-inflammatory, antioxidant, and protective characteristics; thus, further investigations on these activities will provide strong evidence for continued research of this compound. However, many obstacles exist in using it as a regular part of clinical medicine, including the need for additional clinical research regarding its use, lack of dosage guidelines, considerable variation in chemical composition, and the lack of safety information. Continued research on this herb should include randomized clinical trials, molecular studies, toxicity evaluations, improved formulation methods, and collaboration between different scientific disciplines to demonstrate the role that *Sophora mollis* may play in contemporary health care. Combining traditional herbal medicine with scientific evidence will lead to developing safer and more effective methods of treating inflammatory diseases that incorporate this herb. With ongoing advances in science, *Sophora mollis* could play an essential role as a natural source of therapeutic innovation within clinical pharmacy and physiotherapy. (Malik et al., 2021)

Conclusion

The medicinal plant *Sophora mollis* has great potential to manage inflammation due to the presence of a variety of bioactive compounds, in addition to its many pharmacological activities.

The presence of alkaloids, flavonoids, phenolic compounds and other phytochemicals in this plant provide a source of anti-inflammatory, antioxidant and immunomodulatory effects; thus, this may allow *Sophora mollis* to modulate the inflammatory pathway, decrease oxidative stress and assist with tissue protection. Clinical Pharmacy has the potential to create complimentary types of natural therapeutic agents using *Sophora mollis* as a foundation to assist or be used instead of conventional therapies. Physiotherapy-based management may also use *Sophora mollis* to support pain relief, tissue healing and functional restoration. Despite the numerous promising results found in experimental studies, many limitations, such as insufficient clinical trials, lack of standardization of extracts, safety concerns and limited pharmacokinetic data continue to be barriers to the use of *Sophora mollis*. Future research must give attention to a continuum of clinical trial data to establish clinical validation and an understanding of molecular mechanisms, quality control and the development of formulations to create therapeutically reliable products. Integrating traditional knowledge with modern science can further develop and enhance the use of *Sophora mollis* as a safe and effective complementary option for managing inflammation. Overall, *Sophora mollis* has the potential to be the basis for further research in Clinical Pharmacy and Rehabilitation Sciences.

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