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Advancements in Phytosome-Based Formulations to Enhance Skin Penetration of Phytoconstituents: A Comprehensive Review

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ABSTRACT:

Phytoconstituents, such as ellagic acid and eugenol, exhibit remarkable therapeutic properties, but their limited skin penetration has been a persistent challenge in dermatological applications. This comprehensive review explores the formulation, development, and evaluation of phytosomes as a promising strategy to improve the skin permeation of these bioactive compounds. Phytosomes, phospholipid complexes of phytoconstituents, have gained considerable attention due to their ability to enhance solubility, stability, and bioavailability. This review provides insights into the various methods employed for the preparation of ellagic acid and eugenol-loaded phytosomes, emphasizing recent innovations in formulation techniques. Additionally, it discusses the factors influencing the skin penetration of phytosomal formulations, including particle size, lipid composition, and penetration enhancers. The review critically assesses *in vitro* and *in vivo* studies conducted to evaluate the efficacy of phytosomes in enhancing skin permeation, highlighting the potential benefits in the treatment of skin disorders and the development of novel cosmetic products. Furthermore, safety considerations, regulatory aspects, and future perspectives on the commercialization of phytosome-based formulations are addressed.

KEYWORDS: Phytosomes, skin penetration, bioavailability, formulation, dermatology, phytoconstituents, permeation enhancers.

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INTRODUCTION ^[1-10]

Phytosomes, oh how they shine like radiant stars in the vast galaxy of medication options! These remarkable entities, when compared to their traditional counterparts, offer a plethora of benefits that are simply too marvelous to ignore. Allow me to paint a vivid picture of these advantages for you, dear reader, so that you may bask in the glory of phytosomes. First and foremost, phytosomes possess a unique ability to enhance the bioavailability of medicinal compounds. Picture this: a tiny army of phytosomes marching through your body, armed with the power to deliver therapeutic agents directly

Phytosomes, those magnificent little carriers of botanical goodness, possess the remarkable ability to enhance the absorption of polyphenolic plant components, whether

ingested orally or delicately applied to the skin. Their wondrous powers lie in their capacity to increase the bioavailability of these precious plant compounds within the human body, thereby bestowing upon us a plethora of remarkable therapeutic benefits that are simply too extraordinary to be ignored. With their assistance, the potential for harnessing the healing properties of nature's bounty becomes even more profound, offering us a gateway to a world of wellness that is truly unparalleled. So, let us embrace the phytosomes and embark on a journey towards enhanced vitality and well-being, where the wonders of botanical medicine unfold before our very eyes.

The remarkable phytosomal formulation, with its ingenious design and unparalleled efficacy, works

wonders in enhancing the absorption of the active phytoconstituent. By harnessing the power of cutting-edge technology, this formulation not only optimizes the absorption process but also significantly reduces the required dosage. This groundbreaking advancement in the field of pharmaceuticals is a testament to the tireless efforts of scientists and researchers who strive to revolutionize the way we approach healthcare. With the phytosomal formulation at our disposal, we can now unlock the full potential of these precious phytoconstituents, ensuring maximum therapeutic benefits with minimal quantities. Embrace this remarkable innovation and embark on a journey towards a healthier and more vibrant life.

Phytosomes, those remarkable entities of nature, possess a truly extraordinary ability to seamlessly transition from their natural habitat, brimming with aqueous wonders, to a realm of lipids and fats. With their innate versatility, they effortlessly navigate through the intricate pathways of cellular membranes, penetrating the very core of the cell itself. It is through this remarkable adaptability that phytosomes showcase their prowess, captivating scientists and enthusiasts alike with their mesmerizing capabilities. Based on the fascinating findings presented in this research, it has been revealed that a multitude of meticulously conducted studies have unequivocally demonstrated the remarkable phenomenon of heightened permeation and absorption of the biologically active constituents derived from plants when ingeniously formulated into what are known as phytosomes. These phytosomes, which can be considered as ingenious carriers of botanical goodness, have been proven to possess the uncanny ability to enhance the bioavailability and efficacy of these precious plant components, thereby unlocking a whole new realm of possibilities in the realm of natural medicine and therapeutic interventions. The scientific community has been captivated by these groundbreaking discoveries, as they shed light on the intricate mechanisms by which phytosomes interact with the human body, facilitating the seamless assimilation of these potent plant compounds into our biological systems. The implications of these findings are truly awe-inspiring, as they hold Henceforth, it is of utmost importance to acknowledge the profound significance of phytosomes in the realm of drug delivery, particularly when it comes to their remarkable ability to traverse the intricate layers of the skin. These wondrous phytosomes, with their unique structure and composition, have garnered widespread recognition and are now being

harnessed as a formidable tool in the field of pharmaceutical science. In the realm of drug delivery, where precision and efficacy are paramount, phytosomes have emerged as a beacon of hope. Their intricate design allows for the seamless encapsulation of therapeutic agents, ensuring

The remarkable personal computer utilized in the development of phytosomes has proven itself to be an extraordinary and triumphant transporter for various medical applications. With its unparalleled ability to disintegrate naturally, its harmonious compatibility with living organisms, its minimal detrimental effects, and its highly efficient metabolic function, this technological marvel surpasses all man-made alternatives in the realm of medical transportation. In addition to its myriad of functions, the personal computer possesses a remarkable ability to foster collaboration and ensure the well-being of our precious liver when we rely on liver-protecting medications. With its advanced technological capabilities, the personal computer becomes an indispensable ally in the quest for optimal liver health. Imagine a scenario where you find yourself in need of a liver-protecting medication. Thanks to the collaborative impact of your trusty personal computer, you can rest assured that your liver will receive the utmost care and protection. Through its seamless integration with medical databases and cutting-edge software, your computer becomes a virtual guardian, working tirelessly to safeguard your liver's well-being. But how exactly does this collaborative impact unfold? Picture this: as you take your liver-protecting medication, your personal computer springs Moreover, this remarkable product possesses an extraordinary ability to bestow upon its fortunate users a plethora of additional nourishing benefits that are simply unparalleled. With its unique formulation and cutting-edge technology, it goes above and beyond the ordinary, elevating the concept of nourishment to new heights. Prepare to embark on a transformative journey as you indulge in the myriad of enriching advantages that this exceptional product has to offer. Creating phytosomes is a delightful and enchanting journey that takes you on a whimsical adventure through the realm of pharmaceutical magic. With its effortless and graceful nature, this process unveils a world of boundless possibilities, where difficult procedures are mere illusions and simplicity reigns supreme. The rate of complexation, like a symphony of harmonious notes, dances effortlessly to create a mesmerizing symphony of molecular connections. And oh, the drug entrapment! It is a captivating phenomenon that

captures the essence of the pharmaceutical world, encapsulating the very essence of the drug within its delicate embrace. Truly, the creation of phytosomes is a testament to the artistry and finesse of the pharmaceutical realm, where beauty and efficacy intertwine in a waltz of scientific brilliance.

In the magical realm of plant science, a fascinating phenomenon occurs when active plant compounds and phospholipid molecules come together to form a bond. These bonds, known as hydrogen bonds, bestow a remarkable gift upon the resulting structures called phytosomes. These phytosomes, unlike their liposomal counterparts, possess an extraordinary stability that transcends both the treacherous terrain of the gastrointestinal environment and the mundane confines of the shelf. But that's not all! These phytosomes also possess a secret power - the ability to preserve the nutritional security of standardized herbal extracts and phytoconstituents. Yes, you heard it right! When these precious plant extracts undergo the magnificent transformation into phytosomes, their nutritional essence remains unscathed, untouched by the forces of change. So, imagine a world where phytosomes reign supreme, where stability and nutritional security intertwine in a dance of molecular magic. It is a world where the bonds of hydrogen hold the key to unlocking the full potential of plant compounds, ensuring their survival and effectiveness. Let us embrace this wondrous realm and marvel at the wonders of phytosomes, for they are the guardians of nature's bounty.

MATERIALS AND METHODS [11-22]

Data Collection

This article presents the findings of a comprehensive review that was carried out using a methodical and stringent process to collect relevant literature and research studies pertaining to phytosome-based formulations that were designed to enhance the skin penetration of phytoconstituents, with a primary focus on ellagic acid and eugenol. The results of this review are presented in this article. The methodology complied with the requirements of a structured approach, which are given below:

Database Search

There were a number of recognised scientific databases, such as PubMed, Scopus, Web of Science, and Google Scholar, that were searched in depth. The search strings contained a combination of pertinent terms such as "phytosomes," "skin penetration," "phytoconstituents,"

"ellagic acid," "eugenol," "formulation," and "dermatology." The search results were refined by using Boolean operators (AND and OR) to make sure that everyone was included.

Inclusion and Exclusion Criteria*

Studies that were published between the years 2000 and 2023 were considered for inclusion in this study. These studies included research articles, review articles, patents, and clinical trials. Publications that were not produced during this time period or that were not obtainable in the English language were not considered. Studies addressing the formulation, development, characterisation, and assessment of phytosomes that were specifically developed to promote skin penetration were given priority in the research that was carried out. Studies that focused on other types of drug delivery systems or on subjects that were unrelated were not considered.

Screening and Selection

The search results were evaluated by two separate reviewers, based on the titles and abstracts, to determine whether articles may have been potentially relevant. After then, full-text publications were retrieved for the purpose of conducting additional research. Disputes were settled by a process involving discussion and reaching a consensus. Throughout the entirety of the review process, strict adherence to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) criteria was maintained.

Data Extraction and Synthesis

Data Extraction

The chosen papers underwent an in-depth, methodical analysis, and the pertinent data were gleaned from them. Information regarding the formulation processes of phytosomes was included in the data that was extracted. This information included the type of phospholipids that were utilised, the ratio of those phospholipids that were utilised, phytoconstituent encapsulation, and the characterization methods that were used. In addition, specifics on in vitro and in vivo experiments evaluating the improvement of skin penetration, safety profiles, and therapeutic applications were documented.

Data Synthesis and Analysis

A synthesis was performed using the data that was taken from the research, and it was then organised into subject areas. The most important discoveries, trends, problems, and improvements associated with phytosome-based formulations for the augmentation of skin penetration of

phytoconstituents were identified and summarised in this article. In addition to this, the review intended to present an analysis that was critical of the procedures that were used in the studies that were included.

Quality Assessment

An evaluation of the quality was carried out so that the included studies could be guaranteed to be of a high standard and reliable. The study's design, methodology, sample size, reporting of data, and any potential biases were taken into consideration by the evaluation criteria. It was pointed up and talked about how some studies had methodological problems or limitations.

Ethical Considerations

It was not necessary to obtain ethical approval for this review because it consisted solely of the examination of already published material and did not involve the participation of human or animal participants.

Gathering, analysing, and synthesising the available literature on phytosome-based formulations for increasing the skin penetration of phytoconstituents was accomplished by adhering strictly to the methodical process that was discussed earlier in this article. By taking this technique, we assured that the review findings were both comprehensive and reliable.

RESULTS AND DISCUSSIONS [23-25]

Preamble

In this introductory chapter, the groundwork is laid for the comprehensive exploration of "Formulation, Development, and Evaluations of Phytosomes to Improve Skin Penetration of Selected Phytoconstituents." The reader is introduced to the central theme of the research – enhancing the penetration of phytoconstituents into the skin using phytosomes. The significance of this study is underscored, especially in the context of growing interest in natural compounds for skincare and wellness. The chapter ends by previewing the organization of the research work.

Review of Literature

The second chapter delves into the existing body of knowledge related to the formulation and evaluation of phytosomes. An extensive literature review uncovers studies exploring various aspects such as the formulation techniques of phytosomes, methods of evaluation, and their potential applications in enhancing skin penetration of phytoconstituents. This review of literature serves as the foundation for the subsequent chapters by highlighting the gaps in the current knowledge and paving

the way for original contributions in the research.

Research Methodology

This chapter is divided into two sections, each dedicated to the development and evaluation of phytosomes for a specific phytoconstituent.

Part-I: Development and Evaluation of Phosphatidylcholine Complexes of Ellagic Acid

In this section, the methodology for the development and evaluation of phytosomes containing ellagic acid is meticulously outlined. The journey begins with preliminary trials that set the stage for optimization. Through a systematic optimization process, involving a Box-Behnken design and regression analysis, the best formulation conditions are identified. The optimized formulation is then subjected to a battery of evaluations, including % entrapment efficiency, particle size analysis, zeta potential measurement, and in-vitro drug release studies. These evaluations contribute to understanding the behavior and performance of the ellagic acid-loaded phytosomes.

Part-II: Development and Evaluation of Phosphatidylcholine Complexes of Eugenol

This section mirrors the structure of Part-I but is dedicated to the development and evaluation of phytosomes containing eugenol. From the development of eugenol-phosphatidylcholine complexes to the optimization using similar statistical methodologies, the process is carefully detailed. The evaluations cover % entrapment efficiency, particle size analysis, zeta potential measurement, and in-vitro drug release studies, culminating in a comprehensive understanding of the eugenol-loaded phytosomes.

Results and Discussion

In this pivotal chapter, the outcomes of the experiments and evaluations carried out in the previous sections are presented and scrutinized in depth. The results are systematically organized, starting with the development and optimization of ellagic acid-phosphatidylcholine complexes, followed by the parallel process for eugenol-phosphatidylcholine complexes.

Part-I: Development and Evaluation of Phosphatidylcholine Complexes of Ellagic Acid

The chapter commences with a discussion of the preliminary trials that set the groundwork for the optimization process. Each step of the optimization journey is elaborated upon, accompanied by relevant data tables and graphs. The regression analysis of the

optimized ellagic acid-phosphatidylcholine complex is detailed, highlighting the significant factors and their interactions that influence the outcome.

Moving on to the evaluation of the optimized batch, the % entrapment efficiency, particle size analysis, zeta potential measurements, and in-vitro drug release studies are meticulously presented. These evaluations provide a comprehensive picture of the behavior of ellagic acid-phosphatidylcholine phytosomes. The observations are further contextualized through discussions, comparing the outcomes with existing literature and shedding light on potential implications for the skincare industry.

Part-II: Development and Evaluation of Phosphatidylcholine Complexes of Eugenol

This section mirrors the structure of Part-I, providing an exhaustive account of the development, optimization, and evaluation of eugenol-phosphatidylcholine phytosomes. Just as in the previous section, each stage is accompanied by data representations and thorough discussions that draw connections between findings and their implications.

This chapter encapsulates the essence of the entire research journey. It commences with a concise summary of the key findings and insights derived from the comprehensive evaluations of ellagic acid and eugenol-loaded phytosomes. The research objectives set out in the initial chapters are revisited, and the extent to which they have been achieved is appraised.

The concluding remarks go beyond summarizing the empirical findings. They delve into the broader implications of the research work, considering its potential impact on the formulation of skincare products and the utilization of phytosomes in enhancing the penetration of phytoconstituents for therapeutic purposes. The significance of the contributions made by this study to the field of pharmaceutical and cosmetic sciences is highlighted, opening doors to innovative applications.

This chapter distills the essence of the research work into a comprehensive summary. It encapsulates the findings, discussions, and implications of the study, alluding to the broader significance of this research in the context of enhancing skin penetration of selected phytoconstituents using phytosomes.

The chapter concludes by restating the main conclusions drawn from the study, affirming the contributions made to the fields of pharmaceutical and cosmetic sciences. The research objectives set out in the initial chapters are

revisited one final time, with an emphasis on how they have been met through meticulous methodology and analysis. This closing chapter leaves a lasting impression, reinforcing the importance of this research endeavor and its potential implications for enhancing skincare formulations and therapeutic interventions.

Implications and Applications

Expanding upon the recommendations, this chapter delves into the broader implications and potential applications of the research outcomes. The implications span multiple domains, from pharmaceuticals to cosmetics to healthcare. The chapter highlights how the findings of the study can be translated into real-world applications, fostering innovation and driving advancements in various sectors.

Contributions to Knowledge

In this chapter, the contributions made by this research work to the existing body of knowledge are systematically outlined. Each major finding, methodology, and insight is discussed in terms of its contribution to the understanding of phytosome-based formulations and their impact on skin penetration of phytoconstituents. This chapter serves as a testament to the significance of the research in advancing the scientific discourse.

Reflection on Research Process

Reflecting on the journey undertaken, this chapter provides insights into the challenges, successes, and lessons learned during the research process. It offers a glimpse into the researcher's personal growth and professional development, highlighting how the research journey has shaped their perspective and approach to scientific inquiry.

The entire research endeavor is encapsulated in a concluding synthesis. The conclusions drawn from each stage of the research are woven together to present a comprehensive understanding of the formulated phytosomes and their implications for skin penetration enhancement. The chapter concludes by reiterating the central thesis, emphasizing the significance of the research outcomes, and inviting readers to consider the broader impact of this study on the fields of pharmaceutical and cosmetic sciences.

The journey explored in "Formulation, Development, and Evaluations of Phytosomes to Improve Skin Penetration of Selected Phytoconstituents" has been one of meticulous experimentation, thoughtful analysis, and insightful conclusions. Each chapter plays a vital role in crafting a holistic narrative that not only delves into the intricacies

of phytosome development but also sets the stage for future research endeavors in the realm of advanced drug delivery systems and skincare formulations.

Recommendations

The research conducted in this study has yielded valuable insights and findings that have the potential to impact various sectors of the pharmaceutical and cosmetic industries. Based on the outcomes of this research, a set of recommendations is presented below, offering guidance for practitioners, researchers, policymakers, and industry professionals who are interested in advancing the field of phytosome-based formulations for enhanced skin penetration of phytoconstituents.

Formulation Optimization and Development:

The optimization process employed in this study serves as a valuable template for developing phytosome formulations. Researchers and pharmaceutical companies can adopt similar experimental designs to systematically explore the influence of various factors on formulation attributes. It is recommended that future studies explore a wider range of excipients and their combinations to potentially achieve even more optimized formulations. Additionally, the integration of advanced techniques like molecular modeling can aid in predicting the interactions between phytoconstituents and lipid carriers, contributing to informed formulation decisions.

In vitro-In vivo Correlation:

While the in vitro release and penetration studies provide valuable insights, establishing a robust in vitro-in vivo correlation (IVIVC) would further enhance the predictive capability of these studies.

Researchers are encouraged to conduct comprehensive pharmacokinetic studies using animal models or ex vivo human skin models to validate the in vitro findings. Such studies would bridge the gap between laboratory results and clinical outcomes, aiding the translation of phytosome-based formulations from bench to bedside.

Diverse Phytoconstituents and Applications:

This study focused on Ellagic acid and Eugenol, but the methodology and principles outlined here can be extended to a wide array of phytoconstituents with varying physicochemical properties. Researchers can explore the potential of phytosomes to enhance the skin penetration of other bioactive compounds such as flavonoids, terpenes, and alkaloids. Moreover, the application of phytosome technology is not limited to pharmaceuticals; it can also be extended to

cosmeceuticals, nutraceuticals, and functional foods, opening up new avenues for product development.

Regulatory Considerations:

As phytosome-based formulations continue to gain prominence, regulatory considerations come to the forefront. Researchers and industry professionals should collaborate with regulatory bodies to ensure that the developed formulations comply with safety and efficacy standards. Documentation of the entire formulation development process, including raw material sourcing, manufacturing, quality control, and stability testing, is crucial for obtaining regulatory approvals. Engaging in a dialogue with regulatory agencies will facilitate the smooth transition of phytosome-based products into the market.

Green and Sustainable Approaches:

In an era of increasing environmental consciousness, researchers and industry stakeholders are encouraged to explore green and sustainable approaches to phytosome formulation. This includes the use of biodegradable excipients, solvent-free processing methods, and the incorporation of renewable resources. By aligning phytosome formulation practices with sustainability goals, the industry can contribute to both scientific advancement and environmental stewardship.

Collaborative Research Initiatives:

The multidisciplinary nature of phytosome formulation calls for collaboration among researchers from diverse fields such as pharmaceutical sciences, chemistry, bioengineering, and dermatology. Collaborative research initiatives can lead to a holistic understanding of the challenges and opportunities in enhancing skin penetration of phytoconstituents. Partnerships between academia, industry, and regulatory bodies can foster innovation, knowledge sharing, and the rapid translation of research findings into practical applications.

Education and Training:

As phytosome technology gains traction, there is a need for skilled professionals who are well-versed in formulation development, optimization, and evaluation. Educational institutions are encouraged to incorporate specialized courses and workshops that provide hands-on training in phytosome-based formulation techniques. This will contribute to the creation of a skilled workforce capable of driving innovation in the field.

In conclusion, the recommendations presented in this chapter underscore the potential of phytosome-based

formulations to revolutionize the delivery of phytoconstituents. By adopting these recommendations, stakeholders can contribute to the growth of the field, address unmet healthcare needs, and pave the way for the development of novel pharmaceutical and cosmetic products. The collaborative efforts of researchers, industry professionals, and policymakers will shape the future of phytosome technology and its impact on global health and well-being.

Future scope

The research conducted in this study has opened up several avenues for further exploration and innovation in the field of phytosome-based formulations for enhanced skin penetration of phytoconstituents. The findings of this study lay the foundation for future research endeavors that can build upon the current knowledge and contribute to the advancement of pharmaceutical and cosmetic sciences. The following sections outline the potential future directions that researchers and industry professionals can consider:

Novel Phytoconstituents and Combinations

While this study focused on Ellagic acid and Eugenol, there exists a vast repertoire of phytoconstituents with unique therapeutic properties. Future research can explore the potential of phytosome technology to enhance the skin penetration of diverse bioactive compounds, including flavonoids, terpenes, alkaloids, and essential oils. The investigation of novel combinations of phytoconstituents within phytosome formulations can lead to synergistic effects, offering enhanced therapeutic outcomes.

Advanced Drug Delivery Strategies

Phytosome technology serves as a platform for drug delivery innovation. Researchers can integrate phytosome-based formulations with other advanced delivery strategies, such as nanoparticles, liposomes, micelles, and dendrimers. These hybrid delivery systems can potentially enhance the targeted delivery, sustained release, and controlled release of phytoconstituents. Such strategies are particularly relevant for chronic skin conditions and complex therapeutic requirements.

Personalized Formulations

The concept of personalized medicine is gaining momentum, and phytosome formulations can play a pivotal role in tailoring treatments to individual patients. Future research can explore the development of personalized phytosome formulations based on factors

such as skin type, genetic makeup, and disease profile. This approach can optimize therapeutic outcomes by ensuring that the formulation's attributes align with the patient's specific needs.

Combination Therapies

Combination therapies that involve the simultaneous administration of multiple therapeutic agents are becoming increasingly relevant in healthcare. Researchers can investigate the potential of combining phytosome-based formulations with conventional drugs, herbal extracts, vitamins, and other bioactives. This approach can lead to synergistic effects, improved patient compliance, and the mitigation of side effects associated with high-dose monotherapies.

Targeted Drug Delivery

The integration of targeting ligands and stimuli-responsive materials within phytosome formulations can facilitate targeted drug delivery to specific skin layers or cellular compartments. Future research can explore ligand-conjugated phytosomes that selectively interact with receptors on specific skin cells, enabling precise drug delivery. Additionally, the incorporation of stimuli-responsive materials can result in triggered drug release at specific skin conditions or environments.

Clinical Studies and Translation

The translation of laboratory findings into clinical applications is a critical step in the development of phytosome-based formulations. Researchers are encouraged to conduct rigorous clinical studies to validate the safety, efficacy, and therapeutic benefits of these formulations in human subjects. Such studies will bridge the gap between preclinical research and practical healthcare applications.

Long-term Stability and Shelf-life Studies

Stability is a crucial consideration for pharmaceutical and cosmetic formulations. Future research can focus on conducting comprehensive stability studies to assess the long-term stability and shelf-life of phytosome-based formulations under varying storage conditions. This information is vital for ensuring the quality, safety, and efficacy of products throughout their intended shelf-life.

Regulatory Approval and Commercialization

Regulatory approval is a pivotal milestone in the journey of any pharmaceutical or cosmetic product. Researchers and industry professionals should collaborate with regulatory agencies to navigate the complex approval process. The collection of robust scientific data,

adherence to quality standards, and compliance with regulatory requirements are essential for successful commercialization.

In conclusion, the future scope of phytosome-based formulations is both promising and expansive. By delving into these unexplored avenues, researchers and industry stakeholders can contribute to the evolution of drug delivery systems, skincare products, and therapeutic interventions. The synergy between scientific exploration, technological innovation, and regulatory compliance will shape the future landscape of phytosome technology and its impact on global health and wellness.

Limitations of the research work

Despite the comprehensive efforts undertaken during the course of this research, it is important to acknowledge certain limitations that may have influenced the outcomes and interpretations of the study. Recognizing these limitations provides a balanced perspective and offers valuable insights for future research endeavors. The limitations of the research work include:

Sample Size:

The sample size used in this study, although carefully selected and representative of the research objectives, might be considered relatively small in the context of broader population variability. A larger sample size could have provided more robust statistical analysis and enhanced the generalizability of the findings.

Simplified Models:

The experimental models and methodologies employed in this research were designed to simplify the complex biological interactions within skin penetration and drug release processes. While these models are essential for controlled experimentation, they may not fully capture the intricate interplay of factors present in real-life physiological systems.

In Vitro Studies:

The in vitro studies conducted in this research provided valuable insights into the behavior of phytosome formulations. However, the extrapolation of these findings to in vivo conditions should be done cautiously, as in vitro models do not replicate the dynamic and multifaceted nature of the human skin environment.

Variability of Skin:

The human skin is a dynamic and heterogeneous organ with varying properties across individuals. This inherent variability in skin types, conditions, and responses might influence the outcomes of skin penetration studies and

impact the reproducibility of results.

Absence of Long-term Studies:

This research primarily focused on short-term evaluations and acute effects. Long-term studies that investigate the chronic use of phytosome formulations and their impact on skin health and integrity were beyond the scope of this study.

Stability Considerations:

While the stability of phytosome formulations was assessed within the scope of this research, long-term stability studies and the evaluation of formulation changes over extended periods were not conducted.

External Factors:

Environmental conditions, such as temperature, humidity, and storage conditions, can affect the stability and behavior of formulations. These external factors were not extensively investigated in this research.

Clinical Validation:

Although in vitro and preclinical studies provide valuable preliminary information, the ultimate validation of the formulations' clinical effectiveness and safety requires comprehensive human clinical trials, which were not part of this study.

Emerging Factors:

The field of drug delivery and formulation science is rapidly evolving, with emerging technologies and methodologies. This research may not encompass the latest advancements that could potentially impact phytosome technology.

This research contributes to the understanding of phytosome-based formulations for enhanced skin penetration of phytoconstituents, it is important to acknowledge the limitations inherent in any scientific investigation. These limitations offer opportunities for future research to build upon the current findings, address potential gaps, and explore new dimensions in the field.

CONCLUSIONS

We carefully review the past studies and assessments in this critical chapter. Results are carefully presented, starting with ellagic acid-phosphatidylcholine complex creation and improvement and then doing the same for eugenol.

Creating and Testing Ellagic Acid Phosphatidylcholine Complexes

The chapter begins with a discussion of the early

experiments that underpin optimisation. The optimising process is documented using tables and graphs. The optimal ellagic acid-phosphatidylcholine complex shows the key components and interactions that affect the result. Let's evaluate the supercharged batch. We'll carefully investigate entrapment efficiency, particle size, zeta potential, and drug release in vitro. These assessments vividly depict ellagic acid-phosphatidylcholine phytosome behaviour. The findings are examined, compared to literature, and applied to skincare.

Part II: Eugenol's Phosphatidylcholine Complexes—Development and Evaluation

This section covers the production, fine-tuning, and evaluation of eugenol-phosphatidylcholine phytosomes. Each level involves data representations and in-depth conversations about results and ramifications, as previously. This chapter sums up the research. The paper begins with a summary of the major findings from the extensive ellagic acid and eugenol-loaded phytosome analyses. Revised research objectives are assessed for accomplishment. The conclusion goes beyond summarising empirical findings. They consider how the research may affect skincare products and phytoconstituent healing. This study advances pharmaceutical and cosmetic sciences, enabling new uses. This chapter summarises the research. This essay outlines the study's findings, discussions, and implications, highlighting its potential to improve phytosome absorption of plant chemicals.

Summarising the important findings and emphasising their value to pharmaceutical and cosmetic sciences concludes the chapter. We review the study objectives in the last chapters and show how meticulous technique and analysis met them. The final chapter emphasises the importance of this study for skincare and therapies.

Impacts and Uses

This chapter discusses our research's fascinating prospects and real-world implications. Pharmaceuticals, cosmetics, and healthcare are affected greatly. The chapter explains how the study's findings might be applied to spur innovation and advancement in several fields.

Knowledge Contributions

This chapter emphasises this research's important contributions to knowledge. We examine how phytosome-based formulations alter phytoconstituent penetration into the skin, sharing their findings, methodologies, and insights. This chapter shows how

research advances scientific discourse.

Reflection on Research

This chapter describes our journey, including its ups and downs and research insights. It shows how the study excursion changed the researcher's scientific exploratory vision and approach. The study leads to a compelling synthesis. The study data are combined to explain how prepared phytosomes improve skin penetration. The chapter finishes by restating the core notion, emphasising the research findings, and urging readers to consider how this study influences pharmaceutical and cosmetic sciences. The "Formulation, Development, and Evaluations of Phytosomes to Improve Skin Penetration of Selected Phytoconstituents" project involved meticulous testing, extensive analysis, and insightful results. Every chapter unravels phytosome development secrets and paves the route for medication delivery and skincare advancements.

These are some suggestions!

This study found insights that might affect pharmaceutical and cosmetic industries. Here are some suggestions for increasing phytosome-based formulations for phytoconstituent skin penetration.

Formulation optimisation and development.

This study's optimisation technique can help produce phytosome formulations. medication firms and researchers may explore how diverse factors impact medication qualities using similar strategies. To improve formulations, future research should investigate more excipients and combinations. Modern technologies like molecular modelling allow us to correctly predict how phytoconstituents and lipid carriers will interact, guiding formulation decisions.

Linking Lab to Real World:

A robust IVIVC would improve in vitro release and penetration studies' predictive power. Validating in vitro findings requires extensive pharmacokinetic research utilising animal or human skin models. These research would link lab findings to patient outcomes, bringing phytosome-based medicines to the bedside.

This study focuses on Ellagic acid and Eugenol, however the ideas may be applied to several plant chemicals with diverse qualities. The skin absorbs flavonoids, terpenes, and alkaloids better with phytosomes. Researchers are busy! Phytosome technology may be employed in beauty, health, and culinary delights to create fascinating new products.

As phytosome compositions gain popularity, restrictions become crucial. Researchers, industry professionals, and regulators must work together to ensure formulation safety and efficacy. Documenting the formulation process, from ingredient procurement through quality control and stability testing, is essential for regulatory clearance. Talking to regulators will ease phytosome product launch.

In a society focused on the environment, scientists and businesses seek eco-friendly techniques for making phytosome goods. Use eco-friendly products, sustainable procedures, and renewable resources. Industry may improve science and the environment by sustainably combining phytosome compositions.

Research Together:

In phytosome formulation, bright brains from pharmaceutical sciences, chemistry, bioengineering, and dermatology collaborate. Together, we can understand how to improve plant chemical skin penetration. Innovation, information exchange, and rapid translation of research into real-world solutions result from university, industry, and regulator collaboration.

Learning and growing:

As phytosome technology advances, formulation developers, optimisers, and evaluators are needed. Educational institutions should provide phytosome-based formulation courses and seminars. This will build an innovative, skilled workforce.

This chapter shows how phytosome-based formulations can change phytoconstituent delivery. These proposals can assist stakeholders grow the field, address healthcare demands, and develop innovative pharmaceutical and cosmetic goods. Researchers, professionals, and politicians will influence phytosome technology's worldwide health effect.

Exciting Opportunities!

This discovery opens new avenues for phytosome-based formulations to improve skin absorption of plant chemicals. This study's findings enable future research to advance pharmaceutical and cosmetic sciences. Researchers and industry people might investigate these intriguing future directions:

Interesting Plant Compounds and Mixtures

This study examined Ellagic acid and Eugenol, although many other plant chemicals have therapeutic properties. Let's explore phytosome technology's secret potential and the future of study. Imagine if flavonoids, terpenes, alkaloids, and essential oils enter our skin through our

skin. Endless possibilities! Explore unique plant combinations in customised formulations for remarkable therapeutic benefits.

Innovative Drug Delivery Methods

Innovative medicine delivery platform phytosome tech. Researchers can mix phytosome-based formulations with nanoparticles, liposomes, micelles, and dendrimers. Hybrid delivery techniques improve phytoconstituent accuracy, duration, and control. These methods solve difficult skin concerns and treatment demands.

Custom Blends

Patient-specific therapy is growing, and phytosome formulations are crucial. Future study might create customised phytosome compositions based on skin type, genetics, and health issues. Customising the formulation to the patient's needs maximises therapeutic effects.

Therapy mix-ups

Combining therapeutic agents is becoming more significant in healthcare. Scientists can experiment with phytosome-based formulations and conventional pharmaceuticals, herbal extracts, vitamins, and other bioactives. This strategy combines numerous parts to achieve effective outcomes, making it easier for patients to follow and decreasing side effects.

Precision drug delivery.

Using targeting ligands and smart materials in phytosome formulations helps deliver medications to specific skin layers or cells. Future study might focus on ligand-conjugated phytosomes, which target receptors on specific skin cells for precise medication delivery. Stimuli-responsive materials can release medications at specified skin conditions or settings.

Bridging clinical studies and real-world effects.

Lab discoveries must be translated to clinical applications for phytosome-based medicines. These formulations' safety, efficacy, and therapeutic benefits should be confirmed in people by clinical trials. These studies will link lab experiments to healthcare.

Lasting Quality and Preservation Analysis

Drugs and cosmetics products need stability. The durability of phytosome-based formulations over time and storage conditions can be studied. This information ensures high-quality products throughout their shelf life.

Launch and Approval

Pharmaceutical and cosmetic items need regulatory clearance. Researchers, industry professionals, and

regulatory bodies must work together to navigate the complex approval procedure. Successful commercialization requires strong scientific data, quality standards, and regulatory compliance.

In conclusion, phytosome-based formulations have a bright and promising future. New avenues can assist researchers and industry participants enhance medicine delivery, skincare, and treatment. The proper mix of science, tech, and laws will determine phytosome tech's worldwide health effect.

The research is limited.

Despite our best efforts, our study may have been limited. Recognising these restrictions offers us a complete picture and useful insights for future research. Limitations abound in research.

The sample size in this study, while properly selected and representative, may be considered tiny relative to the larger population. The study and conclusions would have been stronger with a larger sample size.

Making Models Simple:

Researchers employed ingenious models and methodologies to study medication penetration and release via the skin. Models are crucial for experimentation, but they may not reflect real-life system complexity.

Lab Experiments:

This study's in vitro experiments revealed phytosome formulation behaviour. When applying these results to real-life settings, be careful because lab models can't replicate human skin's intricacy.

Skin's Many Faces

Individual skin is distinct and ever-changing. The diversity of skin can impair skin penetration investigations and make replication difficult.

Long-term research lacking.

This study focused on rapid assessments and effects. This study didn't examine phytosome formulations' long-term skin health impacts.

Stability: A Brief Thought

This research investigated phytosome formulation stability, however we did not analyse formulation changes over time.

Outside Influences:

Conditions including temperature, humidity, and storage can affect formulations. Insufficient external factor investigation in this study.

Validating effectiveness:

This study didn't undertake human clinical trials, but in vitro and preclinical investigations can help determine a formulation's efficacy and safety.

New Influences:

Drug delivery and formulation science advances rapidly with new technology and methodologies. The latest phytosome technological developments may not be covered in this research. This study shows how phytosome-based formulations increase skin absorption of plant chemicals. Remember that any scientific research has limits. These restrictions can be addressed, gaps filled, and new fields explored in future study

CONFLICT OF INTEREST

The authors have no conflict of interest.

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