## **Review Article**



# Innovations Shaping the Future: Advanced Approaches in Extended-Release Tri-Layer Tablets

# Seetharam Gude<sup>1\*</sup>, Akshay varun. B<sup>2</sup>, J.M. Parameswara<sup>2</sup>

Department of Pharmaceutics, Vignan pharmacy college, Vadlamudi, Guntur, 522213, AP, India.
Department of Pharm.D, St. Johns college of pharmaceutical sciences, Yemmiganur, Kurnool, 518360, AP, India.
\*Corresponding author's E-mail: Seethasaiofficial@gmail.com

Received: 15-04-2024; Revised: 28-05-2024; Accepted: 06-06-2024; Published on: 15-06-2024.

#### ABSTRACT

The pharmaceutical arena is currently undergoing a transformative phase in drug delivery systems, with extended-release tri-layer tablets positioned at the forefront of pioneering innovation. In this article, we navigate through the cutting-edge developments propelling the refinement of this sophisticated drug delivery system. The exploration encompasses breakthroughs in nanotechnology, 3D printing, smart polymers, combination therapies, personalized medicine, data-driven approaches, and *In Vitro-In Vivo* Correlation (IVIVC). Collectively, these advancements illuminate a trajectory towards precision, heightened efficiency, and a more patient-centric paradigm in the realm of pharmaceutical research.

Keywords: Extended-release, Tri-layer tablets, Personalized medicine, Combination therapy, Nano technology, Smart polymers.

## INTRODUCTION

xtended-release tri-layer tablets epitomize an advanced drug delivery system meticulously engineered to ensure the sustained release of medications over a protracted duration. Characterized by a stratified structure, each of the three layers assumes a distinct role in meticulously regulating drug release kinetics. In response to the burgeoning demand for heightened precision, operational efficiency, and patientcentricity within the pharmaceutical landscape, extendedrelease tri-layer tablets have garnered substantial prominence. Positioned as a response to these imperatives, these tablets integrate innovative technologies and formulation strategies to redefine therapeutic delivery paradigms. Critical to the enhanced efficacy of extended-release tri-layer tablets are the integration of nanotechnology. This pivotal factor involves the assimilation of nanoparticles such as liposomes and polymeric nanoparticles. This strategic integration not only achieves superior drug solubility and bioavailability but also facilitates targeted drug delivery, thus affording a heightened degree of control over drug release profiles and, subsequently, augmenting therapeutic outcomes. The advent of 3D printing technology has catalyzed a revolutionary transformation in the manufacturing process of extended-release tri-layer tablets. This avantgarde approach confers a granular level of control over the tablets' geometry and composition. The consequential outcome is the ability to create bespoke drug delivery solutions tailored to the unique needs of individual patients. Smart polymers, characterized by their responsiveness to environmental stimuli, introduce a dynamic dimension to drug release in extended-release trilayer tablets. These polymers adeptly respond to physiological changes within the body, orchestrating an adaptive drug release profile. The net result is an optimization of therapeutic outcomes while concurrently minimizing undesirable side effects. The tablets serve as a versatile platform for combination therapies, facilitating the concurrent release of multiple drugs with distinct kinetics. This holistic approach is particularly germane in addressing complex medical conditions, invariably enhancing treatment efficacy and, notably, promoting heightened patient compliance. The integration of personalized medicine into the design ethos of extendedrelease tri-layer tablets is synonymous with tailoring formulations to align seamlessly with individual patient characteristics. This individualized approach serves to maximize treatment efficacy while concurrently mitigating adverse effects, thereby adhering steadfastly to the overarching goal of patient-centric care. Data-driven methodologies, notably employing artificial intelligence and machine learning, optimize the intricate process of formulation design and the prediction of drug release profiles in extended-release tri-layer tablets. These sophisticated techniques engender a streamlined drug development process, culminating in formulations that are both efficient and precise. Furthermore, the establishment and refinement of In Vitro-In Vivo Correlation (IVIVC) models significantly enhance the predictability and reliability of extended-release tri-layer tablets. This is achieved through the establishment of meaningful correlations between in vitro characteristics and subsequent in vivo performance. Such models, therefore, contribute indispensably to a more accurate assessment of drug release kinetics and the overarching therapeutic efficacy of these innovative pharmaceutical entities. 1,2



Figure 1: Tri-layer tablets



Available online at www.globalresearchonline.net ©Copyright protected. Unauthorised republication, reproduction, distribution, dissemination and copying of this document in whole or in part is strictly prohibited.

## Nanotechnology Integration:

In recent advancements, nanotechnology has become a pivotal element in the evolution of extended-release trilayer tablets, presenting a paradigm shift in drug delivery strategies. This integration is marked by the deliberate incorporation of nanoparticles, including liposomes and polymeric nanoparticles, into the intricate structure of the tri-layer tablet. The primary objective of this integration is to significantly enhance the solubility and bioavailability of the administered drug. Nanoparticles, owing to their diminutive size, offer several advantages in the pharmaceutical context. They possess an increased surface area, facilitating better interaction with the drug substance, thereby improving its solubility. Moreover, the inclusion of these nanoparticles within the tablet structure serves as a sophisticated mechanism to achieve controlled and sustained release profiles of the encapsulated drug. One notable class of nanoparticles, liposomes, consists of phospholipid bilayers enclosing an aqueous core. This unique structure provides liposomes with the ability to encapsulate both hydrophilic and lipophilic drugs, making them versatile carriers within the context of extendedrelease formulations. Similarly, polymeric nanoparticles, which can be tailored for specific drug release kinetics, contribute to the controlled release profile desired in trilayer tablets. The controlled and sustained release achieved through the integration of nanotechnology is instrumental in enhancing drug delivery efficiency. This controlled release mitigates the fluctuation of drug concentrations in the bloodstream, leading to more predictable pharmacokinetics and therapeutic effects. Moreover, it minimizes the likelihood of adverse effects associated with rapid drug release. Beyond mere efficiency gains, the integration of nanotechnology into extendedrelease tri-layer tablets afford the unprecedented capability of targeted drug delivery. By leveraging the unique properties of nanoparticles, these tablets can be engineered to release drugs selectively to specific tissues or cells within the body. This targeted delivery holds immense potential for optimizing therapeutic outcomes while minimizing systemic exposure and potential side effects. 3

# **3D Printing Technology:**

The advent of 3D printing technology stands as a groundbreaking revolution in the manufacturing processes associated with tri-layer tablets. This innovative methodology introduces an unprecedented level of precision, affording meticulous control over both the geometric configuration and compositional attributes of each layer within the tablet structure. The implications of this precise control are profound, as it facilitates the production of highly customized tablets, wherein specific drug release kinetics can be tailored with exceptional accuracy. In more explicit terms, the 3D printing method involves the layer-by-layer deposition of material, allowing for a meticulous construction process that can be modulated according to the specific requirements of each

layer. This level of precision is notably advantageous in the realm of tri-layer tablets, where each layer is meticulously designed to serve a distinct function in controlling drug release kinetics. As a result, this technology empowers pharmaceutical scientists to fine-tune the tablet's architecture, ensuring that the rate and pattern of drug release align precisely with the therapeutic objectives. Beyond the intrinsic advantages of precision, 3D printing technology also opens the door to the prospect of ondemand, personalized medicine within the realm of trilayer tablet production. This potential arises from the inherent flexibility of the 3D printing process, allowing for the rapid adaptation of tablet designs to accommodate the unique needs of individual patients. By tailoring tablet designs to match specific patient requirements, this technology represents a paradigm shift towards a more patient-centric approach, enhancing the overall efficacy and adherence to therapeutic regimens.

## **Smart Polymers:**

The progress in the field of smart polymers represents a substantial contribution to the evolution of intelligent trilayer tablets. These polymers, characterized by their inherent responsiveness to environmental stimuli, constitute a pivotal element in orchestrating controlled drug release within the intricate design of tri-layer tablets. Smart polymers exhibit a remarkable ability to react to specific changes in the tablet's surroundings, particularly fluctuations in pH or temperature. When exposed to these environmental stimuli, the smart polymers undergo conformational changes, thereby triggering a precisely controlled release of the incorporated drug. This sophisticated mechanism allows for a targeted and adaptable drug release profile, aligning with the dynamic conditions presented by the patient's physiological environment. The integration of smart polymers into trilayer tablets imparts a level of adaptability and intelligence to the drug delivery system. By responding to the variations in pH or temperature, which are indicative of different physiological states within the body, the smart polymers effectively modulate the drug release kinetics. This responsiveness ensures that the tablet optimally adjusts to the unique conditions encountered in various regions of the gastrointestinal tract or other targeted anatomical sites. The adaptability of tri-layer tablets through smart polymers plays a crucial role in optimizing therapeutic outcomes. This is achieved by tailoring the drug release to specific physiological requirements, ensuring the drug is delivered precisely when and where it is needed most. Consequently, this approach enhances the overall efficacy of the treatment, as the therapeutic agent is strategically released to exert its pharmacological effects at the desired location. A notable advantage of incorporating smart polymers into tri-layer tablets is the potential to minimize side effects associated with drug administration. The controlled and targeted release facilitated by these polymers not only enhances the therapeutic efficacy but also mitigates the risk of adverse reactions. By avoiding abrupt or excessive drug release,



Available online at www.globalresearchonline.net

smart polymers contribute to a more favourable safety profile, aligning with the imperative of minimizing undesired effects in patient care.<sup>4</sup>

## **Combination Therapies:**

Extended-release tri-layer tablets have become integral in the realm of combination therapies, where the concurrent release of multiple drugs with distinct profiles is strategically orchestrated. This innovative approach serves as a therapeutic cornerstone, particularly in addressing medical conditions that necessitate complex а comprehensive treatment strategy. By facilitating the simultaneous release of multiple drugs, each tailored to address specific facets of the disease, these tri-layer tablets offer a multifaceted solution. The hallmark of this approach lies in the meticulous control over drug release achieved through the distinct layers within each tablet. Each layer is meticulously designed to house a particular drug, exhibiting unique release kinetics. This precision ensures that the drugs are released at specified intervals, allowing for a synergistic effect. The synchronized delivery of multiple therapeutic agents enhances the overall efficacy of the treatment, as each component can act on different aspects of the disease pathology. The significance of extended release tri-layer tablets in combination therapies extends beyond the therapeutic spectrum to encompass patient compliance. The precisely orchestrated release profiles contribute to a streamlined dosage regimen, simplifying the complexities associated with managing multiple medications. This not only enhances the convenience for patients but also plays a pivotal role in fostering improved adherence to the prescribed treatment plan. Moreover, the tailored control over drug release within each layer enables a harmonized therapeutic response, avoiding potential interactions that might compromise the overall effectiveness of the treatment. This aspect is particularly crucial when dealing with intricate medical conditions where drug interactions could have significant implications for patient outcomes.

# **Personalized Medicine:**

The paradigm of personalized medicine is increasingly becoming a focal point in the meticulous design of extended-release tri-layer tablets. This innovative approach involves the meticulous tailoring of formulations to align seamlessly with the unique characteristics of individual patients, including considerations of their genetics, lifestyle, and disease progression. This tailored approach represents a pivotal shift in pharmaceutical strategy, as it seeks to optimize treatment efficacy by acknowledging and accommodating the inherent diversity among patients. The incorporation of genetic information into the formulation design ensures that the extendedrelease tri-layer tablets are crafted with a deep understanding of each patient's genetic makeup. This level of precision allows for the creation of formulations that are not only therapeutically effective but also considerate of genetic factors that may influence drug metabolism and response. As a result, the approach significantly enhances

the overall therapeutic outcomes for each patient. Consideration of lifestyle factors further refines the personalized medicine approach in extended-release trilayer tablet design. Lifestyle variables such as dietary habits, physical activity levels, and other routine behaviours can profoundly impact drug absorption, metabolism, and overall treatment outcomes. By tailoring formulations to accommodate these lifestyle nuances, the model ensures a more personalized medicine comprehensive and effective therapeutic intervention tailored to individual patient needs. In tandem with genetic and lifestyle considerations, the incorporation of disease progression data becomes integral to optimizing treatment efficacy. Understanding the trajectory and specific characteristics of a patient's disease enables the formulation of extended-release tri-layer tablets that adapt to evolving medical conditions. This dynamic approach ensures that the medication remains aligned with the changing needs of the patient, further enhancing therapeutic outcomes over the course of the treatment. The overarching goal of personalized medicine in the design of extended-release tri-layer tablets is to provide a holistic and patient-centric approach to healthcare. By tailoring formulations to individual characteristics, this approach minimizes adverse effects, a crucial factor in promoting patient adherence to prescribed regimens. Improved adherence, in turn, contributes significantly to better treatment outcomes and fosters a positive impact on overall healthcare outcomes. 5

# **Data-Driven Approaches:**

The seamless integration of data-driven approaches, leveraging the power of artificial intelligence (AI) and machine learning (ML), represents a transformative leap forward in the formulation design process for extendedrelease tri-layer tablets. This contemporary methodology involves the systematic analysis of expansive datasets, enabling researchers to make informed predictions and optimize drug release profiles with a level of precision that was previously unattainable. Through the utilization of AI and ML algorithms, researchers can discern intricate patterns within the datasets, extracting valuable insights that inform the formulation design of tri-layer tablets. These advanced computational tools can navigate the complexity of various factors influencing drug release, including the interplay of different layers within the tablet, specific drug properties, and environmental conditions. The predictive capabilities of data-driven approaches play a pivotal role in expediting the development of highly effective tri-layer tablet formulations. By leveraging the insights gained from the analysis of extensive datasets, researchers can make informed decisions regarding the composition, layering, and other critical parameters of the tablet design. This informed approach significantly accelerates the entire drug development process, ensuring that formulations are not only efficient but also tailored to meet specific therapeutic requirements. Moreover, the adoption of this data-driven paradigm brings about a notable reduction in the time and costs associated with



traditional trial-and-error methods. Historically, the formulation design process relied heavily on iterative experimentation, consuming substantial time and resources. The precision afforded by data-driven approaches minimizes the need for extensive empirical testing, allowing researchers to streamline the development path, thereby realizing cost efficiencies and expediting the time to market for new tri-layer tablet formulations. <sup>6</sup>

# In Vitro-In Vivo Correlation (IVIVC):

The process involves meticulously comparing the outcomes of in vitro experiments, conducted in artificial laboratory settings, with the real-world in vivo performance within living organisms. By deciphering and quantifying this relationship, researchers gain insights into how the tablets will behave in the complex and dynamic environment of the human body. Furthermore, improved IVIVC models play a pivotal role in enhancing the reliability of assessments related to drug release kinetics, bioavailability, and the overall therapeutic efficacy of trilayer tablets. The accuracy of these predictions is paramount in guiding subsequent stages of drug development, ensuring that formulations are not only effective but also exhibit the desired release profiles within the physiological context. This refinement contributes to a more informed decision-making process during the development and optimization of tri-layer tablet formulations, ultimately translating to safer and more efficacious medications for patients.<sup>7</sup>

# CONCLUSION

The landscape of extended-release tri-layer tablets is rapidly evolving with these innovative approaches. From nanotechnology to personalized medicine, these advancements hold great promise for enhancing the precision, efficiency, and patient-centric nature of drug delivery systems. As researchers continue to push the boundaries of pharmaceutical science, these developments pave the way for a new era of advanced and tailored therapeutic interventions. **Source of Support:** The author(s) received no financial support for the research, authorship, and/or publication of this article.

**Conflict of Interest:** The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

## REFERENCES

- 1. Nizami F, Malviya Y. Recent Advancement and Challenges in Bilayer Tablet Technology: An Overview. Current Research in Pharmaceutical Sciences. 2021:6(2):91-7.
- 2. JADIYA S, UPMANYU N, JAIN V. How the advanced technologies are becoming the first choice for pharmaceutical competitors: A review on Multi-layered Tablet dosage forms. JJPBS, 2023;9(1):55-61.
- Balducci AG, Magosso E, Colombo G, Sonvico F. From tablets to pharmaceutical nanotechnologies: Innovation in drug delivery strategies for the administration of antimalarial drugs. Journal of Drug Delivery Science and Technology. 2016 Apr 1;32:167-73.
- Asadi-Zaki N, Mardani H, Roghani-Mamaqani H, Wang F. Stimuli-induced adjustment of spatial distribution of fluorescence resonance energy transfer dyads in smart polymers. Coordination Chemistry Reviews. 2024 Feb 1;500:215518.
- Du J, Gao S, Chen CH. A contextual ranking and selection method for personalized medicine. Manufacturing & Service Operations Management. 2024 Jan;26(1):167-81.
- Sen S, Ganta B, Rachel VN, Gogikar SK, Singh V, Sonti R, Dikundwar AG. Mapping Advantages and Challenges in Analytical Development for Fixed Dose Combination Products, a Review. Journal of Pharmaceutical Sciences. 2024 Apr 30;5(3):18-26.
- Bassetto R, Amadio E, Ciampanelli F, Perin S, Ilari P, Gaballo P, Callegari M, Feltrin S, Gobbo J, Zanatta S, Bertin W. Designing an effective dissolution test for bilayer tablets tailored for optimal melatonin release in sleep disorder management. Frontiers in Nutrition. 2024 May 6;11:1394330.

For any questions related to this article, please reach us at: <a href="mailto:globalresearchonline@rediffmail.com">globalresearchonline@rediffmail.com</a> New manuscripts for publication can be submitted at: <a href="mailto:submit@globalresearchonline.net">submit@globalresearchonline@rediffmail.com</a>



Available online at www.globalresearchonline.net