

Recent trends in 3D-printed pharmaceuticals: Personalized dosage forms and their clinical applications

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ABSTRACT

The integration of 3D printing technology into the pharmaceutical industry has revolutionized drug development, enabling the production of personalized dosage forms tailored to individual patient needs. This review explores recent trends in 3D-printed pharmaceuticals, highlighting advancements in drug formulation, customizable dosage forms, and multi-drug delivery systems. By enabling precise dosing and drug release profiles, 3D printing enhances therapeutic efficacy, patient compliance, and treatment outcomes, particularly in areas such as oncology, pediatrics, geriatrics, and rare diseases. Personalized medicine, supported by 3D printing, allows healthcare providers to design medications specific to a patient's genetic, environmental, and lifestyle factors, thereby addressing limitations of conventional mass-produced pharmaceuticals. The review also examines challenges in scalability, regulatory oversight, and manufacturing consistency, while discussing potential future directions involving artificial intelligence, bioprinting, and nanotechnology. As the pharmaceutical landscape evolves, 3D printing holds great promise for the future of personalized healthcare, offering flexibility, patient-centric solutions, and improved therapeutic outcomes.

Keywords: 3D printing, additive manufacturing, customized medicine, drug formulation, multi-drug delivery, patient compliance, personalized dosage forms

Introduction

The integration of 3D printing technology into the pharmaceutical industry represents a transformative shift toward personalized medicine. This technology allows for the on-demand production of tailored dosage forms that can meet individual patient needs, thus enhancing therapeutic efficacy and patient compliance. The importance of personalized medicine in modern healthcare cannot be overstated, as it aims to provide customized treatment plans based on the unique characteristics of each patient, including genetic, environmental, and lifestyle factors.

The objectives of this review are to outline the current state of 3D printing in pharmaceuticals, explore its clinical applications, and discuss the implications for personalized dosage forms. By

examining recent advancements and trends, this review will provide insights into how 3D printing is reshaping drug development and delivery.

Overview of 3D Printing Technology in the Pharmaceutical Industry

3D printing, also known as additive manufacturing, encompasses various techniques that build objects layer by layer from digital models. In pharmaceuticals, it facilitates the creation of complex drug formulations that can be tailored to specific patient requirements. Techniques such as fused deposition modeling (FDM), stereolithography (SLA), and Inkjet printing are commonly employed to fabricate personalized dosage forms that can vary in size, shape, and drug release profiles.^[1]

The flexibility of 3D printing allows for the incorporation of multiple active pharmaceutical ingredients (APIs) into a single dosage form, enabling combination therapies that can enhance treatment outcomes. For instance, the development of multi-layered tablets and polyfills

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illustrates the potential of 3D printing to streamline medication regimens for patients with chronic diseases.^[2,3]

Importance of Personalized Medicine in Modern Healthcare

Personalized medicine aims to optimize therapeutic interventions by considering individual patient differences. The rise of chronic diseases and the need for effective treatment strategies have driven the demand for personalized pharmaceuticals. 3D printing technology aligns with this paradigm by enabling the customization of drug formulations, allowing for precise dosing and tailored release mechanisms.^[4]

The ability to produce medications on demand can also address issues related to drug shortages and accessibility, particularly in remote areas where traditional manufacturing and supply chains may be limited. This capability is especially significant in the context of public health emergencies, such as the COVID-19 pandemic, which highlighted the need for rapid and flexible pharmaceutical manufacturing solutions.

Objectives and Scope of the Review

This review aims to

1. Summarize the current state of 3D printing technologies in pharmaceuticals: Highlighting the various methods and their applications in drug formulation
2. Explore the clinical applications of personalized dosage forms: Discussing how these innovations are being implemented in real-world healthcare settings to improve patient outcomes
3. Assess the future implications of 3D printing in personalized medicine: Considering regulatory challenges, market trends, and potential advancements in technology that could further enhance the role of 3D printing in pharmaceuticals.

By addressing these objectives, this review will contribute to a deeper understanding of how 3D printing is revolutionizing the pharmaceutical landscape and promoting the advancement of personalized medicine.

Fundamentals of 3D Printing in Pharmaceuticals

Overview of 3D printing technology

3D printing, also known as additive manufacturing, is a process of creating three-dimensional objects from digital files by building them up layer by layer. This technology has revolutionized the pharmaceutical industry by enabling the production of personalized dosage forms with tailored drug release profiles, shapes, and sizes.^[5]

Types of 3D Printing Techniques Used in Pharmaceuticals

Several 3D printing techniques have been explored for pharmaceutical applications, including:

1. FDM: In this technique, thermoplastic polymers are melted and extruded through a nozzle to create the desired object layer by layer^[6]
2. SLA: SLA uses a ultraviolet laser to selectively cure a photosensitive resin, layer by layer, to produce the final object^[5]
3. Selective laser sintering (SLS): SLS employs a high-power laser to fuse small particles of polymer powder into a solid structure, layer by layer
4. Inkjet printing: Inkjet printing involves the deposition of liquid droplets containing the drug and excipients onto a powder bed or substrate to create the desired object
5. Binder Jetting: In binder jetting, a liquid binding agent is selectively deposited onto a powder bed to join the material together and create the final product.^[6]

Comparison of Different 3D-Printing Techniques in Terms of Pharmaceutical Applications

Each 3D printing technique has its own advantages and limitations when applied to pharmaceutical manufacturing:^[7]

FDM is suitable for thermoplastic polymers and can produce solid dosage forms with complex geometries and controlled drug release profiles.

SLA allows for the fabrication of intricate structures with high resolution but may not be suitable for heat-sensitive drugs due to the use of UV light.

SLS can process a wide range of materials, including polymers and ceramics, and is suitable for producing porous structures for controlled drug delivery.

Inkjet printing enables the precise deposition of small drug volumes and can be used to produce multi-layered or multi-drug dosage forms.

Binder jetting is a powder-based technique that can produce porous structures and is suitable for producing immediate-release dosage forms.

The choice of 3D printing technique depends on factors such as the drug properties, desired dosage form, and the specific application requirements.

Personalized Dosage Forms: Concept and Need

Definition of personalized dosage forms

Personalized dosage forms are tailored pharmaceutical formulations designed to meet the specific therapeutic needs of individual patients. This customization can involve altering the dosage, release profile, or even the formulation itself based on a patient's unique characteristics such as age, weight, metabolic rate, and genetic profile. The goal is

to enhance treatment efficacy while minimizing adverse effects and improving patient adherence to medication regimens.^[8]

Role of Personalized Dosage in Enhancing Patient Compliance and Treatment Outcomes

Personalized dosage forms significantly enhance patient compliance by providing medications that are easier to administer and more acceptable to patients. For instance, pediatric patients often require liquid formulations or smaller doses that are not typically available in standard commercial products. By customizing medications, healthcare providers can ensure that patients receive the correct dose in a form that is easier for them to take, thereby improving adherence and overall treatment outcomes.^[9]

Moreover, personalized medicine has been shown to reduce the incidence of adverse drug reactions. Traditional mass-manufactured medications may not be suitable for all patients; studies indicate that up to 70% of patients do not achieve therapeutic efficacy from these conventional approaches. Personalized dosage forms can address this issue by providing targeted therapy that aligns with an individual's specific health needs.^[10]

Comparison with Conventional Pharmaceutical Manufacturing Methods

Conventional pharmaceutical manufacturing typically relies on mass production techniques that produce fixed-dose medications in limited strengths. This one-size-fits-all approach often fails to accommodate the diverse needs of patient populations, particularly in chronic disease management where dosing may need to be adjusted frequently due to changes in a patient's condition or concurrent therapies.^[1]

In contrast, 3D-printing technology allows for the on-demand production of personalized dosage forms. This method not only facilitates the creation of individualized doses but also enables the incorporation of multiple APIs into a single dosage form, thus simplifying complex medication regimens for patients.

Examples of Drugs That Benefit from Personalized 3D Printed Dosage Forms

Several drugs have been successfully developed as personalized 3D-printed dosage forms. Notable examples include:

Spritam

A 3D-printed formulation of levetiracetam for epilepsy, which allows for rapid disintegration in the mouth, enhancing ease of administration for patients who may have difficulty swallowing pills.

ZyCoV-D

An example from the field of vaccines where 3D printing has been explored for personalized delivery systems.

These innovations illustrate how 3D printing can create dosage forms that cater specifically to patient needs, thereby improving treatment adherence and therapeutic outcomes.^[6]

Recent Trends and Innovations in 3D Printed Pharmaceuticals

Overview of current research and advancements in 3D printed drug formulations

Recent advancements in 3D printing technology have led to significant innovations in drug formulation. Research is increasingly focused on developing customizable drug release profiles that can be tailored to individual patient requirements. This includes immediate-release formulations for acute conditions and sustained or controlled-release systems for chronic disease management.

Case studies of innovative 3D-printed drugs and dosage forms

One prominent case study is Spritam, which was approved by the Food and Drug Administration (FDA) as the first 3D-printed drug. It utilizes a unique formulation process that allows for rapid disintegration upon contact with moisture, making it particularly beneficial for patients with swallowing difficulties. This case exemplifies how 3D printing can enhance drug delivery systems by creating formulations that are both effective and patient-friendly.^[7]

Customizable drug release profiles

The ability to customize drug release profiles is one of the most significant advantages of 3D-printed pharmaceuticals. Techniques such as varying the material composition or altering the geometric design of the dosage form enable manufacturers to create products with specific release characteristics – such as immediate release for rapid onset or controlled release for prolonged effects – tailored to individual patient needs.^[5]

Exploration of multi-drug delivery systems through 3D printing

Another innovative trend is the development of multi-drug delivery systems using 3D printing technology. These systems allow multiple medications to be combined into a single dosage form, simplifying administration and improving compliance among patients who require polypharmacy, especially relevant in geriatric care or chronic disease management.

Biodegradable and Bioresorbable Materials in 3D Printed Pharmaceuticals

The use of biodegradable and bioresorbable materials in 3D-printed pharmaceuticals is gaining traction as a means to enhance sustainability while maintaining therapeutic efficacy. These materials can reduce environmental impact while also providing safe degradation within the body, making them ideal candidates for applications such as implants or controlled-release systems.

Clinical Applications of 3D Printed Dosage Forms

Key areas of clinical application: Oncology, pediatrics, geriatrics, and rare diseases

The clinical applications of personalized 3D-printed dosage forms span various fields, including oncology, pediatrics, geriatrics, and rare diseases. In oncology, customized dosing can help tailor treatment plans based on individual tumor characteristics and patient responses. In pediatrics and geriatrics, personalized formulations ensure appropriate dosing based on age-related physiological differences.

Personalized dosing for specific patient populations

Personalized dosing is particularly beneficial for specific populations such as children or elderly patients who often require adjustments based on weight or metabolic capacity. For example, pediatric patients may need liquid formulations or smaller doses that are not readily available in standard formulations.

Applications in polypharmacy management

The integration of multiple medications into a single 3D-printed pill can greatly assist in managing polypharmacy – common among elderly patients who often take several medications simultaneously. This approach simplifies medication regimens and enhances adherence by reducing pill burden.^[12]

Use in orphan drugs and rare diseases where patient-specific treatments are needed

In cases involving orphan drugs or rare diseases where patient-specific treatments are essential due to small patient populations, 3D printing offers a viable solution. The ability to produce small batches tailored to individual needs makes it feasible to develop effective therapies without the extensive costs associated with traditional manufacturing methods.

Regulatory and Manufacturing Challenges

Overview of regulatory landscape for 3D printed pharmaceuticals

The regulatory landscape surrounding 3D-printed pharmaceuticals is evolving but remains complex. Regulatory bodies such as the FDA and European Medicines Agency are working towards establishing guidelines that ensure safety and efficacy while accommodating innovative manufacturing processes like 3D printing.

Quality control and consistency in 3D-printed drug production

Quality control remains a critical challenge in ensuring consistency across batches produced through additive manufacturing techniques.

Establishing robust quality assurance protocols is essential to maintain therapeutic standards similar to those achieved through conventional manufacturing methods.^[15]

Manufacturing scalability and cost-effectiveness challenges

While 3D printing offers flexibility in production, scalability remains an issue. The cost-effectiveness of producing personalized medicines at scale compared to traditional mass production methods requires further exploration and optimization.^[17]

Intellectual property and patent-related issues

Intellectual property rights related to novel formulations developed through 3D printing pose additional challenges. As this technology evolves, clear guidelines will be necessary to protect innovations while promoting collaboration within the industry.^[15,18]

Future Directions and Potential of 3D-Printed Pharmaceuticals

Role of artificial intelligence (AI) and machine learning in optimizing formulations

The integration of AI and machine learning into pharmaceutical development processes holds promise for optimizing drug formulations tailored through 3D printing technologies. These tools can analyze vast datasets to predict optimal formulation parameters based on individual patient profiles.^[18]

Future possibilities in bioprinting and organ-on-a-chip models for drug testing

Bioprinting technologies may further revolutionize drug testing by enabling researchers to create organ-on-a-chip models that mimic human physiology more accurately than traditional methods. This advancement could lead to more effective preclinical testing phases before human trials begin.

Integration of nanotechnology with 3D-printed pharmaceuticals for advanced drug delivery

Combining nanotechnology with 3D printing could enhance drug delivery systems by improving bioavailability and targeting capabilities. This integration has potential applications across various therapeutic areas including oncology and infectious diseases.

Prospects for fully personalized treatments at the point of care

The future may see fully personalized treatments produced at point-of-care settings such as hospitals or pharmacies using portable 3D printers. This capability could drastically reduce wait times for medication while ensuring that treatments are tailored precisely to individual patient needs.^[9,10]

Conclusion

In summary, recent trends indicate that personalized medicine through 3D-printed pharmaceuticals is paving the way for significant advancements in healthcare delivery. While challenges remain regarding regulation and manufacturing practices, ongoing innovations promise enhanced treatment outcomes through tailored therapies.

The impact of personalized dosage forms on healthcare is profound; they offer solutions that address individual patient needs more effectively than traditional methods ever could. As research progresses and technologies mature, personalized medicine will likely become an integral part of standard healthcare practices worldwide.

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