Chapter 7: Innovations in pharmaceutical formulation and delivery system

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ABSTRACT

Drug formulation is an evolving discipline that has made significant advancements that will contribute to advancing medicine's future. Drug formulation innovations have taken on many facets including improved drug delivery systems, increased bioavailability, enhanced therapeutic outcomes either through optimized formulations or enhanced patient compliance, and ultimately cost savings in health care through decreases in adverse reactions or improved therapeutic results with less drug consumption. This paper highlights some of the most interesting, including advances in nanotechnology and nanomedicine, new controls in controlled delivery systems, advances in personalized medicine, biologics, and threedimensional (3D) printing. Nanoparticles and nanocarriers allow for targeted drug delivery through site specific and local interventions, while controlled delivery systems permit sustained drug release and regulate concentration thereby allowing for increased compliance and adherence to drug protocols. Personalized medicine via pharmacogenomics and pharmacodynamics or pharmacokinetics uses information about a patient's unique genes and drug action to tailor performances of drugs to an individual based on these characteristics. Additionally, 3D Printing can create and manufacture both complex drug structures and individualized dosage forms. Each of these advancements begins to provide personalized, targeted, and patient-centered medicine that has the potential to change healthcare delivery outcomes.

KEYWORDS: Nanotechnology, Sustained release, Targeted drug delivery, 3D printing, Personalized medicine, Artificial intelligence.



INTRODUCTION

The field of pharmaceutical sciences continues to rapidly change due to the ongoing pursuit of improved therapeutic outcomes and patient experiences. The process of pharmaceutical formulation development and the associated drug delivery methods are integral parts of this process as they represent the fundamental basis for converting drug molecules into therapeutic agents. This introduction highlights the overview of the exciting process of discovering advancements, barriers [1 To achieve this goal, a comprehensive knowledge of drug properties, physiological barriers, the knowledge of how to design drug formulations using components together to address these properties, is necessary [2, 3] In general, nanotechnology is an interdisciplinary field that is focused on controlling matter at the molecular and atomic level [4,5]

With these systems, drugs can be released over a longer period of time, therefore giving a more sustained and therefore controlled therapeutic effect. This controlled sustained drug release system potentially overcomes most of the shortcomings of immediate release formulations that allow patients to remain compliant with prescribed treatment regimes. It also reduces the frequency with which a drug is administered - it is known overall that the greater the frequency of administration, the greater the potential for noncompliance. In turn, this improves treatment overall and provides a better patient experience. [7-9] 3D printing and similar innovations provide incredible opportunities for personalized medicine where dosage forms are fabricated on-demand to be specific for an individual patient. Perhaps augmented even more, the artificial intelligence marketplace will significantly change formulation design for drug development timelines, by allowing researchers to rapidly optimize formulation inputs and predictive modelling [11]

XISTING SYSTEM:[12]

Rapid evolution is occurring in pharmaceutical sciences, driven by the quest for better therapeutic outcomes for patients. Much of this evolution is occurring in the area of pharmaceutical formulation development and in drug delivery systems, both of which design dosage forms that optimize drug delivery in patients. Examples of recent advances in drug delivery systems include applications of nanotechnology to deliver drugs in a targeted manner to specific sites in the body, and the use of sustained release formulations to improve dosing and reduce toxicity. Nevertheless, while new technologies and drug formulation applications provide new strategies to optimize drug delivery, issues like poor solubility and stability of drug products still exist.

RECENT ADVANCEMENTS[13]

The review of literature revealed a plethora of recent advancements in pharmaceutical formulation development and drug deliverv systems. Nanotechnology-based delivery systems, including nanoparticles, liposomes, and micelles, have gained significant traction for their ability to enhance drug solubility, stability, and bioavailability. Sustained release formulations utilizing innovative materials and formulation techniques have shown promise in providing prolonged drug release with minimal variability. Additionally, targeted drug delivery approaches, such as ligand-mediated targeting and stimuli-responsive carriers, have demonstrated remarkable specificity and efficacy in delivering therapeutics to diseased tissues while minimizing off-target effects.

PROPOSED SYSTEM[12]

In response to the challenges and opportunities identified in the existing pharmaceutical formulation development and drug delivery systems, our proposed system aims to integrate cutting-edge technologies and methodologies to address unmet clinical needs and enhance therapeutic outcomes. The proposed system encompasses several key components designed to overcome current limitations and pave the way for the development of next-generation pharmaceutical formulations and drug delivery systems. Future innovations such as 3D printing for personalized medicine and AI for rapid optimization hold promise. This research paper explores the latest trends and technologies in the field, aiming to inspire further advancements for safer, more effective therapies.

ADVANCED NANOTECHNOLOGY-BASED DELIVERY SYSTEMS:[13,14]

Table 4 Advanced Manotechnology-Dased Derivery Systems				
Nanotechnology-	Key Features	Applications	Advantages	
Based System				
Liposomes	Spherical vesicles	Anticancer drugs,	Biocompatible,	
	with lipid bilayers	antifungals,	encapsulate both	
		vaccines	hydrophilic and	
			lipophilic drugs	
Polymeric	Biodegradable	Targeted drug	Controlled release,	
Nanoparticles	polymer-based	delivery, sustained	improved drug	

Table 4 Advanced Nanotechnology-Based Delivery Systems

	systems	release	stability
Solid Lipid	Solid lipid matrix	Brain delivery,	High physical
-	with surfactant	•	U I I
Nanoparticles		oral bioavailability	stability, controlled
(SLNs)	coating	enhancement	release
Nanostructured	Improved version of	Anticancer, anti-	Enhanced loading
Lipid Carriers	SLNs with liquid	inflammatory,	capacity, reduced
(NLCs)	lipid incorporation	topical drug	drug leakage
		delivery	
Dendrimers	Branched, tree-like	Gene delivery,	Precise molecular
	synthetic polymers	imaging,	structure, high drug
		anticancer drugs	loading
Micelles	Amphiphilic block	Poorly soluble	Solubilization of
	copolymers forming	drugs, anticancer	hydrophobic drugs,
	core-shell structures	therapy	small size
Carbon Nanotubes	Cylindrical carbon	Gene therapy,	High cellular uptake,
(CNTs)	molecules with high	cancer targeting	customizable
	surface area		surfaces
Quantum Dots	Semiconductor	Bioimaging, drug	High-resolution
	nanocrystals with	tracking	imaging,
	fluorescent		simultaneous
	properties		diagnosis and
			therapy
Exosomes	Natural nanoscale	Targeted delivery,	Biocompatible,
	vesicles secreted by	cancer and	natural targeting
	cells	regenerative	ability
		medicine	
Metallic	Gold, silver, or iron	Antimicrobial,	Surface plasmon
Nanoparticles	oxide nanoparticles	diagnostic	resonance, magnetic
		imaging, cancer	targeting
		therapy	

Red blood cell membrane-camouflaged nanoparticles drug delivery system

Over time, scientists had realized that nanotechnology has the potential to significantly enhance pharmaceutical delivery techniques. Nanoparticles disguised to look like red blood cell membranes are a novel kind of medication delivery. Table 1 displays advanced delivery systems based on nanotechnology.

Enzymes, drugs, proteins, and macromolecules are just a few of the bioactive substances that have been studied and proven to be well transported.[15]

NANOTECHNOLOGY AND NANO MEDICINE:

These microscopic structures provide special benefits such improved cellular absorption, regulated release, and higher drug stability [23]. A variety of medications, such as proteins, nucleic acids, and tiny compounds, may be encapsulated in nanoparticles, which can be engineered to target certain tissues or cells [24]. Nanomedicine has enormous promise for treating a wide range of illnesses, such as infectious diseases, cancer, and cardiovascular conditions. [25]

Advancements in Drug Formulation:

Personalized medicine optimizes drug dosage and reduces side effects by utilizing technologies such as pharmacogenomics and pharmacokinetics [25]. To address particular patient demands, customized medication formulations can be developed, guaranteeing optimal therapeutic efficacy while reducing toxicity. Novel delivery methods and biologics Gene treatments and monoclonal antibodies are two examples of biologic medications that have become essential components of contemporary medicine.

Drug formulation techniques have changed throughout time to handle the particular difficulties posed by biologics, including distribution, immunogenicity, and stability. The therapeutic potential of biologic pharmaceuticals is increased by novel delivery methods such liposomes, exosomes, and polymer-based carriers, which provide better protection and tailored distribution.

- These patches provide a convenient, painless, and continuous delivery method that can improve patient compliance, especially for chronic conditions.[26]
- Implantable medication Delivery Devices: Anderson's research on implantable devices enables long-term medication delivery. Adherence to treatment programs is improved when patients get regulated dosages without the requirement for frequent administration [27].
- These systems are a good option for treating respiratory disorders and enhancing patient compliance because of the work of researchers like Lavorini.[28].
- Oral Drug Delivery Technologies: The recent advancements in oral drug delivery systems, such as orally dissolving tablets (ODTs) and nanoparticles,

support ease and convenience of administration for patients and, ultimately, patient adherence. Progress has been made with these two examples [29].

• Controlled release systems:

These are taking drug formulation to the next level with the expansion of dose forms that utilize these technologies to promote a constant drug release over an extended dosing interval. By developing a controlled release system one can address the limitations of conventional immediate release dosage forms such as issues of patient adherence to prescribed dosage regimens and the potential side effects of medications. Controlled release technologies include hydrogels, microspheres, and transdermal patches to name a few and enable the ability to modulate specific rates of drug release using the appropriate drug properties and formulation technology. [30].

- **Personalized Medicine:** Personalized medicine describes custom drug delivery systems that take into account pharmacogenomics research to match drug formulations to an individual's genetics. [31]
- Smart Drug Delivery Systems: Researchers like Farokhzad have developed smart drug delivery systems. These systems respond to physiological signals and physiological changing conditions through the release of drugs based on the specific condition within the person. Smart systems allow more precise treatment and a friendlier patient experience. [32]
- Mobile Health (mHealth) Applications: Mobile health (mHealth) applications such as medication reminders, can now be integrated into drug delivery systems to enhance patient engagement and adherence. As researchers continue to explore mHealth's potential to improve patient compliance, we can conclude that novel drug delivery systems are an important and evolving area of healthcare, including the technologies that researchers and pharmaceutical companies involved in drug development are now making accessible, in order to provide medication regimens that patients can better manage and get effective use from. [33]

Personalized Medicine: The idea of customizing drug formulations to fit an individual's genetic, physiological, and clinical profile is gaining traction. Pharmacogenomics, biomarker-driven therapy, and patient-specific dosing plans will become more common. This approach aims to optimize treatment results while reducing side effects. [36]

FUTURE SCOPE

Key findings highlight the potential of integrating advanced technologies such as artificial intelligence, customized drug delivery strategies, sustained release formulations, nanotechnology-based delivery systems, and personalized medicine. These innovations can enhance therapeutic outcomes by tailoring formulations to discrete patient outlines, optimizing drug release, and improving targeting efficiency. Regulatory collaboration will be crucial to ensure safety and efficacy. Embracing innovation and interdisciplinary approaches will drive the development of transformative therapies, ultimately improving patient care worldwide.

REFERENCES

- Bruneau BG. The developmental genetics of congenital heart disease. Nature. 451, 943-948(2008)
- Kargozar S., Mozafari M. Nanotechnology and nanomedicine: Start small, think big. Mater. Today Proc. 2018;5:15492–15500. doi: 10.1016/j.matpr.2018.04.155.
- Faisal N., Kumar K. Polymer and metal nanocomposites in biomedical applications. Biointerface Res. Appl. Chem. 2017;7:2286–2294.
- Kaushik P, Yadav R. Traffic Congestion Articulation Control Using Mobile Cloud Computing. Journal of Advances and Scholarly Researches in Allied Education (JASRAE). 2018;15(1):1439-1442. https://doi.org/10.29070/JASRAE
- Luchini C, Pea A, Scarpa A. Artificial intelligencein oncology: Current applications and futureperspectives. Br J Cancer. 126, 4–9(2022).
- Nemer M. Genetic insights into normal and abnormal heart development Cardiovasc. Pathol.17, 48–54(2008).
- Srivastava D, Olson EN. A genetic blueprintfor cardiac development. Nature. 407, 221–226(2000).
- Mu W, Jiang L, Zhang J et al. Non-invasivedecision support for NSCLC treatment using PET/CT radiomics. Nat Commun. 11, 52281(2020)
- Hua S, De Matos MBC. Biodegradable polymer nanoparticles for drug delivery to solid tumors. Frontiers in Pharmacology. 2013;4:57.
- Hamburg, M. A., & Collins, F. S. (2010). The path to personalized medicine. New England Journal of Medicine, 363(4), 301-304.
- Farokhzad OC, Langer R. Impact of nanotechnology on drug delivery. ACS Nano. 2009 Jan 27;3(1):16-20. doi: 10.1021/nn900002m. PMID: 19206243.

- Rizwanullah M, Alam M, Harshita, Mir SR, Rizvi MMA, Amin S. Polymer-lipid hybrid nanoparticles: a next-generation nanocarrier for targeted treatment of solid tumors. Curr Pharm Des. 2020 Dec;26(11):1206-1215. doi: 10.2174/1381612826666200116150426.
- Ezike tc, okpala us, onoja ul, nwike cp, ezeako ec, okpara oj, okoroafor cc, eze sc, kalu ol, odoh ec, nwadike ug, ogbodo jo, umeh bu, ossai ec, nwanguma bc. Advances in drug delivery systems, challenges and future directions. Heliyon. 2023 jun 24;9(6):e17488. Doi: 10.1016/j.heliyon.2023.e17488. Pmid: 37416680; pmcid: pmc10320272.
- Tripodo G, Trapani A, Torre ML, et al. Hyaluronic acid and its derivatives in drug delivery and imaging: recent advances and challenges. *European Journal of Pharmaceutics and Biopharmaceutics*. 2015;97(Pt B):400-416. doi:10.1016/j.ejpb.2015.01.018
- Fallacara A, Baldini E, Manfredini S, Vertuani S. Hyaluronic acid in the third millennium. *Polymers*. 2018;10(7):701. doi:10.3390/polym10070701
- Lv R, Yang P, Chen G, et al. Hexagonal boron nitride nanosheets: preparation, functionalization and applications. *Materials Chemistry Frontiers*. 2018;2(3):445-467. doi:10.1039/C7QM00568F
- Liu J, Zhang X, Yang Y, et al. Boron nitride nanomaterials for biomedical applications. *Materials Science and Engineering:* C. 2017;70(Pt 1):831-849. doi:10.1016/j.msec.2016.09.035
- Mura S, Nicolas J, Couvreur P. Stimuli-responsive nanocarriers for drug delivery. *Nature Materials*. 2013;12(11):991-1003. doi:10.1038/nmat3776
- Pouton CW. Formulation of self-emulsifying drug delivery systems. *Advanced Drug Delivery Reviews*. 1997;25(1):47-58. doi:10.1016/S0169-409X(96)00490-5
- Tang B, Cheng G, Gu JC, Xu CH. Development of solid self-emulsifying drug delivery systems: preparation techniques and dosage forms. *Drug Discovery Today*. 2008;13(13-14):606-612. doi:10.1016/j.drudis.2008.04.006
- Patra JK, Das G, Fraceto LF, et al. Nano based drug delivery systems: recent developments and future prospects. *Journal of Nanobiotechnology*. 2018;16(1):71. doi:10.1186/s12951-018-0392-8
- Zhang RX, Cai P, Zhang T, et al. Nanoparticles in medicine: therapeutic applications and developments. *Clinical Pharmacology: Advances and Applications*. 2019;11:1-16. doi:10.2147/CPAA.S178176
- Ventola CL. The nanomedicine revolution: part 1: emerging concepts. *Pharmacy and Therapeutics*. 2012;37(9):512-525.
- Prausnitz MR, Langer R. Transdermal drug delivery. Nature Biotechnology. 2008;26(11):1261-1268. doi:10.1038/nbt.1504

- Anderson JM, Shive MS. Biodegradation and biocompatibility of PLA and PLGA microspheres. Advanced Drug Delivery Reviews. 2012;64:72-82. doi:10.1016/j.addr.2012.09.004
- Lavorini F, Fontana GA, Usmani OS. New inhaler devices-the good, the bad and the ugly. Respiratory Medicine. 2014;108(5):661-669. doi:10.1016/j.rmed.2014.01.003
- Rao DA, Bajaj DR, Dua K. Recent advances in orally disintegrating tablets. Pharmaceutical Nanotechnology. 2019;7(3):187-197. doi:10.2174/2211738507666190112111115
- Siepmann J, Peppas NA. Modeling of drug release from delivery systems based on hydroxypropyl methylcellulose (HPMC). Advanced Drug Delivery Reviews. 2012;64:163-174. doi:10.1016/j.addr.2012.09.027
- Hamburg MA, Collins FS. The path to personalized medicine. New England Journal of Medicine. 2010;363(4):301-304. doi:10.1056/NEJMp1006304
- Farokhzad OC, Langer R. Impact of nanotechnology on drug delivery. ACS Nano. 2009;3(1):16-20. doi:10.1021/nn900002m
- Aldeer M, Javanmard M, Martin RP. Mobile health sensing platforms for health management and disease diagnosis. Sensors (Basel). 2020;20(21):5904. doi:10.3390/s20215904
- Goyanes A, Buanz ABM, Basit AW, Gaisford S. Fused-filament 3D printing (3DP) for fabrication of tablets. International Journal of Pharmaceutics. 2014;476(1-2):88-92. doi:10.1016/j.ijpharm.2014.09.044