

Current Insights on Lipid-Based Nanosystems 2023

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Among the different types of nanosystems that have been investigated for therapeutic use, lipid-based ones are the most explored, as they have advantages over non-lipid nanosystems, especially for improving the transport and efficacy of drugs through different routes of administration, such as ocular, cutaneous, intranasal, and intravenous [1–18].

The concept of lipid-based nanosystems is broad and includes solid lipid nanoparticles (SLN), nanostructured lipid carriers (NLC), cationic lipid nanoparticles, liposomes, exosomes, nanoemulsions, microemulsions, and self-nanoemulsifying systems. Studies have shown that these nanosystems are promising for improving the efficacy of lipophilic drugs or nucleic acids in different therapeutic applications, especially those that respond to unmet medical needs [2,4,19–26].

In the second edition of the Special Issue on lipid-based nanosystems, we notice that research in this field remains very active, as we published 16 works, including 10 research articles and 6 review articles, which were the following:

1. Aljuffali, I.A.; Anwer, M.K.; Ahmed, M.M.; Alalaiwe, A.; Aldawsari, M.F.; Fatima, F.; Jamil, S. Development of Gefitinib-Loaded Solid Lipid Nanoparticles for the Treatment of Breast Cancer: Physicochemical Evaluation, Stability, and Anticancer Activity in Breast Cancer (MCF-7) Cells. *Pharmaceuticals* **2023**, *16*, 1549. <https://doi.org/10.3390/ph16111549>.
2. Ocaña-Arakachi, K.; Martínez-Herculano, J.; Jurado, R.; Llaguno-Munive, M.; Garcia-Lopez, P. Pharmacokinetics and Anti-Tumor Efficacy of PEGylated Liposomes Co-Loaded with Cisplatin and Mifepristone. *Pharmaceuticals* **2023**, *16*, 1337. <https://doi.org/10.3390/ph16101337>.
3. Arif, S.T.; Khan, M.A.; Zaman, S.u.; Sarwar, H.S.; Raza, A.; Sarfraz, M.; Bin Jardan, Y.A.; Amin, M.U.; Sohail, M.F. Enhanced Antidepressant Activity of Nanostructured Lipid Carriers Containing Levosulpiride in Behavioral Despair Tests in Mice. *Pharmaceuticals* **2023**, *16*, 1220. <https://doi.org/10.3390/ph16091220>.
4. Tyagi, R.; Waheed, A.; Kumar, N.; Ahad, A.; Bin Jardan, Y.A.; Mujeeb, M.; Kumar, A.; Naved, T.; Madan, S. Formulation and Evaluation of Plumbagin-Loaded Niosomes for an Antidiabetic Study: Optimization and In Vitro Evaluation. *Pharmaceuticals* **2023**, *16*, 1169. <https://doi.org/10.3390/ph16081169>.
5. Ahalwat, S.; Bhatt, D.C.; Rohilla, S.; Jogpal, V.; Sharma, K.; Virmani, T.; Kumar, G.; Alhalmi, A.; Alqahtani, A.S.; Noman, O.M.; et al. Mannose-Functionalized Isoniazid-Loaded Nanostructured Lipid Carriers for Pulmonary Delivery: In Vitro



Citation: Silva, A.C.; Moreira, J.N.; Sousa Lobo, J.M. Current Insights on Lipid-Based Nanosystems 2023. *Pharmaceuticals* **2023**, *16*, 1700. <https://doi.org/10.3390/ph16121700>

Received: 20 November 2023
Accepted: 24 November 2023
Published: 8 December 2023



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- Prospects and In Vivo Therapeutic Efficacy Assessment. *Pharmaceuticals* **2023**, *16*, 1108. <https://doi.org/10.3390/ph16081108>.
6. Peczek, S.H.; Tartari, A.P.S.; Zittlau, I.C.; Diedrich, C.; Machado, C.S.; Mainardes, R.M. Enhancing Oral Bioavailability and Brain Biodistribution of Perillyl Alcohol Using Nanostructured Lipid Carriers. *Pharmaceuticals* **2023**, *16*, 1055. <https://doi.org/10.3390/ph16081055>.
 7. Satyanarayana, S.D.; Abu Lila, A.S.; Moin, A.; Moglad, E.H.; Khafagy, E.-S.; Alotaibi, H.F.; Obaidullah, A.J.; Charyulu, R.N. Ocular Delivery of Bimatoprost-Loaded Solid Lipid Nanoparticles for Effective Management of Glaucoma. *Pharmaceuticals* **2023**, *16*, 1001. <https://doi.org/10.3390/ph16071001>.
 8. Garrós, N.; Bustos-Salgados, P.; Domènech, Ò.; Rodríguez-Lagunas, M.J.; Beirampour, N.; Mohammadi-Meyabadi, R.; Mallandrich, M.; Calpena, A.C.; Colom, H. Baricitinib Lipid-Based Nanosystems as a Topical Alternative for Atopic Dermatitis Treatment. *Pharmaceuticals* **2023**, *16*, 894. <https://doi.org/10.3390/ph16060894>.
 9. Tong, Y.; Shi, W.; Zhang, Q.; Wang, J. Preparation, Characterization, and In Vivo Evaluation of Gentiopicroside-Phospholipid Complex (GTP-PC) and Its Self-Nanoemulsion Drug Delivery System (GTP-PC-SNEDDS). *Pharmaceuticals* **2023**, *16*, 99. <https://doi.org/10.3390/ph16010099>.
 10. Rincón, M.; Espinoza, L.C.; Silva-Abreu, M.; Sosa, L.; Pesantez-Narvaez, J.; Abrego, G.; Calpena, A.C.; Mallandrich, M. Quality by Design of Pranoprofen Loaded Nanostructured Lipid Carriers and Their Ex Vivo Evaluation in Different Mucosae and Ocular Tissues. *Pharmaceuticals* **2022**, *15*, 1185. <https://doi.org/10.3390/ph15101185>.
 11. Paiva, D.d.F.; Matos, A.P.d.S.; Garófalo, D.d.A.; do Nascimento, T.; Monteiro, M.S.d.S.d.B.; Santos-Oliveira, R.; Ricci-Junior, E. Use of Nanocarriers Containing Antitrypanosomal Drugs for the Treatment of Chagas Disease. *Pharmaceuticals* **2023**, *16*, 1163. <https://doi.org/10.3390/ph16081163>.
 12. Korzun, T.; Moses, A.S.; Diba, P.; Sattler, A.L.; Taratula, O.R.; Sahay, G.; Taratula, O.; Marks, D.L. From Bench to Bedside: Implications of Lipid Nanoparticle Carrier Reactogenicity for Advancing Nucleic Acid Therapeutics. *Pharmaceuticals* **2023**, *16*, 1088. <https://doi.org/10.3390/ph16081088>.
 13. Subhan, M.A.; Filipczak, N.; Torchilin, V.P. Advances with Lipid-Based Nanosystems for siRNA Delivery to Breast Cancers. *Pharmaceuticals* **2023**, *16*, 970. <https://doi.org/10.3390/ph16070970>.
 14. Gugleva, V.; Andonova, V. Recent Progress of Solid Lipid Nanoparticles and Nanostructured Lipid Carriers as Ocular Drug Delivery Platforms. *Pharmaceuticals* **2023**, *16*, 474. <https://doi.org/10.3390/ph16030474>.
 15. Richards, T.; Patel, H.; Patel, K.; Schanne, F. Endogenous Lipid Carriers—Bench-to-Bedside Roadblocks in Production and Drug Loading of Exosomes. *Pharmaceuticals* **2023**, *16*, 421. <https://doi.org/10.3390/ph16030421>.
 16. Torres, J.; Costa, I.; Peixoto, A.F.; Silva, R.; Sousa Lobo, J.M.; Silva, A.C. Intranasal Lipid Nanoparticles Containing Bioactive Compounds Obtained from Marine Sources to Manage Neurodegenerative Diseases. *Pharmaceuticals* **2023**, *16*, 311. <https://doi.org/10.3390/ph16020311>.

Among the articles published in this Special Issue on lipid-based nanosystems, the lipid nanoparticles, specifically NLC, are the most explored, which suggests the potential of these systems to reach clinic in the upcoming years. Regarding the most investigated diseases, these include cancer, brain diseases, ocular diseases, skin diseases, microbial infections, and metabolic diseases. Thereby, we hope that all these works will contribute to the advancement of these scientific fields.

We would like to thank all the authors of this Special Issue for contributing high-quality articles, as well as all the reviewers who critically evaluated these works. We also thank to the assistant editor Evelyn Du for her kind help in managing this Special Issue.

Author Contributions: Conceptualization, A.C.S.; writing—original draft preparation, A.C.S.; writing—review and editing, J.N.M. and J.M.S.L. All authors have read and agreed to the published version of the manuscript.

Funding: The Applied Molecular Biosciences Unit—UCIBIO, which is financed by national funds from Fundação para a Ciência e a Tecnologia—FCT (UIDP/04378/2020 and UIDB/04378/2020), supported this work.

Conflicts of Interest: The authors declare no conflict of interest.

References

1. Almeida, H.; Silva, A.C. Nanoparticles in Ocular Drug Delivery Systems. *Pharmaceutics* **2023**, *15*, 1675. [[CrossRef](#)] [[PubMed](#)]
2. Correia, A.C.; Monteiro, A.R.; Silva, R.; Moreira, J.N.; Sousa Lobo, J.M.; Silva, A.C. Lipid nanoparticles strategies to modify pharmacokinetics of central nervous system targeting drugs: Crossing or circumventing the blood–brain barrier (BBB) to manage neurological disorders. *Adv. Drug Deliv. Rev.* **2022**, *189*, 114485. [[CrossRef](#)] [[PubMed](#)]
3. Garcês, A.; Amaral, M.H.; Sousa Lobo, J.M.; Silva, A.C. Formulations based on solid lipid nanoparticles (SLN) and nanostructured lipid carriers (NLC) for cutaneous use: A review. *Eur. J. Pharm. Sci.* **2018**, *112*, 159–167. [[CrossRef](#)] [[PubMed](#)]
4. Kon, E.; Ad-El, N.; Hazan-Halevy, I.; Stotsky-Oterin, L.; Peer, D. Targeting cancer with mRNA–lipid nanoparticles: Key considerations and future prospects. *Nat. Rev. Clin. Oncol.* **2023**, *20*, 739–754. [[CrossRef](#)] [[PubMed](#)]
5. Gugleva, V.; Andonova, V. Recent Progress of Solid Lipid Nanoparticles and Nanostructured Lipid Carriers as Ocular Drug Delivery Platforms. *Pharmaceutics* **2023**, *16*, 474. [[CrossRef](#)] [[PubMed](#)]
6. Han, H.; Li, S.; Xu, M.; Zhong, Y.; Fan, W.; Xu, J.; Zhou, T.; Ji, J.; Ye, J.; Yao, K. Polymer-and lipid-based nanocarriers for ocular drug delivery: Current status and future perspectives. *Adv. Drug Deliv. Rev.* **2023**, *7*, 114770. [[CrossRef](#)] [[PubMed](#)]
7. Wang, X.; Liu, S.; Sun, Y.; Yu, X.; Lee, S.M.; Cheng, Q.; Wei, T.; Gong, J.; Robinson, J.; Zhang, D. Preparation of selective organ-targeting (SORT) lipid nanoparticles (LNPs) using multiple technical methods for tissue-specific mRNA delivery. *Nat. Protoc.* **2023**, *18*, 265–291. [[CrossRef](#)]
8. Mohammed, H.A.; Khan, R.A.; Singh, V.; Yusuf, M.; Akhtar, N.; Sulaiman, G.M.; Albukhaty, S.; Abdellatif, A.A.; Khan, M.; Mohammed, S.A. Solid lipid nanoparticles for targeted natural and synthetic drugs delivery in high-incidence cancers, and other diseases: Roles of preparation methods, lipid composition, transitional stability, and release profiles in nanocarriers' development. *Nanotechnol. Rev.* **2023**, *12*, 20220517. [[CrossRef](#)]
9. Czajkowska-Kośnik, A.; Szekalska, M.; Winnicka, K. Nanostructured lipid carriers: A potential use for skin drug delivery systems. *Pharmacol. Rep.* **2019**, *71*, 156–166. [[CrossRef](#)]
10. Nguyen, T.-T.-L.; Maeng, H.-J. Pharmacokinetics and Pharmacodynamics of Intranasal Solid Lipid Nanoparticles and Nanostructured Lipid Carriers for Nose-to-Brain Delivery. *Pharmaceutics* **2022**, *14*, 572. [[CrossRef](#)]
11. Abla, K.K.; Mehanna, M.M. The battle of lipid-based nanocarriers against blood-brain barrier: A critical review. *J. Drug Target.* **2023**, *31*, 832–857. [[CrossRef](#)]
12. Ilić, T.; Đoković, J.B.; Nikolić, I.; Mitrović, J.R.; Pantelić, I.; Savić, S.D.; Savić, M.M. Parenteral Lipid-Based Nanoparticles for CNS Disorders: Integrating Various Facets of Preclinical Evaluation towards More Effective Clinical Translation. *Pharmaceutics* **2023**, *15*, 443. [[CrossRef](#)]
13. Jiang, Y.; Pan, X.; Yu, T.; Wang, H. Intranasal administration nanosystems for brain-targeted drug delivery. *Nano Res.* **2023**, *12*, 1–23. [[CrossRef](#)]
14. Almawash, S. Solid lipid nanoparticles, an effective carrier for classical antifungal drugs. *Saudi Pharm. J.* **2023**, *31*, 1167–1180. [[CrossRef](#)]
15. Xu, Y.; Fourniols, T.; Labrak, Y.; Prétat, V.; Beloqui, A.; des Rieux, A. Surface modification of lipid-based nanoparticles. *ACS Nano* **2022**, *16*, 7168–7196. [[CrossRef](#)] [[PubMed](#)]
16. Akbari, J.; Saeedi, M.; Ahmadi, F.; Hashemi, S.M.H.; Babaei, A.; Yaddollahi, S.; Rostamkalaei, S.S.; Asare-Addo, K.; Nokhodchi, A. Solid lipid nanoparticles and nanostructured lipid carriers: A review of the methods of manufacture and routes of administration. *Pharm. Dev. Technol.* **2022**, *27*, 525–544. [[CrossRef](#)] [[PubMed](#)]
17. Das, B.; Nayak, A.K.; Mallick, S. Lipid-based nanocarriers for ocular drug delivery: An updated review. *J. Drug Deliv. Sci. Technol.* **2022**, *15*, 103780. [[CrossRef](#)]
18. Henostroza, M.A.B.; Tavares, G.D.; Yukuyama, M.N.; De Souza, A.; Barbosa, E.J.; Avino, V.C.; dos Santos Neto, E.; Lourenço, F.R.; Löbenberg, R.; Bou-Chacra, N.A. Antibiotic-loaded lipid-based nanocarrier: A promising strategy to overcome bacterial infection. *Int. J. Pharm.* **2022**, *621*, 121782. [[CrossRef](#)] [[PubMed](#)]
19. Silva, A.C.; Moreira, J.N.; Lobo, J.M.S. Editorial—Current Insights on Lipid-Based Nanosystems. *Pharmaceutics* **2022**, *15*, 1267. [[CrossRef](#)] [[PubMed](#)]
20. Chapa González, C.; Martínez Saráoz, J.V.; Roacho Pérez, J.A.; Olivas Armendáriz, I. Lipid nanoparticles for gene therapy in ocular diseases. *DARU J. Pharm. Sci.* **2023**, *15*, 75–82. [[CrossRef](#)]
21. Zeng, L.; Gowda, B.J.; Ahmed, M.G.; Abourehab, M.A.; Chen, Z.-S.; Zhang, C.; Li, J.; Kesharwani, P. Advancements in nanoparticle-based treatment approaches for skin cancer therapy. *Mol. Cancer* **2023**, *22*, 10. [[CrossRef](#)] [[PubMed](#)]

22. Correia, A.; Moreira, J.; Sousa Lobo, J.; Silva, A. Design of experiment (DoE) as a quality by design (QbD) tool to optimize formulations of lipid nanoparticles for nose-to-brain drug delivery. *Expert Opin. Drug Deliv.* **2023**, *31*, 1–18. [[CrossRef](#)]
23. Zong, Y.; Lin, Y.; Wei, T.; Cheng, Q. Lipid Nanoparticle (LNP) Enables mRNA Delivery for Cancer Therapy. *Adv. Mater.* **2023**, e2303261. [[CrossRef](#)] [[PubMed](#)]
24. Akanda, M.; Mithu, M.S.H.; Douroumis, D. Solid lipid nanoparticles: An effective lipid-based technology for cancer treatment. *J. Drug Deliv. Sci. Technol.* **2023**, *3*, 104709. [[CrossRef](#)]
25. Jeong, M.; Lee, Y.; Park, J.; Jung, H.; Lee, H. Lipid nanoparticles (LNPs) for in vivo RNA delivery and their breakthrough technology for future applications. *Adv. Drug Deliv. Rev.* **2023**, *7*, 114990. [[CrossRef](#)]
26. Tenchov, R.; Sasso, J.M.; Zhou, Q.A. PEGylated Lipid Nanoparticle Formulations: Immunological Safety and Efficiency Perspective. *Bioconjug. Chem.* **2023**, *34*, 941–960. [[CrossRef](#)]

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