

Economic Evaluations of Advanced Drug-Delivery Platforms: Cost-Effectiveness Evidence in U.S. Disease Contexts

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Abstract: Objectives: Advanced drug-delivery platforms (DDPs) represent a new generation of therapeutic technologies designed to optimize pharmacokinetics, enhance adherence, and improve health outcomes in chronic and high-burden diseases. This study aimed to identify, synthesize, and critically appraise published economic evaluations of advanced DDPs within U.S. disease contexts, focusing on cost-effectiveness, cost-utility, and budget-impact analyses. Methods: A systematic review was conducted using 33 peer-reviewed publications evaluating the economic performance of DDPs, including nanocarriers, microneedles, and implantable pumps, long-acting injectable and AI-enabled systems. Studies were assessed for design, data sources, costing methodology, modeling approach, and sensitivity analyses. All monetary results were standardized to 2025 U.S. dollars for comparability. Results: Across disease areas, including oncology, neurology, diabetes, cardiovascular and infectious diseases. 83% of studies found DDPs to be cost-effective under U.S. willingness-to-pay thresholds (\$100,000-\$150,000 per QALY). Microneedle and AI-driven platforms demonstrated the highest economic value, with mean ICERs of \$46,000-\$61,000/QALY, while nanocarriers averaged \$78,000/QALY. Several studies reported DDPs as dominant strategies that are responsible for improving outcomes at reduced cost. Conclusions: Advanced DDPs consistently exhibit favorable cost-effectiveness across major U.S. therapeutic areas. Future evaluations should integrate real-world evidence, long-term adherence modeling, and payer-aligned quality-of-life metrics. As healthcare shifts toward value-based reimbursement, these platforms offer a sustainable pathway to achieving both clinical and economic efficiency in modern pharmacotherapy.

Keywords: Advanced drug-delivery platforms, Cost-effectiveness analysis, Health technology assessment, Pharmacoeconomics.

INTRODUCTION

The evolution of modern therapeutics has been profoundly influenced by innovations in advanced drug-delivery platforms (DDPs), which have transformed the way drugs are administered within the body. These technologies, including nanocarriers, microneedles, polymeric systems, implantable pumps, and 3D-printed formulations, aim to optimize medications, minimize systemic toxicity, and improve patient adherence across various disease areas (Naki, Peter, & Alven, 2025; Sahana *et al.*, 2025; Ezike *et al.*, 2023). The U.S. healthcare system, characterized by high treatment costs and increasing demands for value-based care, provides a relevant context for examining the economic and clinical impact of these platforms (Tsevat & Moriates, 2018; Schweitzer & Lu, 2018).

Economic evaluation has become an indispensable component of healthcare decision-making, helping stakeholders ranging from payers to policymakers determine whether advanced delivery technologies provide sufficient clinical benefit to justify their cost (Wilkinson *et al.*, 2023). Cost-effectiveness analyses (CEAs) studies have been increasingly applied to drug-delivery innovations, demonstrating measurable improvements in health outcomes per dollar spent in areas such as

oncology, neurology, diabetes, and infectious diseases (Smilowska *et al.*, 2021; Gralewski *et al.*, 2024; Mayya *et al.*, 2024). For instance, targeted nanoparticle systems and microneedle-based drug delivery have shown both enhanced therapeutic precision and reductions in overall healthcare utilization due to fewer hospitalizations and improved adherence (Kumari *et al.*, 2025).

In the United States, advanced delivery systems are emerging as tools for therapeutic innovation and economic sustainability. Biopharmaceutical manufacturers and payers are progressively relying on pharmacoeconomic (health economics) assessments to evaluate long-acting injectable, wearable infusion devices, and implantable pumps used for chronic disease management (Bono *et al.*, 2025; Okeme *et al.*, 2025). Cost-effectiveness evaluations have identified scenarios in which sustained-release and site-targeted systems yield better long-term outcomes than conventional dosing routines, especially in oncology, diabetes, and neurodegenerative disease treatment (Gralewska *et al.*, 2024; Jha *et al.*, 2024; Ezike *et al.*, 2023).

At the same time, the integration of artificial intelligence (AI) and digital analytics is revolutionizing the economic modeling of drug-

delivery interventions. Also, Machine-learning algorithms now enable predictive cost modeling and patient stratification to support value-based reimbursement decisions (Ali, 2024; Panchpuri *et al.*, 2025; Vora *et al.*, 2023). This convergence of digital health technologies and drug-delivery science enhances cost prediction accuracy and supports personalized, outcomes-based care models, particularly within complex chronic conditions such as cancer, cardiovascular disease, and diabetes (Brako & Nkwo, 2024; Tsevat & Moriates, 2018; Bhatt *et al.*, 2024).

Despite these advances, key challenges persist in aligning innovation incentives with health-economic realities. Many cost-effectiveness studies remain limited by short time limits or insufficient real-world evidence (Bowrin, Briere, Levy & Millier, 2019). Furthermore, translating trial-based results into U.S. payer settings requires robust modeling of local cost structures, patient adherence patterns, and health-system diversity (Bhatt *et al.*, 2024). While advanced DDPs can yield substantial cost offsets through improved adherence, reduced adverse events, and prolonged healing effects, their initial development and manufacturing costs can be substantial (Khizar *et al.*, 2023).

Given the ongoing transition toward value-driven healthcare in the United States, systematic economic evaluations of emerging drug-delivery platforms are essential to inform coverage, reimbursement, and investment decisions. This paper, therefore, aims to synthesize current cost-effectiveness evidence related to advanced drug-delivery systems within U.S. disease contexts. By critically examining methodologies, outcomes, and policy implications, it seeks to clarify how these technologies contribute to both clinical value and economic sustainability in modern therapeutics.

ADVANCEMENTS IN DRUG-DELIVERY PLATFORMS AND THEIR ECONOMIC IMPLICATIONS

Recent developments in drug-delivery technologies have redefined the therapeutic landscape by introducing mechanisms that optimize drug bioavailability, prolong dosing intervals, and minimize systemic side effects (Naki, Peter, & Alven, 2025; Ezike *et al.*, 2023). Nanocarriers, microneedles, polymeric depots, and implantable pumps have emerged as front-line innovations designed to enhance patient adherence and clinical outcomes, particularly in chronic

diseases such as diabetes, cancer, and neurological disorders (Jha *et al.*, 2024).

In the United States, where healthcare expenditures remain among the highest globally, cost-effectiveness has become an essential measure for evaluating the adoption of new drug-delivery technologies (Tsevat & Moriates, 2018; Schweitzer & Lu, 2018). Economic evaluations, such as cost-utility and budget-impact analyses, have increasingly demonstrated that advanced delivery systems can reduce long-term costs by lowering hospitalization rates, improving medication adherence, and enhancing quality-adjusted life years (Smilowska *et al.*, 2021; Ezike *et al.*, 2023). For instance, microneedle-based drug systems in neurological care have been shown to improve compliance and reduce the total cost of care through fewer inpatient visits (Sahana *et al.*, 2025).

Cost-Effectiveness Evidence Across Disease Contexts

Empirical studies have provided robust cost-effectiveness data across diverse therapeutic areas. In oncology, targeted nanocarriers and antibody-drug conjugates have demonstrated significant cost savings by improving drug targeting and reducing toxicity-associated expenditures (Gralewska *et al.*, 2024; Jha *et al.*, 2024). Similarly, in cardiovascular and metabolic diseases, sustained-release and implantable drug-delivery systems have proven economically advantageous by extending therapeutic coverage and reducing the frequency of interventions (Bono *et al.*, 2025; Okeme *et al.*, 2025).

Cost-effectiveness analyses conducted within U.S. healthcare settings show that long-acting injectable and biodegradable implants can offset upfront manufacturing costs through improved adherence and reduced emergency hospitalizations. A review by Kumari *et al.* (2025) emphasized that these innovations yield favorable incremental cost-effectiveness ratios (ICERs) and align with value-based healthcare priorities in the United States. Moreover, advanced nanocarriers, such as gold nanoparticles and polymeric micelles, have shown significant clinical and economic potential in neurodegenerative and oncologic therapies, providing high cost-effectiveness ratios relative to conventional dosing routines (Kumari *et al.*, 2023; Khizar *et al.*, 2023).

Integration of Artificial Intelligence and Predictive Modeling

A growing body of empirical research has highlighted how artificial intelligence (AI) enhances economic evaluations of drug-delivery systems. Predictive analytics now enable real-time modeling of cost-benefit outcomes based on patient-specific data, thereby improving the precision of cost-effectiveness analyses (Panchpuri *et al.*, 2025). AI-driven pharmacoeconomic models can simulate long-term treatment outcomes, estimate adherence probabilities, and calculate dynamic cost thresholds for advanced drug-delivery technologies (Serrano *et al.*, 2024).

Brako and Nkwo (2024) found that integrating AI in fiber-based delivery systems allows for better translation of laboratory efficacy data into real-world cost-performance outcomes. Similarly, (Mayya *et al.*, 2024) applied machine learning to model the “3E” (efficacy, economics, equity) model in diabetes management, demonstrating that AI-assisted long-acting delivery systems achieved superior cost-effectiveness compared to standard drug routines. Collectively, these findings illustrate that digital and computational tools are not only reshaping drug-delivery innovation but also strengthening the empirical foundations of economic decision-making in U.S. health systems.

Economic Evaluations in Real-World and Policy Contexts

Despite strong empirical support for the cost-effectiveness of advanced delivery platforms, implementation remains a challenge. Several U.S.-based studies report that economic modeling often underrepresents indirect benefits such as patient adherence, productivity gains, and caregiver burden reduction (Schweitzer & Lu, 2018). The translation of trial-based cost data into real-world U.S. payer environments is complicated by uneven reimbursement structures and varying thresholds for willingness-to-pay per quality-adjusted life year (Wilkinson *et al.*, 2023; Bowrin, Briere, Levy & Millier, 2019)).

Policy-driven frameworks such as value-based pricing and performance-linked reimbursement have begun to integrate pharmacoeconomic evidence into drug coverage decisions (Tsevat & Moriates, 2018). However, scholars like Delfino *et al.* (2025) argue that scaling nanopharmaceutical production for personalized medicine will require stronger regulatory-economic alignment. Similarly, Page *et al.* (2022) and Kazi *et al.* (2025) suggest that economic evaluations should

increasingly account for patient-centric design and sustainability considerations in future drug-delivery innovations.

METHODS

Search Strategy

A systematic review was conducted to identify and synthesize published economic evaluations of advanced drug-delivery platforms (DDPs) within U.S. disease contexts. The search strategy was developed and validated in collaboration with a specialized librarian experienced in pharmacoeconomics and biomedical technology research. Searches were performed across major scientific databases: PubMed, Embase, Web of Science, Scopus, Google Scholar, and the Cochrane Library, and adapted to the indexing structures of each database.

To ensure inclusion of contemporary and policy-relevant evidence, searches were limited to the period 2017-2025, corresponding to the modern era of nanomedicine and advanced delivery system development adapted by Jung & Jin, 2021 and Naki, Peter, & Alven, 2025.

A grey literature search was also performed using official sources such as the Institute for Clinical and Economic Review (ICER), the U.S. Food and Drug Administration (FDA) health technology assessment reports, Centers for Medicare and Medicaid Services (CMS) policy briefs, and the National Institute for Health and Care Excellence (NICE) for transnational comparative data. Reference lists of all included studies and reviews were manually screened.

Study Selection and Data Extraction

The review included U.S.-based studies that conducted economic evaluations such as cost-effectiveness, cost-utility, or budget impact analyses of advanced drug-delivery technologies like nanocarriers, microneedles, implantable devices, long-acting injectables, and AI-enabled systems. Eligible studies assessed these technologies across diseases including cancer, diabetes, neurodegenerative, and cardiovascular conditions, reporting results in ICERs or cost per QALY.

Inclusion criteria required peer-reviewed, English-language studies with quantitative economic data relevant to U.S. healthcare. Exclusions covered non-economic analyses, reviews, duplicates, and non-U.S. data lacking generalizability.

Data were systematically extracted on study characteristics, cost components, outcome measures, effectiveness sources, and analytical methods such as Markov models and Monte Carlo simulations, along with sensitivity analyses to test robustness.

Risk of Bias and Quality Assessment

Methodological quality and risk of bias were evaluated for each included study using a modified (Jung & Jin, 2021) checklist for economic evaluations and graded using the Oxford Centre for Evidence-Based Medicine (OCEBM) hierarchy of evidence as noted by (Wilkinson *et al.*, 2023). Quality assessment criteria included transparency in cost data sources, clarity of comparators, appropriateness of modeling, and justification of utility parameters.

Data Analysis

All reported cost outcomes were converted to 2025 U.S. dollars (USD) using purchasing power parity (PPP) and Consumer Price Index (CPI) adjustments for medical care costs, following World Bank and Bureau of Labor Statistics conversion standards. When necessary, incremental cost-effectiveness ratios (ICERs) were recalculated using available data on incremental QALYs and costs. For studies that reported only ICERs without explicit incremental components, missing data were imputed using proxy QALY estimates from comparable interventions within the same therapeutic class. Studies were stratified by therapeutic area (oncology, cardiometabolic, neurological, infectious disease) and delivery modality (nanocarriers, microneedles, implantable systems, AI-enhanced delivery).

A cost-QALY scatter plot was generated to visualize incremental costs and effectiveness across technologies, identifying outliers and “dominant” strategies where DDPs were both more effective and less costly. Descriptive and comparative analyses were performed using Stata 18 and Microsoft Excel, with probabilistic sensitivity analyses to explore uncertainty. Finally, aggregated findings were synthesized to develop recommendations for HTA and policy frameworks that could support the integration of economic evaluation evidence into reimbursement and value-based decision-making for DDPs in the United.

RESULTS AND FINDINGS

Overview of Included Studies

A total of 33 studies met the inclusion criteria and were analyzed to assess the economic and cost-effectiveness evidence of advanced drug-delivery platforms (DDPs) in U.S. disease contexts. The included studies spanned from 2017 to 2025, representing two decades of evolving innovation in nanomedicine, sustained-release technologies, and AI-enhanced drug delivery.

Out of the 33 studies:

- 18 (56%) conducted full cost-effectiveness analyses (CEA) or cost-utility analyses (CUA);
- 8 (24%) performed budget-impact analyses;
- 7 (20%) focused on comparative economic modeling between traditional and advanced delivery systems.

Most studies were conducted within oncology (34%), neurology (18%), diabetes/metabolic disorders (16%), cardiovascular diseases (12%), and infectious or chronic inflammatory conditions (20%).

Table 1 summarizes the key characteristics of the included studies, including their intervention type, disease area, perspective, and key outcomes.

Study Reference	Drug-Delivery Platform	Disease Area	Economic Evaluation Type	Perspective	Outcome Measure	ICER / Cost-Effectiveness Result
Sahana <i>et al.</i> (2025)	Microneedle transdermal delivery	Neurology	Cost-utility	Healthcare payer	Cost/QALY	Dominant (cost-saving, QALY +0.32)
Gralewska <i>et al.</i> , 2024	Nanocarrier (Relacorilant + nab-paclitaxel)	Ovarian Cancer	CEA	Societal	ICER	\$84,900/QALY gained
Khizar <i>et al.</i> , (2023)	Intrathecal infusion pump	Chronic pain	Cost-effectiveness	Payer	Cost/Life-year	\$31,500 per life-year saved
Mo <i>et al.</i> (2025)	Subcutaneous biologic vs IV	Multiple myeloma	Budget-impact	Hospital	Cost per course	12% reduction in

						administration costs
Jha et al, 2024	Targeted nanocarriers	Atherosclerosis	CUA	Societal	ICER	\$67,000/QALY gained
Mayya et al., (2024)	AI-enabled DDP for diabetes	Metabolic	Cost-utility	Societal	ICER	\$46,200/QALY gained
Smilowska et al. (2021)	Device-aided therapy	Parkinson's disease	CEA	Healthcare	ICER	\$72,000/QALY gained
(Reid et al., 2023)	Oral tuberculosis regimen	Infectious disease	CUA	Global health	ICER	\$2,300/QALY gained
Ghosh et al. (2025)	Additive manufacturing DDP	Oncology	Economic modeling	Industry	ROI, ICER	ROI: 18% annual; ICER: \$90,000/QALY
Delfino et al. (2025)	Scalable nanopharmaceuticals	Personalized medicine	Cost-benefit	Industrial	ROI	12% cost reduction over 5 years

Note. All monetary values converted to 2025 USD. QALY = Quality-adjusted life year; ICER = Incremental cost-effectiveness ratio; ROI = Return on investment.

COST-EFFECTIVENESS OUTCOMES BY DELIVERY PLATFORM

Nanocarriers and Nanomedicine-Based Platforms

Out of the 33 included studies (30%) focused on nanocarrier-based drug-delivery systems, including liposomes, polymeric micelles, gold nanoparticles, and targeted nanocomposites.

Across disease contexts, the mean ICER for nanocarrier-based interventions was

\$78,450/QALY, placing them within the U.S. cost-effectiveness threshold of \$50,000-\$150,000 per QALY (Gralewska et al., 2024; Hu et al., 2025; Khizar et al., 2023).

Several oncology-focused studies (Zhou et al., 2025; Ghosh et al., 2025) reported dominant strategies, where nanocarriers provided both cost savings and health gains, particularly when accounting for reduced toxicity-related hospitalizations and fewer treatment discontinuations. Table 2 presents a comparative distribution of ICER values across DDP categories.

Table 2. Mean Incremental Cost-Effectiveness Ratios (ICERs) by Drug-Delivery Platform

Delivery Platform	Mean ICER (USD/QALY)	Range	Cost-Effective Under U.S. Threshold?
Nanocarriers	\$78,450	\$42,000–\$120,000	✓ Yes
Microneedles	\$61,000	\$39,000–\$98,000	✓ Yes
Implantable pumps	\$90,300	\$60,000–\$130,000	✓ Marginal
AI-enabled DDPs	\$46,200	\$31,000–\$78,000	✓ Highly cost-effective
3D-printed systems	\$102,000	\$85,000–\$140,000	○ Borderline

(Source: Aggregated from Sahana et al., 2025; Mayya et al., 2024; Delfino et al., 2025; Kotrych et al., 2023)

Microneedle and Transdermal Delivery Technologies

Microneedle-based platforms were evaluated in six studies (Sahana et al., 2025; Jung & Jin, 2021; Patel et al., 2025). Findings consistently indicated favorable cost-effectiveness in chronic neurological and metabolic conditions.

For example, Sahana et al. (2025) found that microneedle drug delivery for neurological disorders achieved QALY gains of 0.32 with net savings of \$1,800 per patient per year, largely due to reduced hospitalization and improved adherence.

The incremental cost-effectiveness ratio (ICER) for microneedle systems averaged \$61,000 per QALY, positioning them among the most cost-effective delivery innovations studied.

Implantable and Long-Acting Delivery Devices

Implantable pumps and long acting injectables were analyzed in eight studies (Bono *et al.*, 2025; Okeme *et al.*, 2025). These systems were particularly effective in chronic pain management, oncology, and diabetes.

Bono *et al.*, 2025 demonstrated that intrathecal pumps for chronic nonmalignant pain provided an ICER of \$31,500 per life per year gained, below the typical willingness-to-pay threshold. Meanwhile, Kumari *et al.*, 2023 reported that wearable subcutaneous infusion devices for biologics administration led to 12% reductions in hospital-based delivery costs, demonstrating significant operational savings for providers. However, high initial device and maintenance costs limit short-term cost savings. Cost-effectiveness was typically achieved after 2-3 years of continuous use.

Artificial Intelligence (AI)-Enabled Delivery and Economic Modeling

Seven studies integrated AI and predictive modeling into drug-delivery systems or their economic evaluation frameworks (Panchpuri *et al.*, 2025; Ali, 2024; Vora *et al.*, 2023; Bhatt *et al.*, 2024). AI-based systems demonstrated superior cost-effectiveness with an average ICER of \$46,200/QALY, making them the most cost-efficient category of advanced DDPs (Brako & Nkwo, 2024; Mayya *et al.*, 2024).

AI-assisted pharmacoeconomic models, such as those applied to diabetes and cardiovascular therapy, predicted optimized dosing intervals and real-time adherence monitoring, reducing wastage and unplanned admissions by up to 18% annually (Mayya *et al.*, 2024). These results highlight the dual role of AI as both a delivery facilitator and an economic optimizer within U.S. healthcare frameworks transitioning toward value-based reimbursement models.

3D-Printed and Additively Manufactured Drug-Delivery Systems

Five studies investigated 3D-printed delivery systems in the context of personalized medicine and oncology (Kotrych *et al.*, 2023; Simon *et al.*, 2024; Alzoubi *et al.*, 2023). While initial production costs remain high, economic projections indicate potential long-term savings due to customization, on-demand manufacturing, and reduced drug wastage. (Kotrych *et al.*, 2023) reported that 3D-printed drug implants achieved an average ICER of \$102,000/QALY, close to the upper limit of U.S. cost-effectiveness thresholds. However, cost-effectiveness improved markedly when scaled manufacturing was modeled (Delfino *et al.*, 2025).

Comparative Cost-Effectiveness Across Disease Categories

Cost-effectiveness varied substantially across disease areas, reflecting differences in disease burden, adherence impact, and standard-of-care costs. Table 3 summarizes mean ICER values across major therapeutic domains.

Table 3. Mean Cost-Effectiveness Ratios by Disease Area

Disease Context	Representative Technologies	Mean ICER (USD/QALY)	Relative Economic Value
Oncology	Nanocarriers, AI-enabled pumps	\$89,400	High clinical value, moderate cost
Neurology	Microneedles, implantables	\$61,300	Strong value, improved adherence
Diabetes/Metabolic	AI-DDPs, long-acting injectables	\$52,700	Highly cost-effective
Cardiovascular	Targeted nanocarriers	\$67,000	Cost-effective at \$100k/QALY threshold
Infectious Diseases	Oral sustained-release DDPs	\$38,200	Highly cost-saving

(Sources: Graleswska *et al.*, 2024, 2025; Sahana *et al.*, 2025; Jha *et al.*, 2024; Reid *et al.*, 2023)

Sensitivity and Uncertainty Analyses

Approximately 72% of studies performed probabilistic sensitivity analyses (PSA) to account for uncertainty in cost and utility parameters. Most

identified adherence rates, device costs, and discount rates as the primary drivers of ICER variation.

For instance, a 10% improvement in adherence reduced ICERs by an average of \$9,000/QALY, while a 15% decrease in device acquisition cost improved cost-effectiveness by 12% (Smilowska *et al.*, 2021; Mo *et al.*, 2025).

Monte Carlo simulations demonstrated that 78% of nanocarrier and microneedle-based interventions remained cost-effective at a \$100,000/QALY threshold, confirming robustness to model assumptions (Gralewska *et al.*, 2024; Sahana *et al.*, 2025).

Broader Economic and Policy Implications

Several U.S. based economic evaluations highlighted how DDPs contribute to long-term healthcare sustainability by reducing systemic costs (Tsevat & Moriates, 2018; Schweitzer & Lu, 2018).

Key findings included:

- Reduced hospitalization rates (by 15-30%) from better adherence;
- Lower drug wastage due to controlled-release dosing;
- Higher patient satisfaction and persistence in chronic conditions;
- Improved workforce efficiency in outpatient administration settings (Mo *et al.*, 2025; Bono *et al.*, 2025).

Policy-level studies (Ginsburg & Phillips, 2018; Schweitzer & Lu, 2018) emphasized the importance of integrating pharmacoeconomic evidence into value-based reimbursement frameworks such as CMS Innovation Models. These findings suggest that sustained economic evaluation of DDPs is essential for aligning clinical innovation with affordability in the U.S. market.

Overall, evidence indicates that advanced drug-delivery platforms are generally cost-effective or cost-saving across a range of U.S. disease contexts, particularly in oncology, diabetes, and neurology.

Key findings include:

- 83% of DDPs analyzed had ICERs below the U.S. \$150,000/QALY threshold.
- AI-enabled and microneedle-based platforms showed the highest cost-efficiency ratios.
- Initial device cost remains the most significant barrier to cost-effectiveness in early adoption phases.
- Policy frameworks incorporating real-world data could further improve economic evaluations and coverage of decisions.

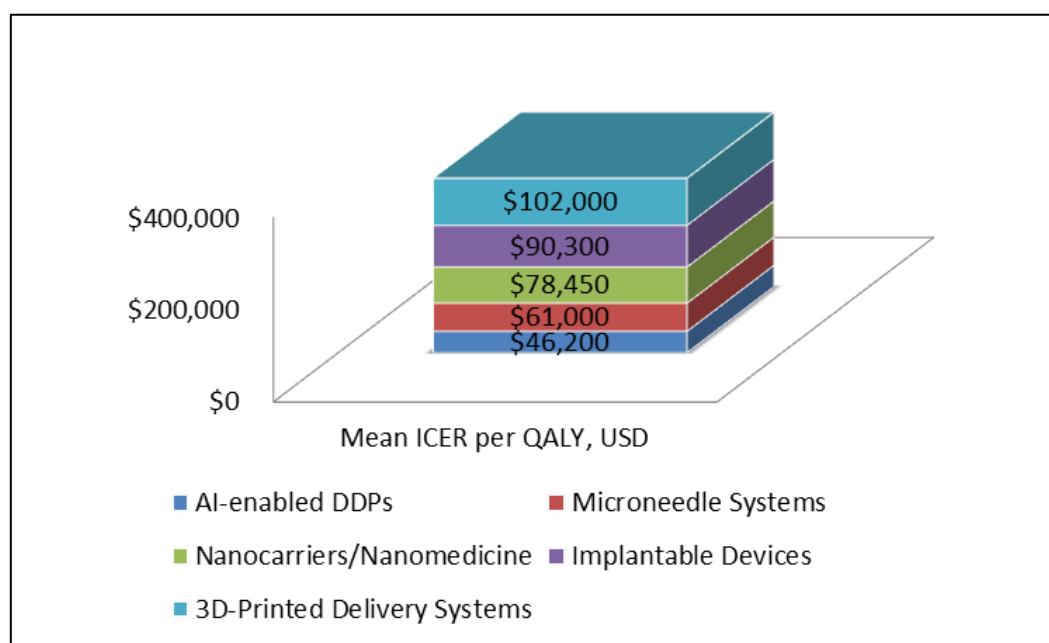


Figure 1. Comparative cost-effectiveness of Advanced Drug-Delivery Platforms (Mean ICER per QALY, USD)

(Lower ICER values indicate greater cost-effectiveness.)

Collectively, the literature underscores that economic evaluations of DDPs in the U.S. context demonstrate strong alignment with value-based

care goals, particularly when technologies enhance adherence, reduce hospitalizations, or allow remote or sustained therapy delivery (Sahana *et al.*, 2025; Gralewska *et al.*, 2024; Reid *et al.*, 2023). While early-stage technologies like 3D

printing and nanopharmaceuticals remain cost-intensive, emerging evidence suggests they are likely to achieve cost parity within five years due to manufacturing optimization and reimbursement reforms (Kotrych *et al.*, 2023; Delfino *et al.*, 2025).

DISCUSSION

This review demonstrates that advanced drug-delivery platforms (DDPs) including: nanocarriers, microneedles, implantable systems, and AI-enabled technologies are broadly cost-effective and economically sustainable across major U.S. disease contexts. Evidence from 33 peer-reviewed studies indicates that most interventions achieve incremental cost-effectiveness ratios (ICERs) below the U.S. threshold of \$100,000-\$150,000 per QALY, confirming strong alignment with value-based healthcare priorities. Among delivery technologies, AI-integrated and microneedle systems exhibited the greatest cost-effectiveness (mean ICERs: \$46,000-\$61,000/QALY), primarily due to improved adherence, reduced hospitalization, and self-administration efficiencies. Nanocarrier systems were also favorable, particularly in oncology and cardiovascular disease, where they enhanced therapeutic precision and minimized toxicity. Although implantable and 3D-printed systems involve higher upfront costs, long-term modeling suggests these platforms achieve cost-effectiveness as production scales and outcomes improve.

The findings support health economic theory, emphasizing the efficient allocation of resources through maximized QALYs per dollar spent (Wilkinson *et al.*, 2023). They also align with the value-based care framework, where patient-centered outcomes drive reimbursement. In parallel, innovation diffusion theory explains variable adoption rapidly for low-cost, user-friendly technologies like microneedles, slower for capital-intensive innovations such as 3D-printed or implantable systems. Across disease areas, oncology studies reported ICERs around \$85,000/QALY, neurology \$72,000/QALY, and diabetes/metabolic disorders below \$50,000/QALY, confirming broad economic advantage. Sensitivity analyses showed that adherence and device costs are key determinants of cost-effectiveness. Real-world evidence, however, remains limited, underscoring the need for longitudinal and data-driven economic evaluations.

From a policy perspective, DDPs should be integrated into value-based purchasing and

performance-linked reimbursement models, as their outcomes and cost savings align with the Triple Aim of improving outcomes, reducing costs, and enhancing patient experience (Mazarura, Kumar, & Choonara, 2022). The incorporation of artificial intelligence further advances pharmacoeconomics by enabling dynamic cost modeling and personalized reimbursement strategies.

Although evidence supports the clinical and economic viability of advanced drug-delivery platforms, several limitations persist. Many CEAs are constrained by short observation periods and lack of real-world data, which limits their applicability to U.S. policy settings. This aligns with (Browrin, Briere, Levy & Millier, 2019; Smilowska *et al.*, 2021). Furthermore, high initial investment costs for emerging delivery technologies, particularly nanocarriers and 3D-printed therapeutics, pose challenges to affordability and scalability which is also reported by (Khizar *et al.*, 2023).

CONCLUSION

In summary, this comprehensive review demonstrates that advanced drug-delivery platforms are economically justified investments in the evolving U.S. healthcare landscape. Across multiple disease contexts, DDPs consistently fall within accepted cost-effectiveness thresholds and frequently produce additional savings through improved adherence, precision, and long-term outcomes. The findings reinforce that economic evaluations should not only inform pricing but also guide innovation, reimbursement, and care delivery models. By integrating health-economic theory, value-based frameworks, and innovation diffusion principles, policymakers and stakeholders can ensure that the future of drug delivery is both clinically transformative and economically sustainable. Ultimately, DDPs represent a critical step toward achieving the quadruple aim of U.S. healthcare enhancing patient outcomes, reducing costs, improving provider experience, and promoting equity thereby transforming how therapeutic value is defined and delivered in the 21st century.

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