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Genetic Engineering of Microorganisms for Industrial Biocompound Production

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Abstract


The production of biocompounds using genetically engineered microorganisms has emerged as a promising and sustainable alternative to conventional extraction and chemical synthesis methods. This review provides a comprehensive overview of recent advances in genetic engineering strategies, including gene insertion, gene knockout, Clustered Regularly Interspaced Short Palindromic Repeats (CRISPR)-based genome editing, metabolic pathway engineering, and metabolic flux optimization, for enhancing biocompound production in microbial systems. The development of microbial cell factories based on bacteria, yeast, filamentous fungi, and emerging platforms has significantly improved the efficiency, scalability, and versatility of biotechnological processes. The applications of microbially derived biocompounds across various industries, including pharmaceuticals, food, agriculture, cosmetics, and chemical manufacturing, are discussed, highlighting their functional diversity and industrial relevance. Despite substantial progress, several challenges remain, such as metabolic complexity, strain stability, process scalability, and economic feasibility. The review further explores current limitations and discusses future perspectives, emphasizing the role of synthetic biology, omics technologies, and artificial intelligence in advancing microbial production systems. This study underscores the transformative potential of engineered microorganisms in the sustainable production of high-value biocompounds and provides insights into future research directions and industrial applications.

Keywords: Genetic engineering, Microbial cell factories, Biocompounds, Metabolic engineering, Clustered regularly interspaced short palindromic repeats–cas systems, Synthetic biology.

1 | Introduction

The growing demand for sustainable, efficient, and scalable production of biologically derived compounds has driven significant advancements in biotechnology and microbial engineering [1–3]. Biocompounds, chemical substances derived from living organisms such as plants, animals, and microorganisms, play a crucial role across a wide range of industrial sectors, including pharmaceuticals, food, agriculture, cosmetics, and fine

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chemicals. These compounds are valued not only for their biological activities but also for their potential as environmentally friendly alternatives to synthetic chemicals [4]. However, traditional extraction from natural sources often faces limitations, including low yields, compositional variability, environmental dependence, and high production costs. As a result, there has been a growing interest in developing alternative production strategies based on engineered microbial systems. Microorganisms, including bacteria, yeasts, and filamentous fungi, have emerged as promising platforms for the biosynthesis of high-value biocompounds [5–7]. Their rapid growth rates, well-characterized genetics, and ease of cultivation make them ideal candidates for industrial-scale production. Moreover, advances in genetic engineering and synthetic biology have enabled the precise manipulation of microbial metabolic pathways, allowing for the efficient production of complex natural and non-natural compounds [8–12]. These developments have transformed microorganisms into "cell factories" capable of producing a wide variety of bioactive molecules, including antibiotics, enzymes, vitamins, alkaloids, terpenoids, and biofuels. Genetic engineering plays a central role in optimizing microbial hosts for enhanced production of target compounds. By introducing, deleting, or modifying specific genes, researchers can redirect metabolic fluxes toward desired biosynthetic pathways while minimizing the formation of unwanted by-products. Techniques such as gene overexpression, gene knockout, and pathway reconstruction have been widely employed to improve yield and productivity [13]. In recent years, the emergence of advanced genome-editing tools, particularly the Clustered Regularly Interspaced Short Palindromic Repeats (CRISPR)/Cas systems, has revolutionized the field by enabling rapid, precise, and cost-effective genetic modifications. These tools have significantly accelerated strain development and expanded the range of achievable metabolic engineering strategies [14]. In parallel, synthetic biology has introduced a new paradigm in microbial engineering by enabling the design and construction of novel biological systems with predictable behaviors. Synthetic biology integrates principles from engineering, biology, and computational sciences to develop standardized genetic parts, regulatory circuits, and modular pathways [11–14]. This approach allows researchers to design synthetic metabolic pathways for the production of compounds that may not naturally occur in a given organism. Additionally, the use of computational modeling and systems biology tools has enhanced our understanding of cellular metabolism, enabling the identification of bottlenecks and optimization targets within complex metabolic networks. Metabolic engineering, as a complementary discipline, focuses on the systematic optimization of metabolic pathways to increase the production of desired compounds. This involves strategies such as enhancing precursor supply, balancing cofactors, eliminating competing pathways, and improving enzyme efficiency [15]. The integration of omics technologies, including genomics, transcriptomics, proteomics, and metabolomics, has provided comprehensive insights into cellular processes, facilitating data-driven approaches to strain optimization. These technologies enable the identification of key regulatory mechanisms and metabolic constraints, which can be targeted for genetic modification [10–12]. The application of genetically engineered microorganisms has already demonstrated significant success in various industrial processes. In the pharmaceutical industry, microbial systems are used to produce antibiotics, anticancer agents, and other therapeutic compounds. In the food and nutraceutical sectors, engineered microbes contribute to the production of vitamins, amino acids, and functional ingredients. Similarly, in the field of bioenergy, microorganisms are used to produce biofuels such as ethanol, biodiesel, and advanced biofuels. The chemical industry has also benefited from microbial production of organic acids, solvents, and biodegradable polymers, supporting the transition toward greener and more sustainable manufacturing processes. Despite these advancements, several challenges remain in the large-scale implementation of microbial biocompound production [14]. One of the major limitations is the metabolic burden imposed on host cells due to the overexpression of biosynthetic pathways, which can negatively affect cell growth and productivity. Additionally, the accumulation of toxic intermediates or end products may inhibit cellular function. Scale-up from laboratory to industrial production also presents significant challenges, including maintaining process stability, optimizing fermentation conditions, and ensuring economic feasibility. Addressing these challenges requires a multidisciplinary approach that combines genetic engineering, process engineering, and systems biology [16–18]. Looking ahead, the future of microbial biocompound production lies in integrating emerging technologies such as artificial intelligence, machine learning, and automation. AI-driven approaches can facilitate the prediction of optimal genetic modifications

and accelerate strain design. Automated high-throughput screening platforms and biofoundries enable rapid testing of engineered strains, significantly reducing development time. Furthermore, the increasing emphasis on sustainability and circular bioeconomy is expected to drive further innovation in the use of renewable resources and waste materials as feedstocks for microbial production [10]. Overall, genetic engineering of microorganisms represents a powerful and versatile approach for the industrial production of biocompounds. The convergence of genetic engineering, synthetic biology, and metabolic engineering has opened new opportunities for the efficient and sustainable production of high-value compounds [3–9]. This review aims to provide a comprehensive overview of the current advances in this field, highlighting key strategies, applications, challenges, and future perspectives in the development of engineered microbial systems for biocompound production. An overview of the genetic engineering strategies and microbial cell factory approaches for the production of biocompounds is illustrated in Fig. 1.

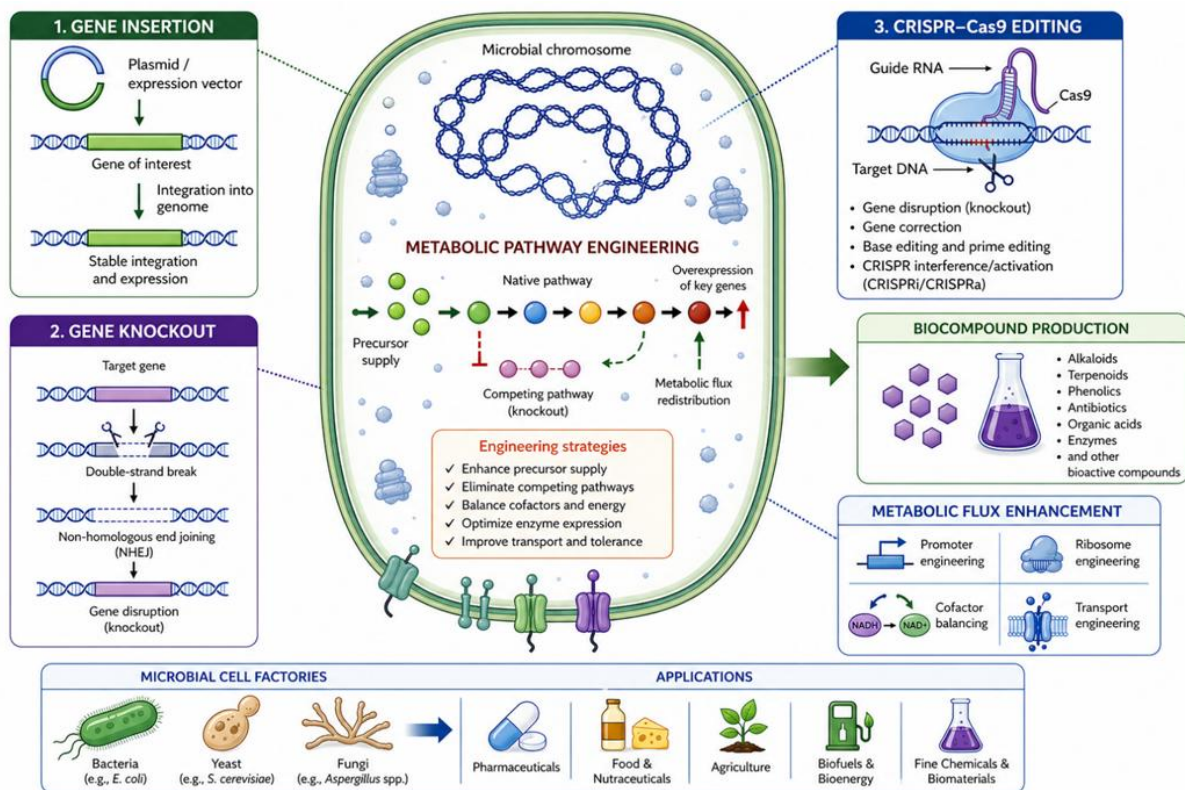


Fig. 1. The diagram also highlights strategies for enhancing metabolic flux, including cofactor balancing, promoter engineering, and pathway optimization.

2 | Genetic Engineering Strategies in Microorganisms

Genetic engineering has emerged as a cornerstone technology for the efficient production of biocompounds using microbial systems. By enabling precise manipulation of cellular genomes and metabolic networks, genetic engineering allows the development of optimized microbial cell factories capable of producing a wide range of high-value compounds. These strategies encompass gene insertion, gene deletion, genome editing, and pathway optimization approaches, all aimed at enhancing productivity, yield, and stability of target metabolites [9].

2.1 | Gene Insertion and Heterologous Expression

Gene insertion is one of the most fundamental strategies in microbial engineering, involving the introduction of foreign genes into host organisms to enable the production of desired compounds. This process is typically achieved using plasmid-based expression systems or chromosomal integration techniques. Plasmid systems offer high expression levels and flexibility, while genomic integration ensures greater genetic stability,

particularly in large-scale industrial processes [3–6]. Heterologous expression enables microorganisms such as *Escherichia coli*, *Saccharomyces cerevisiae*, and various filamentous fungi to produce compounds that are not naturally synthesized in these hosts. This strategy has been widely applied in the production of pharmaceuticals, enzymes, and secondary metabolites. However, challenges such as codon usage bias, protein folding, and metabolic burden must be addressed to achieve optimal expression levels [10].

2.2 | Gene Knockout and Pathway Elimination

Gene knockout strategies involve the targeted disruption or deletion of specific genes to eliminate competing metabolic pathways. By removing genes responsible for undesired by-product formation, metabolic flux can be redirected toward the synthesis of target biocompounds. Modern genome-editing tools have significantly improved the efficiency and precision of gene-knockout approaches. Techniques such as homologous recombination and CRISPR-based systems allow researchers to selectively inactivate genes and optimize metabolic pathways. This approach is particularly important in industrial biotechnology, where minimizing by-products and maximizing yield are critical for economic viability [4–6].

2.3 | Clustered Regularly Interspaced Short Palindromic Repeats–Cas Systems and Advanced Genome Editing

The development of CRISPR–Cas systems has revolutionized microbial genetic engineering by providing a highly efficient and programmable platform for genome editing. CRISPR–Cas9 enables precise gene disruption, insertion, and modification through targeted Deoxyribonucleic Acid (DNA) cleavage guided by Ribonucleic Acid (RNA) sequences. Beyond conventional genome editing, advanced CRISPR-based technologies, such as CRISPR interference (CRISPRi) and CRISPR activation (CRISPRa), enable reversible regulation of gene expression without altering the DNA sequence. In addition, base-editing and prime-editing technologies enable precise introduction of specific nucleotide changes, further expanding the capabilities of genetic engineering. These tools have accelerated strain development and enabled the construction of highly optimized microbial systems for the production of complex biocompounds [19].

2.4 | Metabolic Pathway Engineering

Metabolic pathway engineering focuses on modifying and optimizing biochemical pathways in microbial cells to enhance the production of target compounds. This argument includes overexpression of key enzymes, introduction of novel biosynthetic pathways, and elimination of bottlenecks that limit metabolic efficiency. One important strategy is increasing precursor availability, ensuring sufficient substrates for the biosynthesis of desired products. Additionally, removing competing pathways and optimizing enzyme kinetics can significantly improve production yields. Advances in systems biology and omics technologies have further facilitated pathway engineering by enabling a comprehensive understanding of cellular metabolism and identifying key regulatory nodes for intervention [20].

2.5 | Metabolic Flux Optimization

Metabolic flux optimization is a critical component of microbial engineering that aims to control the distribution of metabolic intermediates within the cell. By redirecting metabolic flux toward desired pathways, researchers can significantly enhance the efficiency of biocompound production. This argument can be achieved through various strategies, including promoter engineering to regulate gene expression levels, ribosome engineering to improve protein synthesis efficiency, and cofactor balancing to ensure adequate availability of essential molecules such as Nicotinamide Adenine Dinucleotide (NADH) and Adenosine Triphosphate (ATP). Transport engineering is also important for improving the uptake of substrates and secretion of products, thereby reducing intracellular accumulation and toxicity [17]. Together, these approaches enable the fine-tuning of microbial metabolism, resulting in improved productivity and scalability

for industrial applications. A summary of the main genetic engineering strategies used for biocompound production is presented in Fig. 2 and Table 1.

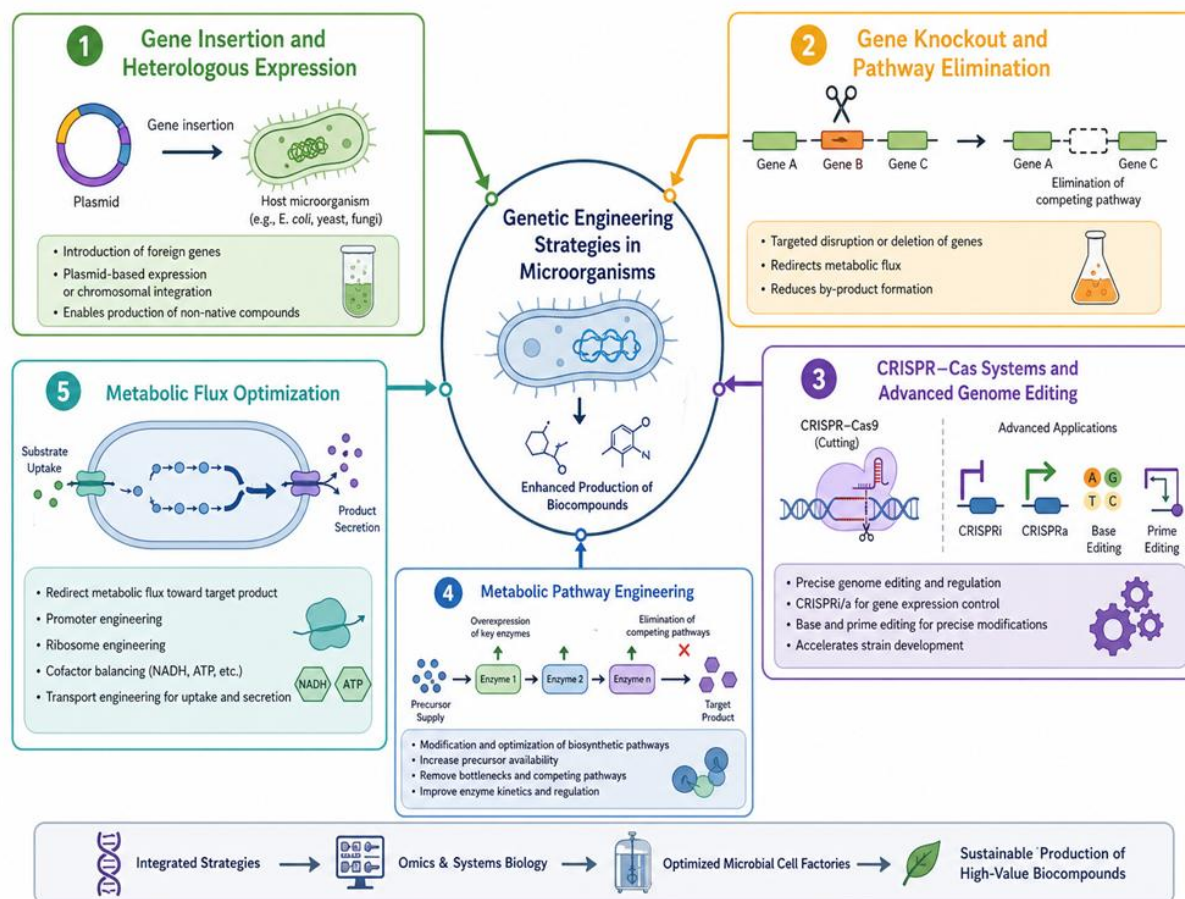


Fig. 2. Microbial genetic engineering strategies for bioproduct production.

Table 1. Genetic engineering strategies for biocompound production.

Strategy	Description	Key Techniques	Advantages	Limitations	Applications
Gene insertion	Introduction of foreign genes into host cells	Plasmids, genome integration	Enables novel product synthesis	Metabolic burden, instability	Recombinant proteins, enzymes
Gene knockout	Removal of competing genes/pathways	CRISPR–Cas, recombination	Improves yield, reduces by-products	Possible growth defects	Fermentation optimization
Crispr systems	Precise genome editing and regulation	CRISPR–Cas9, CRISPRi/a, base editing	High precision, flexible control	Off-target risks, optimization needed	Strain engineering
Pathway engineering	Modification of biosynthetic pathways	Enzyme overexpression, pathway design	Increases productivity	Complex regulation	Natural product synthesis
Flux optimization	Control of metabolic flux distribution	Promoter engineering, cofactor balance	Enhances efficiency	System complexity	Biofuels, metabolites

3 | Microbial Cell Factories for Biocompound Production

Microbial cell factories have emerged as highly efficient and versatile platforms for the sustainable production of biocompounds. By leveraging microorganisms' metabolic capabilities and integrating advanced genetic engineering strategies, it is possible to convert simple, renewable substrates into a wide array of valuable products. Microbial systems offer several advantages over traditional extraction methods, including faster growth rates, scalability, reduced environmental impact, and the ability to produce complex molecules with

high specificity and consistency. The concept of microbial cell factories is based on the design and optimization of microorganisms to function as efficient production units. This argument involves integrating synthetic pathways, optimizing native metabolic networks, and fine-tuning regulatory mechanisms to maximize product yield and minimize by-product formation. The selection of an appropriate host organism is a critical factor that significantly influences the efficiency of biocompound production [21].

3.1 | Bacterial Cell Factories

Bacteria, particularly *Escherichia coli*, are among the most widely used microbial hosts for the production of biocompounds. Their rapid growth rate, well-characterized genetic background, and availability of a wide range of molecular tools make them highly attractive for industrial applications. *E. coli* has been extensively engineered for the production of recombinant proteins, enzymes, organic acids, and various secondary metabolites. One of the key advantages of bacterial systems is their ability to achieve high cell densities and fast production cycles. Additionally, genetic manipulation in bacteria is relatively straightforward, allowing for efficient implementation of metabolic engineering strategies. However, bacterial systems also have certain limitations, including the lack of post-translational modifications required for the production of complex eukaryotic proteins and the potential accumulation of toxic intermediates. Other bacterial hosts, such as *Bacillus subtilis* and *Corynebacterium glutamicum*, are also widely used due to their ability to secrete proteins directly into the culture medium and their robustness in industrial fermentation processes. These organisms have been successfully engineered for the production of amino acids, enzymes, and bio-based chemicals [10–14].

3.2 | Yeast-Based Cell Factories

Yeasts, especially *Saccharomyces cerevisiae*, serve as important eukaryotic platforms for the production of biocompounds. As eukaryotes, yeasts possess the cellular machinery for post-translational modifications, making them suitable for the production of complex proteins and secondary metabolites. Yeast-based systems have been extensively utilized for the production of bioethanol, pharmaceuticals, and nutraceuticals. Their ability to tolerate harsh industrial conditions, such as low pH and high osmotic pressure, further enhances their applicability in large-scale production processes. Moreover, advances in synthetic biology have enabled the construction of sophisticated metabolic pathways in yeast, allowing for the production of non-native compounds such as terpenoids and alkaloids. Despite these advantages, yeast systems may exhibit slower growth rates compared to bacteria and may require more complex optimization strategies to achieve high productivity [14].

3.3 | Filamentous Fungi as Production Platforms

Filamentous fungi, including *Aspergillus niger* and *Trichoderma reesei*, are widely used in industrial biotechnology for the production of enzymes, organic acids, and secondary metabolites. These organisms are particularly known for their high secretion capacity, which simplifies downstream processing and reduces production costs. Fungal systems are highly efficient in producing extracellular enzymes such as cellulases, proteases, and amylases, which are essential for various industrial applications. In addition, filamentous fungi have complex metabolic networks that enable the biosynthesis of diverse natural products. However, the genetic manipulation of filamentous fungi is generally more challenging than that of bacteria and yeast, due to their complex cellular structure and regulatory mechanisms. Nevertheless, recent advances in genome-editing technologies have significantly increased the feasibility of engineering these organisms [20].

3.4 | Emerging Microbial Platforms

In addition to traditional hosts, emerging microbial platforms are gaining attention for their unique metabolic capabilities and potential for biocompound production. These include non-conventional yeasts, cyanobacteria, and extremophiles. Non-conventional yeasts, such as *Yarrowia lipolytica*, have shown great potential for lipid production and the synthesis of hydrophobic compounds. Cyanobacteria, as photosynthetic

microorganisms, offer the advantage of utilizing carbon dioxide and sunlight as feedstocks, making them attractive for sustainable bioproduction. Extremophiles, which thrive in harsh environments, offer opportunities to develop robust production systems capable of operating under extreme conditions. These emerging platforms expand the range of available tools for metabolic engineering and open new avenues for the production of novel biocompounds [21].

3.5 | Industrial Considerations and Scale-Up Challenges

While laboratory-scale production of biocompounds using engineered microorganisms has achieved significant success, translating these processes to industrial scale remains a major challenge. Factors such as process optimization, bioreactor design, substrate availability, and economic feasibility must be carefully considered. One of the key challenges is maintaining the stability and productivity of engineered strains under industrial conditions. Additionally, issues such as oxygen transfer, nutrient limitations, and accumulation of inhibitory compounds can affect process performance. Strategies such as adaptive laboratory evolution, process control optimization, and fed-batch or continuous fermentation are commonly employed to address these challenges. Economic considerations, including the cost of substrates, downstream processing, and overall process efficiency, play a critical role in determining the feasibility of large-scale production. The integration of bioprocess engineering with genetic and metabolic engineering is essential for achieving commercially viable production systems [18]. A comparative overview of commonly used microbial cell factories, along with their characteristics and applications, is presented in *Table 2*.

Table 2. Microbial cell factories for biocompound production.

Microorganism	Type	Key Features	Advantages	Limitations	Representative Products
<i>Escherichia coli</i>	Bacterium	Well-characterized genetics, fast growth	High expression, easy genetic manipulation	No post-translational modification, toxicity issues	Recombinant proteins, enzymes, and organic acids
<i>Bacillus subtilis</i>	Bacterium	Strong secretion system	Direct protein secretion, GRAS status	Lower expression for some proteins	Enzymes, industrial proteins
<i>Corynebacterium glutamicum</i>	Bacterium	Robust metabolism	Efficient amino acid production	Limited pathway diversity	Amino acids (lysine, glutamate)
<i>Saccharomyces cerevisiae</i>	Yeast	Eukaryotic system, well-studied	Post-translational modification, stress tolerance	Slower growth than bacteria	Bioethanol, pharmaceuticals, terpenoids
<i>Yarrowia lipolytica</i>	Yeast	Lipid metabolism	High lipid accumulation	Requires optimization	Lipids, biofuels
<i>Aspergillus niger</i>	Filamentous fungus	High secretion capacity	Efficient enzyme production	Complex genetics	Organic acids, enzymes
<i>Trichoderma reesei</i>	Filamentous fungus	Cellulase producer	High enzyme yield	Slow growth	Cellulases, bio-based enzymes
Cyanobacteria	Photosynthetic bacteria	CO ₂ utilization	Sustainable production	Low productivity	Biofuels, biochemicals
Extremophiles	Various	Tolerance to extreme conditions	Robust industrial performance	Limited genetic tools	Specialty enzymes

4 | Applications of Biocompounds in Industry

Biocompounds derived from genetically engineered microorganisms have gained significant attention due to their wide range of applications across multiple industrial sectors. Their natural origin, structural diversity, and functional properties make them highly valuable alternatives to synthetic chemicals. In recent years, advances in metabolic engineering, synthetic biology, and bioprocess optimization have enabled the large-scale production of these compounds, facilitating their integration into pharmaceutical, food, agricultural, cosmetic, and chemical industries. The growing demand for sustainable and environmentally friendly products has further accelerated the adoption of biocompounds in industrial applications [17]. Microbial production platforms offer a reliable, scalable approach to producing high-value compounds with consistent quality, reduced environmental impact, and improved economic feasibility.

4.1 | Pharmaceutical and Medical Applications

One of the most important applications of biocompounds is in the pharmaceutical and medical fields. Many naturally derived compounds exhibit potent biological activities, including antimicrobial, anticancer, anti-inflammatory, and antioxidant effects. These properties make them ideal candidates for drug discovery and development. Genetically engineered microorganisms are widely used to produce antibiotics, therapeutic proteins, vaccines, and bioactive secondary metabolites. Advances in synthetic biology have enabled the design of novel drug molecules and improved the efficiency of existing production pathways. Additionally, microbial systems are increasingly used for the biosynthesis of complex molecules that are difficult to obtain through chemical synthesis or natural extraction. Biocompounds also play a critical role in drug delivery systems, particularly in the development of nano-based carriers and targeted delivery mechanisms that enhance drug efficacy and reduce side effects [20].

4.2 | Food and Nutraceutical Applications

In the food industry, biocompounds are widely used as functional ingredients, natural additives, and nutraceuticals. These include vitamins, amino acids, organic acids, flavors, pigments, and antioxidants that improve the nutritional value, safety, and shelf life of food products. Microbial fermentation is a key process for producing food-grade biocompounds, offering advantages such as scalability, cost-effectiveness, and controlled production conditions. Engineered microorganisms have been successfully utilized to produce high-value nutraceuticals, including omega-3 fatty acids, probiotics, and bioactive peptides. The increasing consumer preference for natural, health-promoting products has driven demand for bio-based ingredients, further underscoring the importance of microbial biocompound production in the food sector [21].

4.3 | Agricultural Applications

Biocompounds play a crucial role in sustainable agriculture by providing environmentally friendly alternatives to chemical fertilizers and pesticides. Microbially derived compounds such as biopesticides, biofertilizers, and plant growth regulators contribute to improved crop productivity and soil health. Genetically engineered microorganisms can be used to produce bioactive compounds that enhance plant resistance to pests, diseases, and environmental stress. These applications not only reduce the reliance on synthetic agrochemicals but also support the development of sustainable agricultural practices. Additionally, biocompounds are increasingly used in seed treatments and soil conditioning, promoting plant growth and improving nutrient uptake efficiency [5].

4.4 | Cosmetic and Personal Care Applications

The cosmetics industry has increasingly adopted biocompounds due to their natural origin and beneficial biological properties. These compounds are widely used in skincare, haircare, and personal care products for their antioxidant, anti-aging, moisturizing, and antimicrobial effects. Microbial production provides a sustainable, consistent source of bioactive ingredients, including hyaluronic acid, peptides, enzymes, and natural pigments. The use of engineered microorganisms enables the production of high-purity compounds with controlled quality, meeting the strict regulatory requirements of the cosmetic industry. Furthermore, the shift toward green, sustainable cosmetics has significantly increased demand for bio-based ingredients [8].

4.5 | Chemical and Industrial Applications

Biocompounds are also extensively used in the chemical and industrial sectors as sustainable alternatives to petroleum-based products. These include biofuels, bioplastics, organic acids, solvents, and specialty chemicals. Microbial fermentation processes enable the production of these compounds from renewable resources, reducing dependence on fossil fuels and minimizing environmental impact. Advances in metabolic engineering have further improved the efficiency and scalability of these processes. In addition, biocompounds are used in various industrial applications such as coatings, adhesives, and biodegradable

materials, contributing to the development of a circular bioeconomy [13]. A summary of major industrial applications of biocompounds and their representative products is presented in *Table 3*.

Table 3. Industrial applications of biocompounds and their representative products.

Industry	Biocompounds	Function/Application	Examples
Pharmaceutical and medical	Antibiotics, alkaloids, peptides	Drug development, therapeutics	Penicillin, anticancer agents
Food and nutraceuticals	Vitamins, amino acids, antioxidants	Functional foods, additives	Vitamin b12, omega-3 fatty acids
Agriculture	Biopesticides, biofertilizers	Crop protection, growth enhancement	Bacillus-based biopesticides
Cosmetics	Hyaluronic acid, enzymes, pigments	Skincare, anti-aging, preservation	Natural pigments, bioactive peptides
Chemical and industrial	Biofuels, organic acids, bioplastics	Sustainable materials, energy	Ethanol, lactic acid, biopolymers

5 | Challenges and Future Perspectives

The rapid development of genetic engineering and microbial biotechnology has significantly advanced the production of biocompounds; however, several scientific, technical, and economic challenges still limit their full industrial implementation. One of the primary challenges lies in the complexity of microbial metabolic networks. Engineering microorganisms for high-yield production often requires extensive modification of interconnected pathways, which can lead to unintended metabolic imbalances, accumulation of toxic intermediates, and reduced cellular fitness. Achieving an optimal balance between cell growth and product formation remains a critical bottleneck in the design of efficient microbial cell factories. Another major challenge is the stability of engineered strains in long-term, large-scale industrial processes. Genetic modifications introduced in laboratory conditions may not remain stable under industrial fermentation environments, where fluctuations in temperature, pH, oxygen levels, and nutrient availability can negatively affect productivity. Additionally, the metabolic burden imposed by heterologous gene expression can reduce cell viability and limit overall process efficiency. Strategies such as genome integration, adaptive laboratory evolution, and dynamic regulation of gene expression are increasingly being explored to address these issues [12]. The scalability of biocompound production is also a significant concern. While many engineered systems demonstrate high performance at laboratory scale, translating these results to pilot and industrial-scale bioreactors introduces new challenges. Limitations in mass transfer, oxygen diffusion, mixing efficiency, and heat dissipation can affect microbial activity and reduce yields. Furthermore, the accumulation of inhibitory by-products and substrate limitations can complicate large-scale operations. The integration of advanced bioprocess engineering techniques, including fed-batch and continuous fermentation systems, is essential to overcome these limitations. Economic feasibility remains a key factor influencing the commercialization of microbial biocompound production [14–16]. The cost of raw materials, fermentation processes, and downstream purification can be substantial, particularly for complex molecules. Although renewable feedstocks and waste-derived substrates offer promising alternatives, their efficient utilization requires further optimization. Downstream processing, including the separation and purification of target compounds, often accounts for a major portion of production costs and remains an area requiring significant innovation [1–3]. From a regulatory perspective, the use of genetically engineered microorganisms in industrial production raises concerns about biosafety, environmental impacts, and public acceptance. Strict regulatory frameworks govern the approval and use of Genetically Modified Organisms (GMOs), particularly in food, pharmaceutical, and environmental applications. Ensuring compliance with international standards while maintaining cost-effectiveness is an ongoing challenge for researchers and industry stakeholders. Despite these challenges, the future of biocompound production using engineered microorganisms is highly promising. Advances in synthetic biology, systems biology, and computational modeling are providing new tools for the rational design of microbial systems. The integration of omics technologies, including genomics,

transcriptomics, proteomics, and metabolomics, enables a comprehensive understanding of cellular processes and facilitates the identification of novel engineering targets [7]. Machine learning and artificial intelligence are also emerging as powerful approaches for predicting metabolic behavior, optimizing pathways, and accelerating strain development. Moreover, the development of next-generation genome editing tools, such as CRISPR-based systems, continues to enhance the precision and efficiency of genetic modifications. These technologies enable multiplex genome editing and dynamic control of gene expression, opening new possibilities for the production of complex and high-value biocompounds. Emerging microbial platforms, including non-conventional hosts and synthetic consortia, further expand the potential of microbial biotechnology. Sustainability considerations are also shaping the future direction of this field. The use of renewable resources, carbon capture technologies, and environmentally friendly production processes aligns with global efforts toward a circular bioeconomy. Photosynthetic microorganisms and engineered systems capable of utilizing carbon dioxide as a feedstock represent promising avenues for sustainable biocompound production. In conclusion, while significant challenges remain in the development and industrialization of microbial cell factories, ongoing advancements in genetic engineering, systems biology, and bioprocess optimization are rapidly addressing these limitations. Continued interdisciplinary collaboration and technological innovation will be essential to unlock the full potential of microbial systems and to establish biocompound production as a cornerstone of sustainable industrial biotechnology [16].

6 | Conclusion

In this review, the role of genetic engineering in the development of microbial cell factories for the production of biocompounds has been comprehensively discussed. The integration of advanced genetic tools, including CRISPR-based technologies, synthetic biology, and metabolic engineering, has significantly enhanced the ability to design and optimize microorganisms for the efficient and sustainable production of a wide range of high-value compounds [16]. These advancements have enabled the transition from traditional extraction methods to scalable and controlled biotechnological processes. Microbial cell factories, including bacteria, yeast, filamentous fungi, and emerging microbial platforms, offer versatile and robust systems for biocompound production. Their adaptability, combined with continuous improvements in pathway engineering and metabolic regulation, has expanded their applications across pharmaceutical, food, agricultural, cosmetic, and industrial sectors. The growing demand for sustainable, eco-friendly production further underscores the importance of these systems in modern biotechnology. Despite significant progress, several challenges remain, including metabolic complexity, strain stability, process scalability, and economic feasibility. Addressing these limitations requires a multidisciplinary approach that integrates advances in systems biology, computational modeling, and bioprocess engineering. In particular, the development of more efficient downstream processing techniques and cost-effective production strategies will be critical for large-scale commercialization [17–20]. Looking forward, the continued evolution of genome editing technologies, omics-driven approaches, and artificial intelligence is expected to further accelerate innovation in this field. The emergence of novel microbial platforms and the increasing focus on sustainable production strategies, such as carbon-neutral and waste-based bioprocesses, will play a key role in shaping the future of biocompound production. Overall, the combination of genetic engineering and microbial biotechnology holds immense potential to revolutionize the production of biocompounds, advancing sustainable industries and the global bioeconomy.

Authors' Contributions

The author carried out all aspects of the research and manuscript preparation. The author has read and approved the final version of the manuscript.

Data Availability

All data are included in the text.

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Conflict of Interest

The author declares that he has no conflicts of interest.

Consent for Publication

The author has given consent for the publication of this manuscript.

Ethics Approval and Consent to Participate

This study does not involve any research conducted on human participants or animals.

References

- [1] Bution, M. L., Molina, G., Abrahao, M. R. E., & Pastore, G. M. (2015). Genetic and metabolic engineering of microorganisms for the development of new flavor compounds from terpenic substrates. *Critical reviews in biotechnology*, 35(3), 313–325. <https://doi.org/10.3109/07388551.2013.855161>
- [2] Singh, H. (2024). Bioengineering for production of biologically active compounds in plants. In *Biosynthesis of natural products in plants: Bioengineering in post-genomics era* (pp. 1-37). Singapore: Springer Nature Singapore. https://doi.org/10.1007/978-981-97-2166-5_1
- [3] Kothari, R., Singh, H. M., Azam, R., Gorla, K., Bharti, A., Singh, A., & Tyagi, V. V. (2023). Potential avenue of genetic engineered algal derived bioactive compounds: Influencing parameters, challenges and future prospects. *Phytochemistry reviews*, 22(4), 935–968. <https://doi.org/10.1007/s11101-023-09859-y>
- [4] Lin, H., Wang, Q., Shen, Q., Zhan, J., & Zhao, Y. (2013). Genetic engineering of microorganisms for biodiesel production. *Bioengineered*, 4(5), 292–304. <https://doi.org/10.4161/bioe.23114>
- [5] Sibirny, A., Madzak, C., & Fickers, P. (2018). Genetic engineering of nonconventional yeasts for the production of valuable compounds. In *Microbial biotechnology* (pp. 78–127). CRC Press. <https://doi.org/10.1201/b17587-5>
- [6] Virklund, A., Jensen, S. I., Nielsen, A. T., & Woodley, J. M. (2023). Combining genetic engineering and bioprocess concepts for improved phenylpropanoid production. *Biotechnology and bioengineering*, 120(3), 613–628. <https://doi.org/10.1002/bit.28292>
- [7] Adegboye, M. F., Ojuederie, O. B., Talia, P. M., & Babalola, O. O. (2021). Bioprospecting of microbial strains for biofuel production: Metabolic engineering, applications, and challenges. *Biotechnology for biofuels*, 14(1), 5. <https://doi.org/10.1186/s13068-020-01853-2>
- [8] Araque Gelves, D. I., de Souza, G. C., Salvador, M. J., Diaz Rodriguez, A. M., Campos Avelar, I., & de los Santos Villalobos, S. (2026). Impact of plant-microorganism interaction: A key driver for the production of bioactive metabolites of interest in the pharmaceutical, agricultural, cosmetic, and food industries. *Journal of plant growth regulation*, 45(2), 833–860. <https://doi.org/10.1007/s00344-025-11921-2>
- [9] Plamada, D., Nemes, A. S., Teleky, B. E., Pascuta, M. S., Odocheanu, R., Mitrea, L., ... & Vodnar, D. C. (2025). Microbial production of aromatic phenolic compounds. In *Microbial production of food bioactive compounds* (pp. 785-808). Cham: Springer Nature Switzerland. https://doi.org/10.1007/978-3-031-90974-0_53
- [10] Sekurova, O. N., Schneider, O., & Zotchev, S. B. (2019). Novel bioactive natural products from bacteria via bioprospecting, genome mining and metabolic engineering. *Microbial biotechnology*, 12(5), 828–844. <https://doi.org/10.1111/1751-7915.13398>
- [11] Hendrawati, O., Woerdenbag, H. J., Hille, J., & Kayser, O. (2012). Metabolic engineering of medicinal plants and microorganisms for the production of natural products. In *Pharmaceutical biotechnology: Drug discovery and clinical applications* (pp. 491-526). Wiley. <https://doi.org/10.1002/9783527632909.ch19>

- [12] Baniasad, A., Baei, M. S., & Tala Tapeh, S. M. (2025). Chitosan-PEGylated niosomes and liposomes as biomacromolecule carriers for Alzheimer's disease treatment: Galantamine drug delivery carrier. *Materials chemistry and physics*, 352, 132003. <https://doi.org/10.1016/j.matchemphys.2025.132003>
- [13] Sadanov, A. K., Baimakhanova, B. B., Orasymbet, S. E., Ratnikova, I. A., Turlybaeva, Z. Z., & Baimakhanova, G. B. (2025). Engineering useful microbial species for pharmaceutical applications. *Microorganisms*, 13(3), 599. <https://doi.org/10.3390/microorganisms13030599>
- [14] Singh, A., & Negi, P. S. (2025). Biotechnological application of health-promising bioactive compounds. In *Biotechnological intervention in production of bioactive compounds: Biosynthesis, characterization and applications* (pp. 73-94). Cham: Springer Nature Switzerland. https://doi.org/10.1007/978-3-031-76859-0_5
- [15] Choi, K. R., & Lee, S. Y. (2023). Systems metabolic engineering of microorganisms for food and cosmetics production. *Nature reviews bioengineering*, 1(11), 832–857. <https://doi.org/10.1038/s44222-023-00076-y>
- [16] Singh, B., Kumar, A., Saini, A. K., Saini, R. V., Thakur, R., & Mohammed, S. A. (2023). Strengthening microbial cell factories for efficient production of bioactive molecules. *Biotechnology and genetic engineering reviews*, 39(2), 1345–1378. <https://doi.org/10.1080/02648725.2023.2177039>
- [17] Olaniyan, O. T., & Adetunji, C. O. (2021). Biological, biochemical, and biodiversity of biomolecules from marine-based beneficial microorganisms: Industrial perspective. In *Microbial rejuvenation of polluted environment: Volume 3* (pp. 57-81). Singapore: Springer Singapore. https://doi.org/10.1007/978-981-15-7459-7_4
- [18] Liu, Y., Zhang, D., Yang, T., Chen, R., & Luo, X. (2024). Pathway engineering of plant-derived bioactive compounds in microbes. In *Engineering biology for microbial biosynthesis of plant-derived bioactive compounds* (pp. 73-87). Academic Press. <https://doi.org/10.1016/B978-0-443-15558-1.00009-6>
- [19] Dixit, S., Shukla, A., Singh, V., & Upadhyay, S. K. (2021). Bioprospecting of natural compounds for industrial and medical applications: Current scenario and bottleneck. In *Bioprospecting of plant biodiversity for industrial molecules* (pp. 53-71). Wiley. <https://doi.org/10.1002/9781119718017.ch3>
- [20] Tapeh, S. M. T., Baei, M. S., & Keshel, S. H. (2021). Synthesis of thermogel modified with biomaterials as carrier for hUSSCs differentiation into cardiac cells: Physicomechanical and biological assessment. *Materials science and engineering: C*, 119, 111517. <https://doi.org/10.1016/j.msec.2020.111517>
- [21] Kowalczyk, T., Merecz Sadowska, A., Picot, L., Brčić Karačonji, I., Wieczfinska, J., Śliwiński, T., & Sitarek, P. (2022). Genetic manipulation and bioreactor culture of plants as a tool for industry and its applications. *Molecules*, 27(3), 795. <https://doi.org/10.3390/molecules27030795>