

AgriCos e-Newsletter

Open Access Multidisciplinary Monthly Online Magazine

Volume: 06 Issue: 01 January 2025

Article No: 03

RNA Interference in Pest Management: A Transformative Approach

Chethankumar M.¹, Sneha Sharma¹, Archna Anokhe², G. P. Darshan³ and Uday Kumar B. V.⁴

¹IARI-Raipur hub, ICAR-National Institute of Biotic Stress Management, Raipur

²ICAR- Directorate of Weed Research, Jabalpur. Madhyapradesh

³Department of Genetics and Plant Breeding, Institute of Agricultural Sciences, Banaras Hindu University,

Varanasi

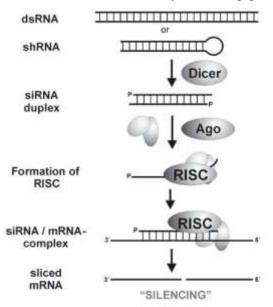
⁴Department of Biotechnology, University of Agricultural Sciences Bangalore

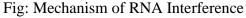
SUMMARY

Pest management has traditionally depended on chemical pesticides, which, while effective, pose significant environmental and health risks and accelerate pesticide resistance in target species. RNA interference (RNAi), a sequence-specific gene-silencing mechanism, has emerged as a highly precise and sustainable alternative. RNAi exploits endogenous cellular pathways to degrade messenger RNA (mRNA), suppressing the expression of vital genes necessary for pest survival, reproduction, or metabolism. This targeted approach minimizes off-target impacts on non-target organisms, addressing one of the key shortcomings of chemical pesticides. Advances in delivery systems, including nanoparticle carriers, plant-mediated RNAi, and sprayable formulations, have significantly enhanced RNAi's field applicability. RNAi-based biopesticides offer reduced environmental toxicity, aligning with the goals of sustainable agriculture. However, challenges such as RNA stability in field conditions, delivery efficiency, and potential resistance evolution require targeted solutions. This article discusses the mechanisms, recent advancements, and limitations of RNAi technology, underscoring its role as an innovative, eco-friendly pest management strategy.

INTRODUCTION

Global food security is continuously threatened by pest infestations that causes 20-40% crop losses worldwide leading to economic losses worth billions of dollars (Savary et al. 2019). While chemical pesticides have been the cornerstone of pest management, their indiscriminate use has led to environmental degradation, pesticide residues in food, and the emergence of resistant pest populations. Furthermore, excessive reliance on these chemicals has disrupted ecological balance by affecting beneficial organisms, pollinators, and natural predators of pests. The search for sustainable, targeted, and environmentally safe alternatives has directed scientific attention toward RNA interference (RNAi), a naturally occurring gene-silencing mechanism.





Discovered in the late 1990s, RNAi has revolutionized molecular biology and now presents significant opportunities for agriculture, particularly in pest management (Hasan et al. 2024). By specifically targeting essential genes in pest species, RNAi provides a highly specific and environmentally friendly approach to pest

AgriCos e-Newsletter (ISSN: 2582-7049)

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control (Ortola & Daros 2024). This paper explores the principles of RNAi, its mechanisms, recent advancements, delivery methods, and challenges in pest management. The review underscores the transformative potential of RNAi in overcoming the limitations of conventional pest control methods. It further highlights the importance of combining RNAi with other emerging technologies to develop integrated, multi-target pest management systems.

2. Mechanism of RNA Interference

RNA interference is a post-transcriptional gene-silencing mechanism triggered by double-stranded RNA (dsRNA) (Stanisławska & Olszewski, 2005). The process begins with the introduction of dsRNA, either naturally or artificially, into the target organism. The cellular machinery processes the dsRNA into small interfering RNAs (siRNAs) through the activity of Dicer, a ribonuclease enzyme (Hammond, 2005; Campbell & Choy, 2005). These siRNAs are subsequently incorporated into the RNA-induced silencing complex (RISC), where they bind to complementary messenger RNA (mRNA) sequences (Kabir et al. 2008). The Argonaute protein, a key component of RISC, cleaves the target mRNA, resulting in its degradation and the subsequent inhibition of gene expression (Hammond, 2005).

Key steps of RNAi include:

- dsRNA Recognition: Entry of double-stranded RNA into the cell via ingestion or topical delivery.
- Dicer Processing: Cleavage of dsRNA into siRNA fragments (~21-25 nucleotides) in a highly sequence-specific manner.
- RISC Assembly: siRNAs guide RISC to complementary mRNA through base-pair recognition.
- Target mRNA Degradation: Specific mRNA cleavage and silencing of gene expression, leading to impaired protein synthesis and eventual disruption of critical biological pathways.

RNAi's precision makes it a powerful tool in pest management as it ensures that only the target gene is silenced, with minimal risk to non-target species. Studies on RNAi mechanisms across diverse pest species have revealed conserved pathways, supporting its applicability across multiple taxa, including insects, mites, and nematodes.

Applications of RNAi in Pest Management

Targeting Essential Genes

RNAi has demonstrated its potential to target essential genes that regulate survival, reproduction, and metabolism in pest species. Genes involved in critical physiological processes, such as chitin biosynthesis, digestion, neural signalling, and reproductive pathways, can be silenced to induce pest mortality or impair their ability to damage crops. For instance, silencing genes associated with chitin synthase in lepidopteran pests disrupts exoskeleton development, leading to death during moulting stages. Additionally, silencing neuropeptide genes in hemipteran pests such as aphids has shown significant reductions in fecundity and feeding activity, further limiting crop damage (Willow et al. 2021).

Plant-Mediated RNAi

Plant-mediated RNAi involves the expression of dsRNA sequences within genetically modified (GM) crops. These plants produce dsRNA that is ingested by pests during feeding, triggering RNAi and silencing target genes. For example, GM maize expressing dsRNA targeting *Diabrotica virgifera virgifera* (Western corn rootworm) has shown significant pest suppression without harming non-target organisms. Plant-mediated RNAi offers long-term protection and reduces the need for repeated pesticide applications, providing a cost-effective and sustainable solution for large-scale farming. Ongoing advancements in genome-editing tools, such as CRISPR/Cas9, have further enhanced the precision and efficiency of GM crops designed for RNAi (Lu et al. 2023).

Sprayable RNAi-Based Biopesticides

RNAi can be delivered exogenously through foliar sprays containing dsRNA. Advances in nanoparticlebased carriers have improved dsRNA stability and uptake in field conditions. Sprayable RNAi biopesticides offer a non-GMO alternative to plant-mediated RNAi, providing flexibility and scalability for use in conventional agriculture. These formulations degrade naturally, reducing environmental toxicity. Recent research has shown that combining RNAi sprays with adjuvants and protective agents significantly enhances dsRNA longevity on plant surfaces, increasing the efficacy of field applications (Lu et al. 2023).

AgriCos e-Newsletter (ISSN: 2582-7049)

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Non-Target Effects and Selectivity

RNAi is inherently selective due to its sequence-specific mechanism. Careful design of dsRNA sequences minimizes the risk of silencing non-target genes in beneficial organisms, such as pollinators, natural enemies, and soil microbes. Bioinformatics tools are used to ensure sequence specificity during dsRNA design. Studies assessing the impact of RNAi on honeybees and other beneficial insects have reported negligible off-target effects when dsRNA is appropriately designed, underscoring its safety in integrated pest management (Munawar, 2023).

Advances in RNAi Delivery Systems

Efficient delivery of dsRNA is critical for the success of RNAi-based pest management. Recent advances in delivery systems include:

Nanoparticle-Based Delivery

Nanoparticles protect dsRNA from degradation and facilitate cellular uptake in pest organisms. Lipid-based nanoparticles, chitosan, and clay nanosheets have been developed to enhance dsRNA stability and bioavailability in the field. These nanoparticles enable controlled release and targeted delivery to specific pest tissues, improving the overall efficiency of RNAi treatments (Chen et al. 2018).

Plant-Mediated Delivery

Genetically modified crops expressing dsRNA provide a continuous source of RNAi, eliminating the need for repeated applications. Advances in plant transformation techniques have enabled the expression of dsRNA targeting specific pest genes. Emerging research focuses on stacking multiple dsRNAs to delay resistance evolution (Rivera & Yuan, 2012).

Topical and Sprayable Formulations

Sprayable dsRNA formulations offer a non-GMO alternative to plant-mediated RNAi. Encapsulation technologies ensure stability under field conditions and enhance dsRNA uptake. Recent innovations include biodegradable polymers that protect dsRNA from UV degradation while ensuring rapid uptake in target pests (Zhou et al. 2013).

Challenges and Future Perspectives

Despite its promise, RNAi-based pest management faces several challenges:

- RNA Stability: dsRNA is susceptible to degradation by environmental factors such as UV radiation, moisture, and microbial activity. Encapsulation technologies are being developed to address these challenges.
- Delivery Efficiency: Efficient uptake of dsRNA by target pests remains a bottleneck. Research is ongoing to optimize delivery methods, such as nanoparticles and carrier molecules.
- Resistance Evolution: Repeated exposure to dsRNA may lead to the development of resistance in pest populations. Strategies such as gene stacking and targeting multiple genes simultaneously are being explored to mitigate resistance.
- Regulatory and Public Acceptance: The deployment of RNAi-based technologies requires regulatory approval and public acceptance. Transparent risk assessments and education on RNAi's safety are essential for widespread adoption.

CONCLUSION

RNA interference represents a paradigm shift in pest management, offering a highly specific, environmentally friendly alternative to chemical pesticides. By targeting essential genes in pest organisms, RNAi minimizes off-target effects and environmental risks. Advances in delivery systems, such as nanoparticle-based carriers, plant-mediated RNAi, and encapsulated spray formulations, have enhanced the feasibility of this technology in agricultural settings. However, challenges such as RNA stability, delivery efficiency, and resistance evolution must be addressed to unlock the full potential of RNAi in pest control. As research progresses, RNAi-based solutions are poised to transform modern agriculture, providing sustainable and effective pest management strategies that align with global efforts to reduce chemical pesticide use and promote environmental conservation. Furthermore, the integration of RNAi with precision agriculture tools and bioinformatics will ensure targeted and optimized pest control, leading to a greener, safer, and more resilient agricultural future.

AgriCos e-Newsletter (ISSN: 2582-7049)

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