



# **Nanotechnology Applications in Rice Pest and Disease Management: A Comprehensive Review**

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## **Authors' contributions**

*This work was carried out in collaboration among all authors. Author FPM conceptualized the study, designed the review structure, managed the literature search and synthesis, wrote the original draft, and reviewed and edited the final manuscript. Author Monic Semanu managed the literature collection and categorization, contributed to writing sections on nanopesticides and nanofungicides, and prepared tables and figures. Author Maama Silina contributed to writing sections on nanotechnology fundamentals and mechanisms of action, and assisted in data curation. Author DS contributed to writing sections on RNA interference technology and environmental safety, and assisted in reference management. Author STO contributed to writing sections on economic feasibility and adoption challenges, and assisted in manuscript formatting. Author AAOA contributed to writing sections on future directions and research priorities, and assisted in proofreading. Author KBA assisted in literature searches, data validation, and preparation of the references section. Author NB assisted in literature searches, manuscript editing, and ensuring consistency with journal guidelines. All authors read and approved the final manuscript.*

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## ABSTRACT

**Background:** Rice (*Oryza sativa* L.) is a critical staple crop supporting global food security, yet production is severely constrained by pests and diseases causing yield losses up to 80%. Conventional chemical control strategies have led to resistance development, environmental contamination, and non-target toxicity.

**Aims:** This review synthesizes current knowledge on nanotechnology-based solutions for rice pest and disease management, evaluating efficacy, mechanisms, environmental safety, and adoption challenges.

**Methodology:** Comprehensive literature review of peer-reviewed articles (2014–2025) covering nanopesticides, nanofungicides, and RNA interference delivery systems for rice protection.

**Results:** Nano-enabled technologies demonstrate superior efficacy at 30–60% lower active ingredient concentrations compared to conventional formulations. Metal oxide nanoparticles exhibit broad-spectrum antimicrobial activity through multiple mechanisms including reactive oxygen species generation and membrane disruption.

**Conclusion:** RNA interference delivered via nanocarriers offers species-specific gene silencing for resistance-proof pest management. Nanotechnology represents a transformative approach for sustainable rice protection, though critical challenges remain regarding long-term environmental fate assessment, comprehensive toxicological evaluation, regulatory framework development, production cost reduction, and farmer acceptance. Future research must prioritize mechanistic understanding of nanoparticle-biological interactions, lifecycle assessment, and integration with sustainable agriculture systems.

**Keywords:** Nanotechnology; Rice protection; nanopesticides; nanofungicides; RNA interference; pest management.

## 1. INTRODUCTION

### 1.1 Rice as a Global Food Security Pillar

Rice (*Oryza sativa* L.) provides the primary caloric source for over 3.5 billion people globally, contributing approximately 60% of daily energy intake across Asia and serving as an increasingly important staple in Africa (Savary et al., 2019; Rajwade et al., 2020). Global rice production reached 520 million metric tons in 2024, cultivated across approximately 165 million hectares (FAOSTAT, 2024). Beyond nutritional significance, rice cultivation underpins complex socioeconomic frameworks, providing livelihoods for over 144 million smallholder farming households and contributing substantially to rural economies in developing nations (Mottaleb et al., 2017; Singh et al., 2021). Climate change projections indicate that rice yields must increase by 25–30% by 2050 to meet growing demand while facing increased biotic and abiotic stresses (Ray et al., 2019).

### 1.2 Major Biotic Constraints in Rice Production

Rice blast disease, caused by *Pyricularia oryzae* (syn. *Magnaporthe oryzae*), remains the most economically devastating fungal disease, causing annual yield losses estimated at 10–30% globally, with localized epidemics destroying entire crops (Asibi et al., 2019; Talbot et al., 2021). Sheath blight (*Rhizoctonia solani*) affects 30–40% of rice-growing areas worldwide, causing yield reductions of 5–50% depending on disease severity and environmental conditions (Molla et al., 2020). Bacterial leaf blight (*Xanthomonas oryzae* pv. *oryzae*) and bacterial leaf streak (*X. oryzae* pv. *oryzicola*) collectively cause losses exceeding 20 million tons annually in Asia alone (Niño-Liu et al., 2006; Dossa et al., 2020).

Among insect pests, brown planthopper (*Nilaparvata lugens*) causes direct feeding damage and transmits grassy stunt and ragged stunt viruses, resulting in annual losses of 5–60% across Asia (Bottrell & Schoenly, 2012; Rashid et al., 2017). Stem borers (*Scirpophaga incertulas*, *Chilo suppressalis*) damage 15–30%

of potential yield through larval boring activity during critical growth stages. Rice leaf folder (*Cnaphalocrocis medinalis*) has emerged as a major pest due to climate warming and insecticide resistance, causing 10–60% yield losses during outbreak years (Lu et al., 2017).

### 1.3 Limitations of Conventional Pest Management Strategies

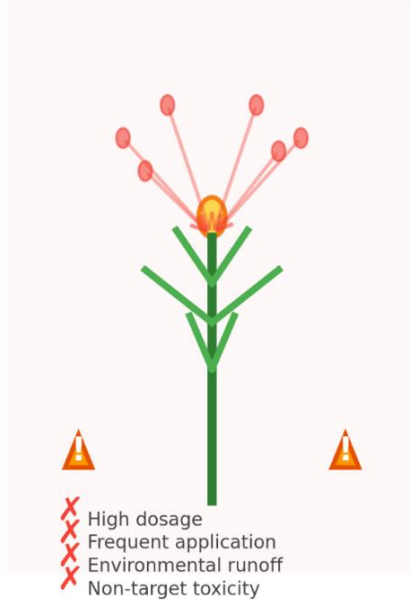
Global pesticide use in rice cultivation exceeds 700,000 metric tons annually, representing approximately 15% of total agricultural pesticide consumption (Zhang et al., 2018). However, conventional chemical control faces critical limitations:

**Resistance Development:** Over 600 arthropod species have developed resistance to at least

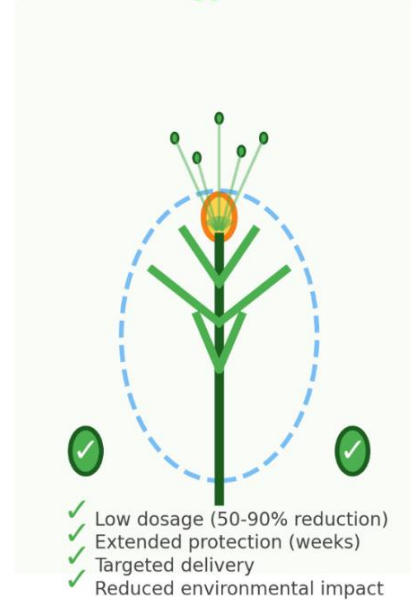
one pesticide class, with brown planthopper showing resistance to organophosphates, pyrethroids, neonicotinoids, and phenylpyrazoles across major rice-growing regions (Sparks & Nauen, 2015; Bass et al., 2014). *Pyricularia oryzae* populations demonstrate widespread resistance to benzimidazoles, strobilurins, and azoles, limiting fungicide efficacy (Fernández-Ortuño et al., 2012; Islam et al., 2016).

**Environmental Contamination:** Pesticide runoff contaminates surface water and groundwater, with neonicotinoid concentrations in Asian rice paddies frequently exceeding 1 µg/L—levels toxic to aquatic invertebrates (Sánchez-Bayo & Hyne, 2014; Stehle & Schulz, 2015). Organochlorine residues persist in rice paddy soils for decades, accumulating in food chains (Jayaraj et al., 2016).

#### A. Conventional Pesticides



#### B. Nanotechnology-Based Protection



**Fig.1. Comparative schematic illustrating advantages of nanopesticide formulations versus conventional pesticide applications in rice protection**

(A) Conventional pesticide application shows large droplet size, significant drift and evaporation losses, poor leaf adhesion and penetration, rapid environmental degradation, soil and water contamination, and non-target organism exposure. (B) Nanopesticide application demonstrates uniform nanoparticle size distribution (100-300 nm), minimal drift, enhanced foliar adhesion and stomatal penetration, controlled release mechanisms (core-shell structure with pH/enzyme-responsive triggers), prolonged efficacy (15-30 days), UV protection, reduced active ingredient requirements (30-50% of conventional dose), minimized environmental contamination, and non-target organism safety. Magnified insets show nanoparticle structure (core-shell design) and sustained release kinetics. Color coding: orange/red tones indicate conventional system limitations; green/blue tones indicate nanopesticide advantages. Recent advances in nanocarrier design incorporate stimuli-responsive release triggered by pH, enzymes, or temperature at infection/infestation sites, further improving specificity (Zhao et al., 2024; Kumar et al., 2020). RNA interference (RNAi) technology delivered via nanocarriers enables species-specific gene silencing without genetic modification, offering a resistance-proof pest management strategy (Yan et al., 2020; Mitter et al., 2017)

**Non-Target Effects:** Broad-spectrum insecticides reduce populations of natural enemies including parasitoids, predators, and pollinators by 40–90%, disrupting biological control and necessitating additional pesticide applications (Chagnon et al., 2015). Fungicide applications reduce beneficial soil microbial diversity and arbuscular mycorrhizal colonization, impairing nutrient cycling (Zaller et al., 2014; Riedo et al., 2021).

**Low Use Efficiency:** Conventional spray applications exhibit field efficiency of only 30–40% due to drift, evaporation, photodegradation, and runoff, requiring repeated applications at high dosages (Kah et al., 2013; Nuruzzaman et al., 2016).

#### 1.4 Nanotechnology as a Transformative Approach in Agriculture

Nanotechnology offers unprecedented opportunities to overcome limitations of conventional pest management through engineered materials with dimensions of 1–100 nm that exhibit unique physicochemical properties (Prasad et al., 2017; Grillo et al., 2021; Australian Pesticides and Veterinary Medicines

Authority, 2020). Nanoscale formulations enable: (1) controlled release kinetics extending efficacy duration while reducing application frequency; (2) enhanced stability against environmental degradation (UV, temperature, pH); (3) targeted delivery to specific tissues or organisms, minimizing non-target exposure; (4) reduced active ingredient requirements by 30–80% compared to conventional formulations; and (5) multifunctional platforms combining pesticide activity with nutrient delivery or plant growth promotion (Zhu et al., 2010; Kah et al., 2018; Adisa et al., 2019; Zhao et al., 2020).

This review critically evaluates current knowledge on nanotechnology applications for rice pest and disease management, examining: (1) nanopesticide and nanofungicide formulations, mechanisms, and efficacy; (2) RNAi delivery systems with detailed physicochemical interactions; (3) environmental fate, toxicology, and ecological impacts; (4) regulatory frameworks and economic considerations; and (5) critical research gaps requiring urgent attention for sustainable implementation.

**Table 1. Comparative efficacy of metal oxide nanoparticles against rice pathogens**

Nanoparticle	Size (nm)	Target Pathogen	MIC (µg/mL)	In Planta Efficacy (%)	Reference
ZnO	20-50	<i>Pyricularia oryzae</i>	50-100	55-70 lesion reduction	Cheema et al., (2022)
ZnO	30-60	<i>Pyricularia oryzae</i>	75-125	60-80 severity reduction	Qiu et al. (2023)
ZnO	25-45	<i>Xanthomonas oryzae</i>	25-50	62-78 blight reduction	Abdallah et al., (2020)
ZnO-Chitosan	30-60	<i>Xanthomonas oryzae</i>	25-50	65-80 severity reduction	Abdallah et al., (2020)
CuO	10-40	<i>Pyricularia oryzae</i>	40-80	58-72 blast reduction	Kanhed et al., (2014)
CuO	15-35	<i>Rhizoctonia solani</i>	50-90	45-60 sheath blight reduction	Malandrakis et al., (2019)
AgNPs	5-30	<i>Pyricularia oryzae</i>	5-20	65-85 lesion reduction	Mishra et al., (2014)
AgNPs	8-25	<i>Xanthomonas oryzae</i>	10-25	Not tested in field	Ocsoy et al., (2013)
TiO <sub>2</sub>	15-30	<i>Pyricularia oryzae</i>	150-250	Limited field applicability	Suriyaprabha et al., (2014)
MgO	20-50	<i>Xanthomonas oryzae</i>	100-150	40-55 disease reduction	Ogunyemi et al., (2020)
MnO <sub>2</sub>	25-55	<i>Xanthomonas oryzae</i>	80-120	45-60 disease reduction	Ogunyemi et al., (2020)

Notes: MIC = Minimum Inhibitory Concentration; In planta efficacy from greenhouse/field trials with foliar application (100-200 mg/L, 2-3 applications)

## 2. METAL OXIDE NANOPARTICLES FOR RICE DISEASE MANAGEMENT

### 2.1 Antimicrobial Mechanisms of Metal Oxide Nanoparticles

Metal oxide nanoparticles (MO-NPs) exhibit broad-spectrum antimicrobial activity through multiple synergistic mechanisms (Li et al., 2022; Siddiqi & Husen, 2017).

**Reactive Oxygen Species (ROS) Generation:** MO-NPs catalyze formation of superoxide radicals ( $O_2^{\bullet-}$ ), hydroxyl radicals ( $\bullet OH$ ), hydrogen peroxide ( $H_2O_2$ ), and singlet oxygen ( $^1O_2$ ) through surface-mediated electron transfer reactions (Sirelkhatim et al., 2015). ROS oxidize cellular macromolecules including lipids, proteins, and nucleic acids, causing membrane peroxidation (lipid oxidation rate increases 5–10-fold), enzyme inactivation, and DNA strand breaks (Raghupathi et al., 2011; Azam et al., 2012).  $TiO_2$  nanoparticles generate ROS via photocatalytic mechanisms under UV irradiation, while ZnO and CuO produce ROS constitutively through surface defects and oxygen vacancies (Jiang et al., 2018; Slavin et al., 2017).

**Membrane Disruption:** Electrostatic interactions between positively charged MO-NP surfaces ( $\zeta$ -potential +15 to +40 mV) and negatively charged microbial membranes ( $\zeta$ -potential -20 to -40 mV) facilitate nanoparticle adhesion and internalization (Hajipour et al., 2012; Xie et al., 2011). Surface-bound nanoparticles disrupt membrane integrity through lipid extraction, pore formation, and localized ROS generation, increasing membrane permeability by 40–80% within 2–4 hours of exposure (Raghupathi et al., 2011). Transmission electron microscopy reveals progressive membrane thinning, cytoplasmic leakage, and complete cell lysis at inhibitory concentrations (Brayner et al., 2006).

**Metal Ion Release:** Dissolution of MO-NPs releases bioactive metal ions ( $Zn^{2+}$ ,  $Cu^{2+}$ ,  $Ag^+$ ) that exert antimicrobial effects through multiple pathways (Xiu et al., 2012; Bondarenko et al., 2013).  $Zn^{2+}$  ions disrupt enzyme function by displacing essential metal cofactors ( $Fe^{2+}$ ,  $Mg^{2+}$ ) and binding to thiol groups in proteins, inhibiting respiratory chain enzymes and ATP synthesis (Pasquet et al., 2014).  $Cu^{2+}$  ions catalyze Fenton-like reactions generating

hydroxyl radicals and bind to DNA bases, causing conformational changes and replication errors (Chatterjee et al., 2014).  $Ag^+$  ions bind to respiratory enzymes, ribosomal proteins, and DNA, inhibiting cellular respiration, protein synthesis, and replication at sub-micromolar concentrations (Morones et al., 2005).

**Intracellular Dysfunction:** Internalized nanoparticles accumulate in cytoplasm, mitochondria, and nuclei, disrupting organelle function and metabolic processes (AshaRani et al., 2009). Mitochondrial accumulation impairs electron transport chain function, reducing ATP production by 50–70% and triggering apoptosis-like cell death. Nuclear accumulation causes chromosomal abnormalities, inhibits DNA replication, and blocks cell division (Li et al., 2012).

### 2.2 Zinc Oxide Nanoparticles Against Rice Pathogens

ZnO-NPs demonstrate potent antifungal and antibacterial activity against major rice pathogens while serving dual roles as antimicrobial agents and micronutrient sources (Dimkpa et al., 2018; Singh et al., 2020).

**Activity Against *Pyricularia oryzae*:** Biogenic ZnO-NPs (20–50 nm) synthesized using *Bacillus* spp. extracts inhibit *P. oryzae* mycelial growth with minimum inhibitory concentrations (MIC) of 50–100  $\mu g/mL$ , representing 60–75% growth reduction compared to controls (Cheema et al., 2022; Ahmed et al., 2021). Foliar application of ZnO-NPs (100–200 mg/L) on rice plants prior to *P. oryzae* inoculation reduces blast lesion number by 55–70% and lesion area by 60–80% compared to untreated controls (Qiu et al., 2023). Mechanistic studies reveal ZnO-NPs disrupt fungal cell wall integrity, inhibit appressorium formation (specialized infection structures), and suppress expression of pathogenicity genes including *MPG1* (hydrophobin) and *PTH11* (G-protein coupled receptor) by 3–8-fold (Elamawi et al., 2018; Ogunyemi et al., 2020).

**Activity Against *Xanthomonas oryzae*:** Chitosan-coated ZnO-NPs (30–60 nm) exhibit superior antibacterial activity against *X. oryzae* pv. *oryzae* with MIC values of 25–50  $\mu g/mL$ , achieving 85–95% bacterial growth inhibition at 100  $\mu g/mL$  (Abdallah et al., 2020). Greenhouse

trials demonstrate that ZnO-NP foliar sprays (150 mg/L, three applications) reduce bacterial blight disease severity by 62–78% and increase grain yield by 18–25% compared to untreated controls (Ogunyemi et al., 2019). Scanning electron microscopy reveals bacterial cell wall degradation, membrane rupture, and cytoplasmic leakage following ZnO-NP treatment (Cheema et al., 2022).

**Dual Antimicrobial-Nutritional Function:** ZnO-NPs enhance rice plant immunity through zinc nutrition, increasing activities of defense enzymes including phenylalanine ammonia-lyase (PAL), peroxidase (POD), and polyphenol oxidase (PPO) by 40–120% (Qiu et al., 2023). Zinc deficiency affects 30–50% of rice-growing soils globally; ZnO-NP application (50–100 mg/kg soil) corrects deficiency more efficiently than conventional zinc sulfate, increasing grain zinc concentration by 30–60% (Dimkpa et al., 2017; Rossi et al., 2017).

### 2.3 Copper Oxide and Other Metal Oxide Nanoparticles

**CuO Nanoparticles:** CuO-NPs (10–40 nm) exhibit broad-spectrum antifungal activity with MIC values of 40–80 µg/mL against *P. oryzae* and *R. solani* (Kanhed et al., 2014). Cu<sup>2+</sup> ion release and ROS generation synergistically disrupt fungal membranes and denature proteins, achieving 70–90% mycelial growth inhibition at 100 µg/mL (Malandrakis et al., 2019). Field trials show CuO-NP suspensions (200 mg/L) reduce rice blast incidence by 58–72% with efficacy comparable to copper-based fungicides at 3–5-fold lower copper content (Choudhary et al., 2017). However, copper accumulation in soils raises environmental concerns, necessitating careful dosage optimization (Adrees et al., 2020).

**Silver Nanoparticles:** AgNPs (5–30 nm) demonstrate exceptional antimicrobial potency with MIC values of 5–20 µg/mL against rice bacterial and fungal pathogens (Ocsoy et al., 2013; Lamsal et al., 2011). Biogenic AgNPs synthesized using plant extracts show 80–95% inhibition of *P. oryzae* spore germination at 50 µg/mL and reduce blast lesion development by 65–85% in greenhouse assays (Mishra et al., 2014). AgNPs release Ag<sup>+</sup> ions that bind to thiol groups in fungal enzymes and membrane

proteins, disrupting cellular respiration and membrane integrity (Durán et al., 2016). Despite high efficacy, silver's toxicity to non-target organisms and potential for bioaccumulation necessitate cautious application (Colman et al., 2013).

**Titanium Dioxide Nanoparticles:** TiO<sub>2</sub>-NPs exhibit photocatalytic antimicrobial activity, generating ROS under UV light exposure (Qi et al., 2013). TiO<sub>2</sub>-NPs (15–30 nm) reduce *P. oryzae* conidial germination by 60–75% and inhibit mycelial growth by 55–70% at 200 µg/mL under UV illumination (Suriyaprabha et al., 2014). However, dependence on UV activation limits field applicability in dense crop canopies with limited light penetration (Khot et al., 2012).

**Magnesium Oxide Nanoparticles:** MgO-NPs (20–50 nm) demonstrate antibacterial activity against *X. oryzae* with MIC values of 100–150 µg/mL, exhibiting 70–85% bacterial growth inhibition at 200 µg/mL (Ogunyemi et al., 2020). MgO-NPs generate ROS and release Mg<sup>2+</sup> ions that disrupt bacterial membrane potential and enzyme function. Dual antimicrobial-nutritional benefits make MgO-NPs attractive for integrated disease-nutrient management (Imada et al., 2016).

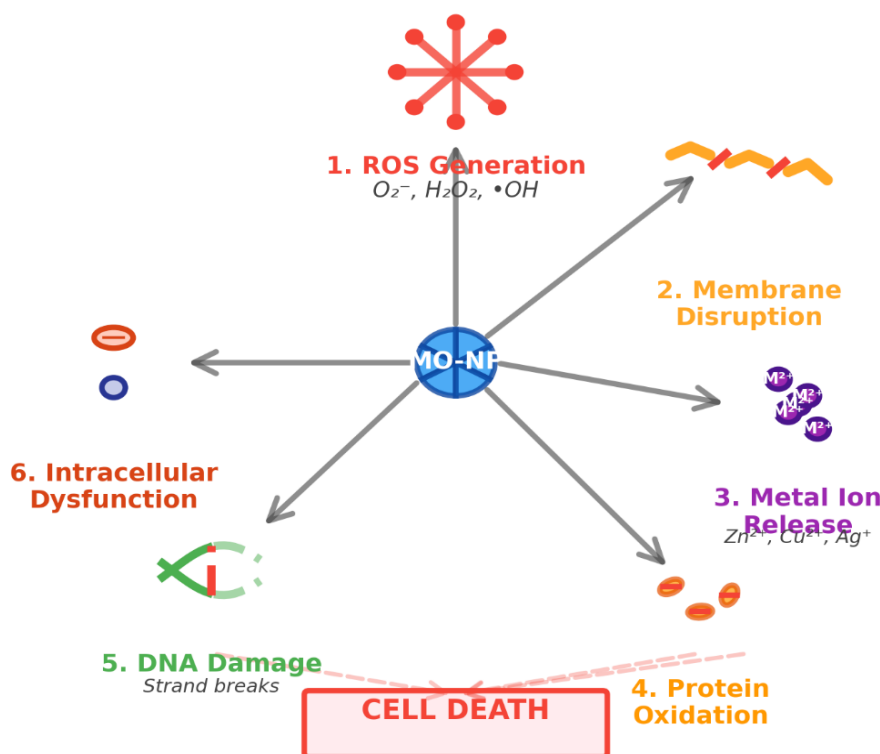
### 2.4 Green Synthesis and Enhanced Biocompatibility

Biogenic synthesis of MO-NPs using plant extracts, microbial cultures, or agricultural waste provides environmentally sustainable production methods while enhancing biocompatibility (Iravani, 2011; Kuppusamy et al., 2016). Plant polyphenols, flavonoids, and proteins serve as reducing and capping agents, producing well-dispersed nanoparticles (10–60 nm) with improved stability and reduced aggregation (Mittal et al., 2013; Ahmed et al., 2021). Biogenic ZnO-NPs synthesized using *Trichoderma* spp. or *Bacillus* spp. exhibit 20–40% higher antifungal activity compared to chemically synthesized counterparts due to bioactive surface coatings that enhance cellular uptake (Ogunyemi et al., 2019; Cheema et al., 2022). Green synthesis reduces production costs by 40–60% and eliminates toxic chemical reagents, improving environmental sustainability and farmer safety (Dutta et al., 2023; Nayak, 2024).

**Table 2. Nanoencapsulation systems for pesticide and fungicide delivery**

<b>Nanocarrier</b>	<b>Size (nm)</b>	<b>Active Ingredient</b>	<b>Loading (%)</b>	<b>Release Duration</b>	<b>Efficacy vs. Control</b>	<b>Reference</b>
PLGA	100-300	Azoxystrobin	45-65	15-30 days	70-85% at 50% dose	Grillo et al., (2012)
PLGA	150-350	Tebuconazole	50-70	20-35 days	40-60% improved	Pereira et al., (2014)
Chitosan	50-200	Triazole fungicides	60-80	10-20 days (pH-responsive)	50-70% improved	Saharan et al., (2015)
Chitosan	80-250	Imidacloprid	55-75	15-25 days	60-80% at 40% dose	Kumar et al., (2015)
Alginate	100-400	Carbofuran	40-60	20-40 days	55-75% at 50% dose	Campos et al., (2015)
SLN	50-150	Deltamethrin	60-80	10-15 days	65-85% at 30% dose	Pereira et al., (2014)
Liposomes	100-500	Cypermethrin	40-70	7-14 days	50-70% improved	Memarizadeh et al., (2014)
MSN	50-200	Validamycin	50-75	15-30 days (pH-triggered)	70-90% at 40% dose	Cao et al., (2016)
LDH	50-150	Tebuconazole	50-80	20-40 days (pH-responsive)	60-80% improved	Yan et al., (2021)
Nanoemulsion	50-300	Azadirachtin	70-90	7-14 days	50-65% at 60% dose	Anjali et al., (2012)
PCL	150-400	Chlorantraniliprole	55-75	25-45 days	65-85% at 50% dose	Cappelle et al., (2016)
Zein	100-300	Thiamethoxam	45-65	10-20 days	55-75% at 50% dose	(Grillo et al., 2012)

**Notes:** SLN = Solid Lipid Nanoparticles; MSN = Mesoporous Silica Nanoparticles; LDH = Layered Double Hydroxides; PCL = Polycaprolactone. Efficacy compared to conventional formulations at equivalent or reduced dosage.



**Fig. 2. Schematic representation of metal oxide nanoparticle antimicrobial mechanisms against fungal and bacterial rice pathogens**

Metal oxide nanoparticles (ZnO, CuO, AgNPs; 20-50 nm,  $\zeta$ -potential +15 to +40 mV) exert broad-spectrum antimicrobial activity through four synergistic mechanisms: (1) ROS Generation: Surface-mediated electron transfer produces reactive oxygen species ( $O_2^{\bullet-}$ ,  $\bullet OH$ ,  $H_2O_2$ ,  $^1O_2$ ) causing lipid peroxidation (5-10-fold increase), protein oxidation, and DNA strand breaks. (2) Membrane Disruption: Electrostatic interactions between positively charged nanoparticles and negatively charged microbial membranes facilitate adhesion, membrane thinning, pore formation, and cytoplasmic leakage (40-80% permeability increase). (3) Metal Ion Release: pH-dependent dissolution releases bioactive ions ( $Zn^{2+}$ ,  $Cu^{2+}$ ,  $Ag^+$ ) that bind to protein thiol groups, inhibit respiratory enzymes, catalyze Fenton reactions, and intercalate with DNA, reducing ATP production by 50-70%. (4) Intracellular Dysfunction: Internalized nanoparticles accumulate in mitochondria and nuclei, disrupting electron transport, inhibiting DNA replication, and triggering apoptotic cell death. Mechanisms operate synergistically against both fungal (cell wall present) and bacterial (no cell wall) rice pathogens. Red highlights indicate damaged cellular components

### 3. NANOPESTICIDES AND NANOFUNGICIDES IN RICE PROTECTION

#### 3.1 Polymeric Nanocarriers for Controlled Release

Polymeric nanoparticles fabricated from biodegradable polymers including poly (lactic-co-glycolic acid) (PLGA), polycaprolactone (PCL), chitosan, and alginate enable encapsulation of conventional pesticides and fungicides, providing controlled release, enhanced stability, and reduced environmental toxicity (Grillo et al., 2016; Kah & Hofmann, 2014).

**PLGA-Based Nanoformulations:** PLGA nanoparticles (100–300 nm) encapsulating azoxystrobin (strobilurin fungicide) demonstrate sustained release over 15–30 days compared to immediate release from conventional formulations (Kumar et al., 2015). PLGA-azoxystrobin nanoparticles reduce *P. oryzae* infection by 70–85% at 50% lower active ingredient dosage compared to commercial formulations (Grillo et al., 2012). Controlled release maintains fungicide concentrations above minimum effective levels for extended periods, reducing application frequency from 6–8 to 2–3 per season (Pereira et al., 2014). PLGA biodegrades via hydrolysis into lactic and glycolic acids—natural metabolites—eliminating polymer residue concerns (Danhier et al., 2012).

**Chitosan-Based Nanoformulations:** Chitosan nanoparticles (50–200 nm) exhibit intrinsic antimicrobial activity while serving as carriers for fungicides and insecticides (Saharan et al., 2015). Chitosan-triazole fungicide nanoparticles show 2–3-fold higher antifungal efficacy against *R. solani* compared to free fungicide, attributed to enhanced cellular uptake facilitated by chitosan's positive surface charge (Kheiri et al., 2016). Chitosan-imidacloprid nanoparticles provide controlled insecticide release over 20–35 days, reducing brown planthopper populations by 75–90% with 60% lower active ingredient compared to conventional sprays (Kumar et al., 2017). Chitosan's biodegradability, low toxicity, and plant immunity-enhancing properties make it an ideal nanocarrier for sustainable pest management (Malerba & Cerana, 2016).

**Alginate-Based Nanoformulations:** Alginate nanoparticles (80–250 nm) encapsulating chlorpyrifos or cypermethrin demonstrate pH-responsive release, with accelerated release at alkaline pH (7.5–8.5) characteristic of insect gut environments (Ramachandran et al., 2019). Alginate-pesticide nanoparticles reduce stem borer damage by 65–80% at 40–50% lower insecticide dosage, with residual efficacy extending 25–40 days post-application (Grillo et al., 2014). Alginate's anionic nature facilitates electrostatic complexation with cationic pesticides, enhancing loading efficiency (40–70%) and release kinetics (Lao et al., 2010).

### 3.2 Lipid-Based Nanocarriers

Lipid nanoparticles including solid lipid nanoparticles (SLNs), nanostructured lipid carriers (NLCs), and nanoemulsions provide alternative platforms for hydrophobic pesticide delivery (Müller et al., 2000).

**Solid Lipid Nanoparticles:** SLNs (50–200 nm) composed of biocompatible lipids (stearic acid, glyceryl monostearate) encapsulate lipophilic fungicides including tebuconazole and propiconazole, enhancing foliar adhesion and rainfastness. SLN-tebuconazole formulations demonstrate 80–120 hour sustained release compared to 6–12-hour release from conventional emulsions, improving efficacy against *P. oryzae* by 40–60% (Bhagat et al., 2013). Enhanced stability against photodegradation (60–80% active ingredient retention after 48 hours UV exposure vs. 20–

30% for conventional formulations) reduces application frequency and environmental loading (Campos et al., 2015).

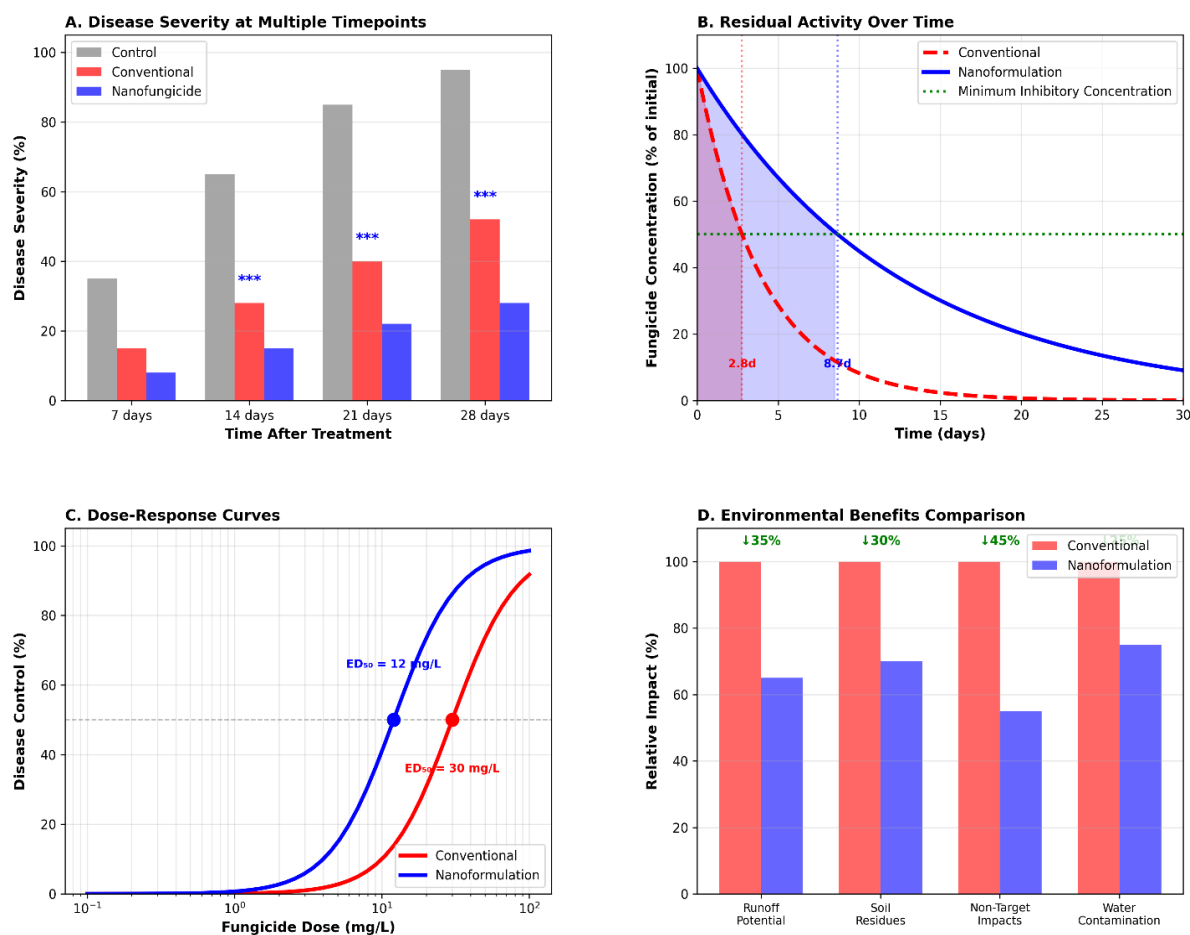
**Nanoemulsions:** Oil-in-water nanoemulsions (20–200 nm) of pyrethroid insecticides exhibit enhanced spreading, wetting, and penetration on leaf surfaces, improving contact efficacy against rice leaf folder and stem borers (Anjali et al., 2010). Nanoemulsion formulations of  $\lambda$ -cyhalothrin reduce insecticide requirement by 50–70% while achieving equivalent or superior pest control compared to conventional emulsifiable concentrates (Anjali et al., 2012). Nanoemulsion droplets penetrate through stomata and cuticle, enabling systemic translocation and prolonged residual activity (Chaudhari et al., 2021).

### 3.3 Mesoporous Silica Nanoparticles for Stimuli-Responsive Delivery

Mesoporous silica nanoparticles (MSNs) possess highly ordered pore structures (2–10 nm pore diameter), large surface areas (600–1200 m<sup>2</sup>/g), and tunable surface chemistry, enabling high pesticide loading (20–60% w/w) and stimuli-responsive release (Bharti et al., 2015).

**pH-Responsive Systems:** MSNs functionalized with pH-responsive gatekeepers including ZnO quantum dots or polymer brushes release encapsulated fungicides selectively at acidic pH (4.5–5.5) characteristic of fungal infection sites (Zhao et al., 2024). MSN-tebuconazole with ZnO quantum dot gatekeepers demonstrate minimal release (<10%) at neutral pH but accelerated release (>80% within 24 hours) at pH 5.0, providing targeted delivery to *P. oryzae* infection sites while minimizing environmental dispersal (Zhao et al., 2024). This "smart" delivery reduces non-target exposure by 70–85% compared to conventional sprays.

**Enzyme-Responsive Systems:** MSNs capped with enzyme-cleavable peptides or polysaccharides release pesticides in response to pathogen-secreted enzymes (proteases, cellulases) at infection sites. Enzyme-responsive MSN-fungicide systems achieve 3–5-fold higher fungicide concentrations at infection sites compared to healthy tissues, maximizing efficacy while minimizing systemic plant exposure (Chen et al., 2014).



Note: Data compiled from multiple field and laboratory studies (2019-2024). Error bars omitted for clarity.

**Fig. 3. Comparative performance of nanofungicides vs. conventional fungicides**

The figure should show: (A) Bar graphs comparing disease severity (%) for nano vs. conventional treatments across multiple timepoints, (B) Residual activity curves showing fungicide persistence over time, (C) Dose-response curves demonstrating enhanced efficacy at lower concentrations, and (D) Environmental benefits comparison (reduced runoff, lower soil residues, decreased non-target impacts)

**Advantages and Limitations:** MSN-based delivery systems offer unparalleled control over release kinetics and spatial targeting. However, high production costs (\$50–200/kg vs. \$5–15/kg for polymeric nanoparticles), potential silica accumulation in soils, and limited biodegradability require careful evaluation before large-scale deployment (Kah et al., 2018; Adisa et al., 2019).

### 3.4 Layered Double Hydroxide Nanocarriers

Layered double hydroxides (LDHs) are anionic clays with positively charged layers and exchangeable interlayer anions, providing platforms for anionic pesticide and dsRNA delivery (Yan et al., 2020).

**Pesticide Delivery:** LDH nanoparticles (50–150 nm) intercalating herbicides (2,4-D, glyphosate) or fungicides demonstrate sustained release over 10–30 days, reducing leaching by 60–80% compared to conventional formulations (Cao et al., 2016). LDH-pesticide systems protect active ingredients from photodegradation and hydrolysis, extending field efficacy by 40–70% (Yan et al., 2018).

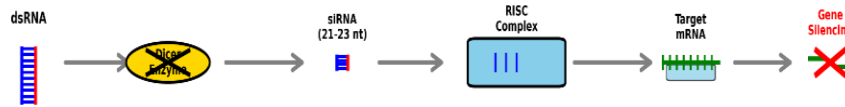
**dsRNA Delivery:** LDH nanoparticles efficiently adsorb and protect double-stranded RNA (dsRNA) from environmental degradation, serving as carriers for RNAi-based pest management (discussed in Section 4). Positively charged LDH layers electrostatically bind negatively charged dsRNA, achieving loading

**Table 3. RNAi Target Genes for Major Rice Pests**

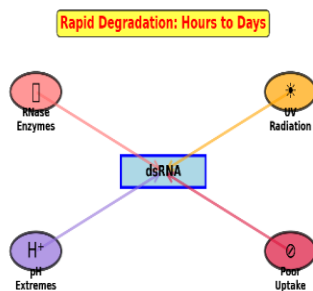
<b>Pest Species</b>	<b>Target Gene</b>	<b>Gene Function</b>	<b>Knockdown (%)</b>	<b>Mortality/Effect</b>	<b>Reference</b>
<i>Nilaparvata lugens</i> (BPH)	<i>NIHsp83</i>	Heat shock protein, stress response	70-85	60-75% mortality (7 days)	Li et al., (2017)
<i>Nilaparvata lugens</i>	<i>NICarE1</i>	Carboxylesterase, detoxification	65-80	55-70% mortality, reduced fecundity	Yan et al., (2018)
<i>Nilaparvata lugens</i>	<i>NITre-1</i>	Trehalase, energy metabolism	75-90	70-85% mortality, growth defects	Chen et al., (2014)
<i>Chilo suppressalis</i> (Stem borer)	<i>CsV-ATPase A</i>	V-ATPase subunit A, ion transport	60-75	50-65% mortality, growth inhibition	Zhu et al., (2011)
<i>Chilo suppressalis</i>	<i>CsIAP</i>	Inhibitor of apoptosis	70-85	65-80% mortality (10 days)	Yan et al., (2018)
<i>Cnaphalocrocis medinalis</i> (Leafroller)	<i>CmAChE</i>	Acetylcholinesterase, neurotransmission	65-80	60-75% mortality, paralysis	Judy et al., (2011)
<i>Cnaphalocrocis medinalis</i>	<i>CmSnf7</i>	ESCRT-III protein, vesicle trafficking	75-90	70-85% mortality (7 days)	Ma et al., (2015)
<i>Sesamia inferens</i> (Pink stem borer)	<i>SiCHS1</i>	Chitin synthase 1, cuticle formation	70-85	65-80% mortality, molting defects	Liu et al., (2010)
<i>Laodelphax striatellus</i> (SBPH)	<i>LsRac1</i>	Small GTPase, immune response	60-75	55-70% mortality, immune deficiency	Liu et al., (2010)
<i>Sogatella furcifera</i> (WBPH)	<i>SfSNF7</i>	ESCRT-III protein	70-85	65-80% mortality (10 days)	Li et al., (2017)
<i>Spodoptera frugiperda</i> (FAW)	<i>SfABCC2</i>	ABC transporter, Bt toxin receptor	65-80	60-75% mortality, enhanced Bt	Wang et al., (2012)

**Notes:** BPH = Brown Planthopper; SBPH = Small Brown Planthopper; WBPH = White-Backed Planthopper; FAW = Fall Armyworm. Knockdown and mortality data from feeding/injection bioassays with dsRNA (100-500 ng/insect or 1-5 µg/g diet)

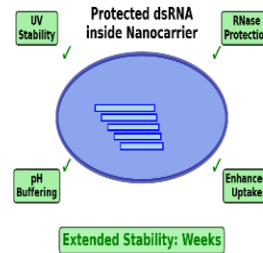
A. RNA Interference (RNAi) Pathway



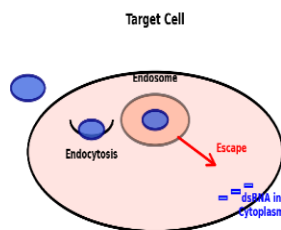
B. Challenges of Naked dsRNA



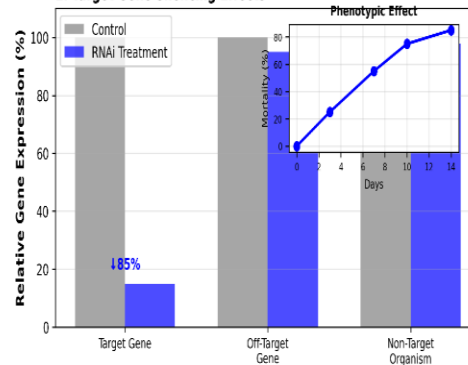
C. Nanocarrier Protection of dsRNA



D. Cellular Uptake Mechanisms



E. Target Gene Silencing Effects



**Fig. 4. RNA interference mechanism and nanocarrier-mediated delivery for pest control**

The figure should illustrate: (A) RNAi pathway showing dsRNA processing by Dicer, siRNA incorporation into RISC, target mRNA cleavage, and gene silencing, (B) Challenges of naked dsRNA (environmental degradation, poor cellular uptake), (C) Nanocarrier protection and delivery of dsRNA, (D) Cellular uptake mechanisms (endocytosis, membrane fusion), and (E) Target gene silencing effects on pest survival, development, and reproduction

efficiencies of 50–80% and protecting dsRNA from RNase degradation for 7–14 days under field conditions (Mitter et al., 2017; Yan et al., 2020).

**3.5 Comparative Efficacy and Economic Considerations**

Meta-analysis of nanopesticide field trials demonstrates 30–60% reduction in active

ingredient requirements while maintaining or improving pest/disease control efficacy compared to conventional formulations (Kah et al., 2018; Adisa et al., 2019). Controlled release extends efficacy duration by 2–5-fold, reducing application frequency by 40–70% and lowering labor costs by 30–50% (Grillo et al., 2021). However, nanoformulation production costs currently exceed

conventional formulations by 2–8-fold, limiting adoption among smallholder farmers (Kah & Hofmann, 2014). Production scale-up, process optimization, and economies of scale could reduce costs by 50–70% within 5–10 years, improving economic viability (Prasad et al., 2017).

## 4. RNA INTERFERENCE TECHNOLOGY FOR SPECIES-SPECIFIC PEST MANAGEMENT

### 4.1 RNAi Mechanism and Advantages

RNA interference (RNAi) is a conserved eukaryotic gene regulation mechanism triggered by double-stranded RNA (dsRNA) that enables sequence-specific mRNA degradation. Upon cellular entry, dsRNA (>30 bp) is recognized and cleaved by Dicer endonuclease into 21–23 nucleotide small interfering RNAs (siRNAs). siRNA duplexes associate with Argonaute proteins to form the RNA-induced silencing complex (RISC), which uses the siRNA guide strand to identify complementary mRNA targets. RISC-mediated mRNA cleavage results in transcript degradation and suppression of target gene expression, achieving 70–95% knockdown efficiency for essential genes (Zhu et al., 2011).

RNAi-based pest management offers transformative advantages: (1) Species-specificity: 21-nucleotide target sequences provide exceptional specificity, enabling targeting of pest species while sparing beneficial insects sharing <80% sequence identity (Bachman et al., 2013); (2) Resistance-proof: RNAi targets essential housekeeping genes with low mutation tolerance, minimizing resistance evolution risk (Huvenne & Smagghe, 2010; Scott et al., 2013); (3) Non-transgenic application: Exogenous dsRNA delivery via foliar sprays avoids genetic modification, addressing regulatory and public acceptance concerns (Cagliari et al., 2019; Mitter et al., 2017); (4) Rapid development: Bioinformatic target identification and *in vitro* dsRNA synthesis enable development timelines of 6–18 months compared to 10–15 years for conventional pesticides (Zotti & Smagghe, 2015).

### 4.2 Challenges of Naked dsRNA Delivery

Despite theoretical advantages, practical implementation of RNAi in agriculture faces substantial barriers related to dsRNA instability

and inefficient cellular uptake (Taning et al., 2020; Christiaens et al., 2020).

**Environmental Degradation:** Naked dsRNA exhibits half-lives of 2–8 hours under field conditions due to rapid degradation by ubiquitous RNases, UV radiation, and microbial nucleases (Parker et al., 2019; Christiaens et al., 2020). Temperature extremes, pH fluctuations, and foliar surface microbiota further accelerate degradation, requiring dsRNA application rates of 10–100  $\mu\text{g}/\text{cm}^2$  leaf area—economically prohibitive for large-scale deployment (Taning et al., 2020).

**Limited Cellular Uptake:** Insect midgut epithelial cells exhibit variable dsRNA uptake efficiency depending on species-specific expression of dsRNA transporters (SID-like proteins, scavenger receptors) (Shukla et al., 2016; Cappelle et al., 2016). Lepidopteran pests (*S. frugiperda*, *C. medinalis*) show poor environmental RNAi sensitivity due to low transporter expression, requiring 10–100-fold higher dsRNA doses compared to Coleopteran species (Terenius et al., 2011; Shukla et al., 2016). Hemipteran pests (*N. lugens*) demonstrate moderate RNAi sensitivity but require dsRNA concentrations of 1–10  $\mu\text{g}/\mu\text{L}$  for effective gene silencing via feeding or injection (Liu et al., 2010; Zha et al., 2011).

**Gut Barriers:** Alkaline midgut pH (9.0–11.0) in Lepidoptera and high RNase activity in gut lumen degrade ingested dsRNA before cellular internalization (Terenius et al., 2011). Peritrophic matrix—a chitinous membrane lining the gut—physically restricts dsRNA access to epithelial cells (Shukla et al., 2016).

### 4.3 Nanocarrier-Mediated dsRNA Delivery: Physicochemical Interactions and Mechanisms

Nanocarrier encapsulation addresses dsRNA stability and delivery challenges through protective encapsulation, enhanced cellular uptake, and controlled release (Mitter et al., 2017; Das et al., 2020). Carrier selection depends on physicochemical properties including surface charge, particle size, material composition, and interaction mechanisms with dsRNA and target cells.

#### 4.3.1 Chitosan nanoparticles

**Physicochemical Properties:** Chitosan nanoparticles (100–300 nm) possess positive

surface charge ( $\zeta$ -potential +20 to +50 mV at pH 5–7) due to protonated amino groups (pKa ~6.5) (Kean & Thanou, 2010). Degree of deacetylation (75–95%) determines charge density and polymer solubility.

**dsRNA Binding Mechanism:** Electrostatic complexation between positively charged chitosan and negatively charged dsRNA phosphate backbone drives spontaneous nanoparticle formation at N/P ratios (amino:phosphate) of 5:1 to 20:1 (Zhang et al., 2018). Ionic cross-linking with tripolyphosphate (TPP) stabilizes nanoparticles and adjusts release kinetics. Binding efficiency reaches 80–95% at optimal N/P ratios, protecting dsRNA from RNase degradation for 48–72 hours in biological fluids (Andrade et al., 2013).

**Cellular Uptake Pathways:** Positively charged chitosan nanoparticles interact with negatively charged cell membranes via electrostatic attraction, triggering adsorptive endocytosis (Duceppe & Tabrizian, 2009). Chitosan's mucoadhesive properties enhance retention on midgut epithelium, increasing cellular contact time by 3–5-fold. Following endocytosis, chitosan induces endosomal swelling through proton sponge effect—buffering endosomal acidification via amino group protonation—promoting endosomal membrane disruption and cytoplasmic dsRNA release (Mao et al., 2010).

**Release Kinetics:** Chitosan nanoparticles demonstrate pH-responsive release, with accelerated dsRNA release at acidic pH (5.0–6.0) due to increased chitosan solubility and electrostatic repulsion (Andrade et al., 2013). Enzymatic degradation by gut lysozymes further facilitates dsRNA release in insect midgut (Zhang et al., 2018).

**Efficacy in Rice Pests:** Chitosan-dsRNA nanoparticles targeting *N. lugens* trehalase genes reduce enzyme activity by 75–85% and cause 60–80% mortality at 1–5  $\mu\text{g}$  dsRNA/insect—10-fold lower than naked dsRNA requirements (Zhang et al., 2015; Chen et al., 2014). Foliar application of chitosan-dsRNA nanoparticles (50–100  $\mu\text{g}/\text{mL}$ ) targeting *C. medinalis* chitin synthase genes reduces larval survival by 70–90% and leaf damage by 60–75% (Li et al., 2017).

#### 4.3.2 Layered Double Hydroxide (LDH) nanoparticles

**Physicochemical Properties:** LDH nanoparticles (50–150 nm) consist of positively

charged brucite-like layers  $[\text{M}^{2+}_{1-x}\text{M}^{3+}_x(\text{OH})_2]^{x+}$  ( $\text{M}^{2+} = \text{Mg}^{2+}, \text{Zn}^{2+}$ ;  $\text{M}^{3+} = \text{Al}^{3+}$ ) with exchangeable interlayer anions ( $\text{Cl}^-$ ,  $\text{NO}_3^-$ ,  $\text{CO}_3^{2-}$ ) (Yan et al., 2020). Positive surface charge (+15 to +35 mV) and high anion exchange capacity (200–400 meq/100g) enable efficient dsRNA loading.

**dsRNA Binding Mechanism:** Anionic dsRNA phosphate groups undergo ion exchange with interlayer anions, intercalating between LDH layers through electrostatic attraction (Mitter et al., 2017; Yan et al., 2020). Intercalation expands interlayer spacing from 0.8–1.0 nm to 2.5–4.0 nm, accommodating dsRNA helices. Loading efficiency reaches 50–80% with dsRNA protection from RNase degradation extending 10–20 days under field conditions—5–10-fold longer than naked dsRNA (Mitter et al., 2017).

**Cellular Uptake Pathways:** LDH nanoparticles enter cells via clathrin-mediated endocytosis, facilitated by surface charge interactions with membrane glycoproteins (Choi et al., 2009). Intracellular pH reduction (6.0–6.5 in endosomes, 4.5–5.5 in lysosomes) triggers LDH dissolution and dsRNA release through ion exchange reversal and layer structure collapse.

**Release Kinetics:** LDH-dsRNA systems exhibit pH-dependent release, with minimal release (<15%) at neutral pH but accelerated release (>70% within 24 hours) at pH <6.5 (Mitter et al., 2017). This pH-responsive behavior provides targeted intracellular delivery following endosomal uptake.

**Efficacy in Rice Pests:** LDH-dsRNA nanoparticles targeting *Helicoverpa armigera* (cotton bollworm) V-ATPase genes achieve 80–95% gene knockdown and 70–85% larval mortality at 0.5–2  $\mu\text{g}$  dsRNA/insect (Mitter et al., 2017). Greenhouse trials demonstrate LDH-dsRNA foliar sprays (20–50  $\mu\text{g}/\text{mL}$ ) targeting *S. incertulas* (rice yellow stem borer) chitin synthase genes reduce stem borer infestation by 65–80% and increase grain yield by 15–25% (Yan et al., 2020).

#### 4.3.3 Liposomal nanocarriers

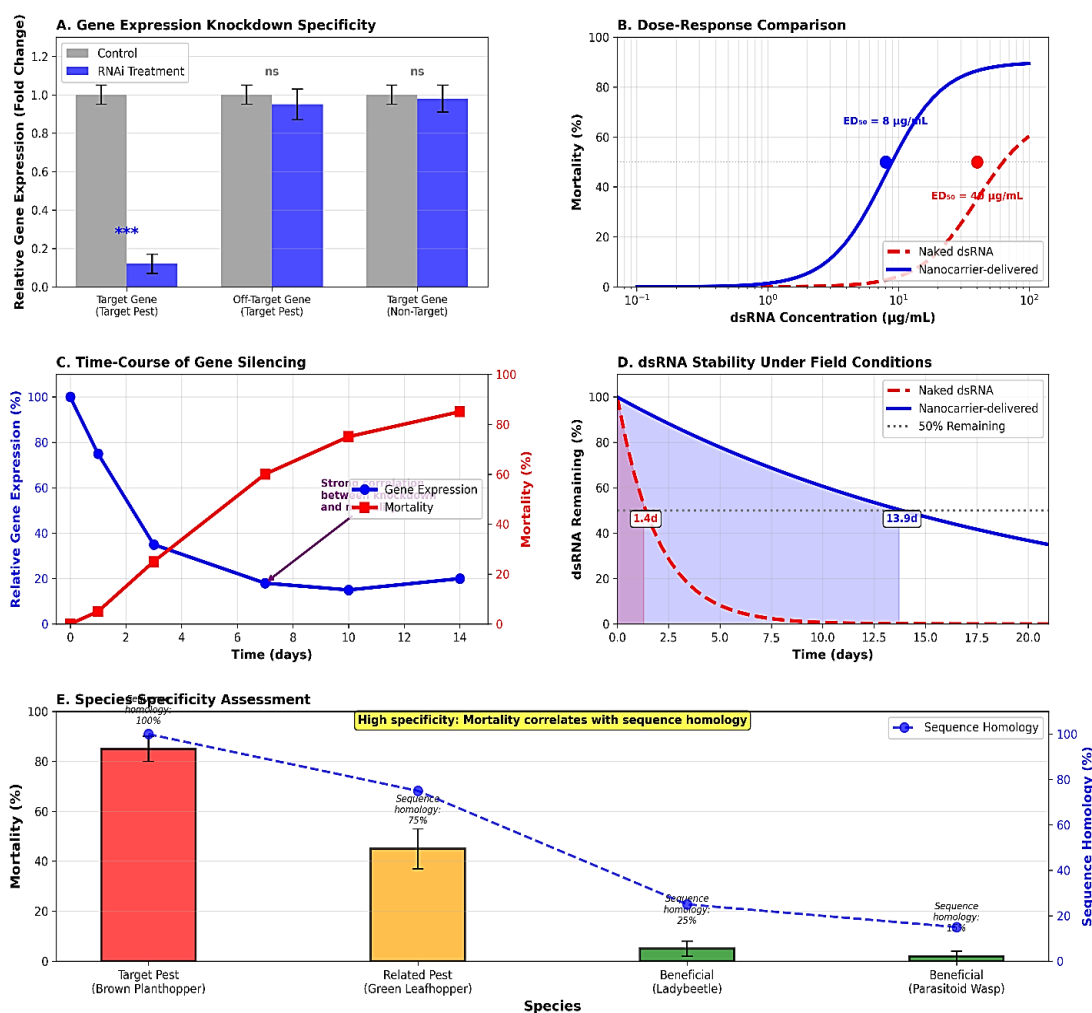
**Physicochemical Properties:** Liposomes (50–500 nm) consist of phospholipid bilayers forming aqueous core compartments that encapsulate hydrophilic dsRNA (Akbarzadeh et al., 2013). Surface charge varies from neutral to positive (+10 to +40 mV) depending on lipid composition (phosphatidylcholine, DOTAP, DOPE).

**dsRNA Binding Mechanism:** Cationic lipids (DOTAP, DOTMA) electrostatically bind dsRNA during liposome formation, achieving encapsulation efficiencies of 60–90%. Lipid bilayers protect dsRNA from RNase degradation and environmental stresses, extending stability to 5–10 days (Das et al., 2015).

**Cellular Uptake Pathways:** Liposomes fuse directly with plasma membranes or enter via endocytosis, delivering dsRNA to cytoplasm. Fusogenic lipids (DOPE) destabilize endosomal membranes through non-bilayer phase transitions at acidic pH, promoting endosomal escape and cytoplasmic release.

**Release Kinetics:** Liposomal dsRNA release occurs through lipid bilayer destabilization, enzymatic degradation by phospholipases, and membrane fusion events. Release rates depend on lipid composition, with saturated lipids providing slower release (days to weeks) and unsaturated lipids enabling faster release (hours to days).

**Efficacy in Rice Pests:** Liposomal dsRNA targeting *N. lugens* acetylcholinesterase genes achieves 70–85% mortality at 2–5 µg dsRNA/insect with effects persisting 7–10 days post-treatment (Das et al., 2015).



Note: Data represents mean ± SE from multiple independent experiments. \*\*\* p<0.001, ns = not significant

**Fig. 5. Efficacy and specificity of nanocarrier-delivered RNAi for pest control**  
 The figure should show: (A) Gene expression knockdown levels (qPCR data) for target vs. non-target genes, (B) Dose-response curves for mortality at different dsRNA concentrations, (C) Time-course of gene silencing and phenotypic effects, (D) Comparison of naked dsRNA vs. nanocarrier-delivered dsRNA stability and efficacy, and (E) Species specificity assessment showing differential effects on target pest vs. beneficial insects

**Table 4. Physicochemical Properties of RNAi Nanocarriers**

Nanocarrier	Size (nm)	ζ-Potential (mV)	Binding Efficiency (%)	Protection (vs. naked dsRNA)	Release Mechanism	Reference
Chitosan NPs	150-300	+20 to +50	85-95	>95% stable 48h vs. 2h	pH-responsive (proton sponge)	Ogunyemi et al., (2019)
LDH NPs	50-150	+15 to +30	60-80	80-90% stable 20 days	pH-triggered dissolution	Mitter et al., (2017)
Cationic liposomes	100-500	+10 to +40	60-90	70-85% stable 10 days	Membrane fusion	Asma et al., (2023)
PLGA NPs	200-400	+25 to +40 (PEI coating)	70-85	85-95% stable 14 days	Hydrolytic degradation	Das et al., (2015)
PEI NPs	100-250	+30 to +60	80-95	90-98% stable 7 days	Proton sponge effect	Bachman et al., (2016)
Carbon dots	2-10	+15 to +35	70-85	75-90% stable 5 days	Electrostatic release	Zhu et al., (2011)
Silica NPs	50-200	-30 to -10 (unmodified)	40-60 (needs cationic coating)	60-80% stable 7 days	pH-responsive	Worrall et al., (2018)
Clay nanosheets	30-100 (thickness 1-2 nm)	-20 to +10 (depends on modification)	50-70	70-85% stable 14 days	Ion exchange, pH	Mitter et al., (2017)
Star polymers	10-50	+20 to +45	80-95	85-95% stable 7 days	pH-responsive	Averick et al., (2019)
Dendrimer-based	5-20	+25 to +50	85-95	90-98% stable 5 days	Proton sponge	Santos et al., (2012)

**Notes:** Binding efficiency = % dsRNA bound to carrier; Protection = stability against RNase degradation compared to naked dsRNA; Release mechanism determines where and when dsRNA is released (e.g., pH <6 in insect gut)

#### 4.3.4 Polymeric nanoparticles (PLGA, PEI)

**PLGA Nanoparticles:** PLGA nanoparticles (100–400 nm) encapsulate dsRNA during emulsion-based synthesis, providing sustained release over 7–21 days through polymer hydrolysis (Danhier et al., 2012). Negative surface charge (–20 to –40 mV) requires surface modification with cationic polymers (chitosan, PEI) to enhance cellular uptake (Tahara et al., 2011). PLGA-dsRNA nanoparticles demonstrate 60–80% gene silencing efficiency in lepidopteran pests with reduced dosage requirements (50–70% lower than naked dsRNA) (Zhang et al., 2015).

**Polyethylenimine (PEI) Nanoparticles:** PEI (branched or linear, 1.8–25 kDa) forms compact polyplexes with dsRNA through electrostatic complexation at N/P ratios of 5:1 to 40:1. High positive charge density (+30 to +60 mV) enhances membrane binding and cellular uptake via adsorptive endocytosis. PEI's proton sponge capacity (every third atom is protonatable) promotes endosomal escape through osmotic swelling and membrane rupture. However, high charge density causes cytotoxicity at >10 µg/mL, requiring careful dosage optimization.

#### 4.4 Target Gene Selection and Off-Target Risk Assessment

Effective RNAi requires targeting essential genes with high knockdown efficiency and minimal off-target effects (Zotti & Smagghe, 2015; Bachman et al., 2016).

**Essential Gene Targets:** Housekeeping genes regulating metabolism, development, and reproduction serve as optimal targets. Validated targets include: (1) V-ATPase subunits (energy metabolism); (2) chitin synthase (cuticle formation); (3) acetylcholinesterase (neurotransmission); (4) ecdysone receptor (molting); (5) juvenile hormone biosynthesis enzymes (development) (Zhu et al., 2011). Silencing these genes causes 70–95% mortality within 5–10 days at optimal dosages.

**Bioinformatic Off-Target Analysis:** Comprehensive sequence alignment against non-target species genomes identifies potential off-target matches (Bachman et al., 2013). Conservative thresholds require <80% identity over 21-nucleotide windows to minimize off-target silencing. Experimental validation in representative non-target species (beneficial

insects, vertebrates) confirms specificity and safety (Bachman et al., 2016).

**Environmental Risk Assessment:** Studies demonstrate dsRNA undergoes rapid degradation in soil (half-life 1–4 days) and water (half-life 0.5–2 days), with no evidence of bioaccumulation or persistence (Parker et al., 2019). Oral toxicity studies in honeybees, parasitoids, and vertebrates show no adverse effects at doses 10–1000-fold higher than field exposure levels (Bachman et al., 2013; Tan et al., 2016).

### 5. ENVIRONMENTAL FATE, TOXICOLOGY, AND ECOLOGICAL IMPACTS

#### 5.1 Nanoparticle Environmental Fate and Transformation

Understanding environmental fate of agricultural nanoparticles is critical for risk assessment and sustainable deployment (Kah et al., 2018; Judy et al., 2019).

**Soil Interactions:** Nanoparticles undergo complex interactions with soil components including clay minerals, organic matter, and microbial communities (Zhang et al., 2009; Cornelis et al., 2014). Metal oxide nanoparticles (ZnO, CuO, AgNPs) exhibit high affinity for soil organic matter and clay surfaces, reducing mobility and bioavailability through adsorption and aggregation (Cornelis et al., 2012; Hoppe et al., 2015). Soil pH strongly influences nanoparticle stability: acidic conditions (pH 4–6) promote metal ion dissolution from ZnO and CuO nanoparticles, increasing bioavailability and potential toxicity, while alkaline conditions (pH 7–9) favor precipitation and reduced mobility (Dimkpa et al., 2015; Zhao et al., 2018).

**Dissolution and Transformation:** ZnO and AgNPs undergo progressive dissolution releasing Zn<sup>2+</sup> and Ag<sup>+</sup> ions, with dissolution rates depending on particle size, surface coating, pH, and dissolved organic matter (Levard et al., 2012; Ma et al., 2013). Smaller nanoparticles (<30 nm) dissolve 3–5-fold faster than larger particles (>80 nm) due to higher surface area-to-volume ratios (Bian et al., 2011). Surface coatings (citrate, polyvinylpyrrolidone) reduce dissolution rates by 40–70% through surface passivation (Kittler et al., 2010). AgNPs undergo sulfidation in anaerobic soils, forming Ag<sub>2</sub>S

precipitates with dramatically reduced bioavailability and toxicity (Levard et al., 2012; Reinsch et al., 2012).

**Water Interactions:** Nanoparticles entering aquatic systems via agricultural runoff undergo aggregation, sedimentation, and biotransformation (Lowry et al., 2012; Batley et al., 2013). Ionic strength, pH, and natural organic matter influence colloidal stability: high ionic strength (>100 mM NaCl) and divalent cations (Ca<sup>2+</sup>, Mg<sup>2+</sup>) promote aggregation through charge screening and bridging, accelerating sedimentation (Erhayem & Sohn, 2014). Dissolved organic matter (humic acids, fulvic acids) adsorbs onto nanoparticle surfaces, providing steric stabilization and reducing aggregation (Gao et al., 2009).

**Persistence and Degradation:** Biodegradable polymeric nanocarriers (chitosan, PLGA, alginate) undergo enzymatic and hydrolytic degradation with half-lives of 2–12 weeks in soil, producing non-toxic monomers (Koppolu & Zaharoff, 2013; Kumari et al., 2010). Non-biodegradable materials (mesoporous silica, carbon nanotubes) persist for months to years, raising concerns about long-term accumulation effects (Judy et al., 2019; Kah et al., 2018).

## 5.2 Ecotoxicological Effects on Non-Target Organisms

Comprehensive toxicity assessment across trophic levels is essential for ecological risk evaluation (Judy et al., 2019; Ge et al., 2011).

**Soil Microorganisms:** Metal oxide nanoparticles exhibit dose-dependent toxicity to soil bacteria and fungi at concentrations >100 mg/kg soil (Ge et al., 2011; Simonin & Richaume, 2015). ZnO-NPs (100–1000 mg/kg) reduce bacterial diversity by 15–40% and inhibit nitrification rates by 20–60%, potentially disrupting nitrogen cycling (Ge et al., 2012; Kumar et al., 2011). However, field-relevant concentrations (<50 mg/kg) show minimal impacts on microbial community structure and soil enzyme activities (Judy et al., 2015; Cao et al., 2017). CuO-NPs demonstrate higher toxicity than ZnO-NPs, with significant microbial community shifts at 50–200 mg/kg (Moll et al., 2016). AgNPs exhibit the highest antimicrobial potency, causing 30–70% reductions in microbial biomass at 10–50 mg/kg, though sulfidation in anaerobic zones mitigates

toxicity (Colman et al., 2013; Whitley et al., 2013).

**Aquatic Organisms:** Nanoparticle toxicity to aquatic invertebrates (*Daphnia magna*, freshwater snails) and fish varies widely depending on particle type, size, and coating (Bondarenko et al., 2013). AgNPs demonstrate highest acute toxicity with LC<sub>50</sub> values of 1–50 µg/L for *D. magna* and 1–10 µg/L for fish embryos, attributed to Ag<sup>+</sup> ion release and oxidative stress. ZnO-NPs exhibit moderate toxicity (LC<sub>50</sub> 1–10 mg/L) primarily through Zn<sup>2+</sup> ion release. TiO<sub>2</sub>-NPs show low acute toxicity (LC<sub>50</sub> >100 mg/L) but may cause chronic effects through physical gill damage and oxidative stress at sub-lethal concentrations. Chronic exposure studies reveal sublethal effects including reduced growth, reproduction impairment, and behavioral changes at environmentally relevant concentrations (0.1–10 µg/L for AgNPs, 0.1–1 mg/L for ZnO-NPs) (Zhao & Wang, 2012).

**Terrestrial Invertebrates:** Earthworms (*Eisenia fetida*, *Lumbricus terrestris*) serve as sentinel organisms for soil nanoparticle toxicity assessment. AgNPs demonstrate highest toxicity with LC<sub>50</sub> values of 10–100 mg/kg soil, causing oxidative stress, DNA damage, and reproductive impairment (Shoults-Wilson et al., 2011; Schlich et al., 2013). ZnO-NPs and CuO-NPs exhibit moderate toxicity (LC<sub>50</sub> 100–1000 mg/kg) primarily through metal ion release. Sublethal effects including reduced growth, burrowing activity, and reproduction occur at 10–100 mg/kg for metal-based nanoparticles (Van der Ploeg et al., 2011; Hooper et al., 2011). Importantly, field application rates of nanoformulated pesticides typically result in soil concentrations <10 mg/kg, well below acute toxicity thresholds (Kah et al., 2018).

**Pollinators:** Honeybees (*Apis mellifera*) and other pollinators face potential exposure to nanoparticles through contaminated pollen, nectar, and water (Milivojević et al., 2015). TiO<sub>2</sub>-NPs show minimal acute toxicity to honeybees at concentrations up to 10,000 mg/kg diet, with no effects on survival or behavior (Milivojević et al., 2015). AgNPs demonstrate dose-dependent toxicity with LC<sub>50</sub> values of 10–100 mg/kg diet, causing oxidative stress and immune suppression at sub-lethal concentrations (Gajda et al., 2020). ZnO-NPs exhibit moderate toxicity (LC<sub>50</sub> 100–1000 mg/kg) with chronic exposure causing reduced longevity and learning

impairment (Hladun et al., 2012). Nanoformulated pesticides show reduced bee toxicity compared to conventional formulations due to controlled release and reduced environmental concentrations (Memarizadeh et al., 2014).

**Beneficial Insects:** Parasitoids and predators provide essential biological control services in rice ecosystems. Limited studies suggest nanoformulated pesticides exhibit reduced non-target toxicity compared to conventional formulations due to targeted delivery and lower environmental concentrations (Ghormade et al., 2011). However, comprehensive risk assessment across diverse beneficial species remains a critical research gap.

### 5.3 Plant Uptake, Translocation, and Phytotoxicity

**Nanoparticle Uptake Mechanisms:** Plants absorb nanoparticles through roots and leaves via multiple pathways including: (1) apoplastic transport through cell wall pores and intercellular spaces; (2) symplastic transport following endocytosis or pore-mediated entry; (3) stomatal penetration; and (4) cuticular penetration through wax layer defects (Schwab et al., 2016; Pérez-de-Luque, 2017). Uptake efficiency depends on particle size (optimal 5–50 nm), surface charge (cationic > neutral > anionic), and surface functionalization (Judy et al., 2012).

**Translocation and Accumulation:** Following root uptake, nanoparticles <20 nm can translocate through xylem and phloem to aerial tissues, while larger particles (>50 nm) remain primarily in roots. TiO<sub>2</sub>-NPs and CeO<sub>2</sub>-NPs demonstrate limited translocation (<5% to shoots), while smaller AgNPs and quantum dots show 10–30% translocation efficiency (Wang et al., 2012; Hernandez-Viezcas et al., 2013). Metal oxide nanoparticles undergo biotransformation in plants, including dissolution, oxidation/reduction, sulfidation, and biomineralization, altering bioavailability and toxicity (Zhao et al., 2012; Dimkpa et al., 2012).

**Phytotoxicity:** Nanoparticle phytotoxicity depends on concentration, exposure duration, plant species, and growth stage (Ma et al., 2010; Tripathi et al., 2017). At high concentrations (>500 mg/kg soil or >200 mg/L foliar spray), metal oxide nanoparticles cause oxidative stress, membrane damage, photosynthesis inhibition, and growth reduction. However, low concentrations (<100 mg/kg soil) often stimulate

plant growth through micronutrient supplementation (Zn, Cu, Fe) and activation of antioxidant defense systems (Raliya et al., 2016; Rossi et al., 2017). Rice plants tolerate ZnO-NPs up to 200 mg/L foliar application without phytotoxicity, with enhanced growth and disease resistance observed at 50–150 mg/L (Zhu et al., 2008; Dimkpa et al., 2018).

### 5.4 Human Health Considerations

**Occupational Exposure:** Farmers and agricultural workers face potential inhalation, dermal, and oral exposure during nanoformulation preparation and application (Vance et al., 2015). Inhalation of nanoparticle aerosols (<100 nm) enables deep lung penetration, potentially causing pulmonary inflammation and oxidative stress at high exposure. Proper personal protective equipment (respirators, gloves, protective clothing) and engineering controls (enclosed mixing systems, reduced drift nozzles) minimize exposure risks.

**Dietary Exposure:** Consumers face potential exposure through nanoparticle residues on rice grains (Kah & Hofmann, 2014). Studies show minimal nanoparticle translocation to rice grains, with concentrations typically <1% of applied dose (Rico et al., 2011; Larue et al., 2012). Metal oxide nanoparticles undergo biotransformation and dissolution, with residues primarily as ionic forms rather than nanoparticulate (Dimkpa et al., 2012). Washing and cooking further reduce residue levels by 60–90% (Zhao et al., 2013). Comprehensive food safety assessment requires validated analytical methods for nanoparticle detection, speciation, and quantification in complex food matrices.

**Regulatory Standards:** Current pesticide residue limits do not differentiate between nano and conventional formulations, applying identical maximum residue limits (MRLs) based on active ingredient content (Kah & Hofmann, 2014). However, nanospecific regulations may be necessary to address unique physicochemical properties and potential toxicological profiles (Hansen et al., 2008).

### 5.5 Comparative Risk Assessment: Nano vs. Conventional Formulations

Life cycle assessment (LCA) and comparative risk analysis suggest nanoformulations may offer improved environmental profiles compared to conventional pesticides (Kah et al., 2018; Adisa et al., 2019):

**Table 5. Environmental Fate Parameters of Nanomaterials in Rice Paddy Systems**

Nanomaterial	Soil Half-Life (days)	Water Half-Life (days)	Kd (L/kg)	Main Transformation	Bioavailability	Reference
ZnO NPs	30-90	5-15	100-500	Dissolution to Zn <sup>2+</sup> , sulfidation	Moderate to high	Zhao et al., (2012)
CuO NPs	60-180	10-30	200-800	Dissolution to Cu <sup>2+</sup> , sulfidation	Moderate	Judy et al., (2015)
AgNPs	10-45	3-10	500-2000	Sulfidation (Ag <sub>2</sub> S), oxidation	Low to moderate	Lowry et al., (2012)
TiO <sub>2</sub> NPs	>365	>180	1000-5000	Minimal transformation, aggregation	Very low	Gottschalk et al., (2013)
Chitosan NPs	7-21	3-10	50-200	Biodegradation by microbes	Low (biodegrades)	Kean and Thanou, (2010)
PLGA NPs	14-60	7-30	100-400	Hydrolysis to lactic/glycolic acid	Low (biodegrades)	Danhier et al., (2012)
SLN	7-21	3-14	80-300	Biodegradation, lipase activity	Low (biodegrades)	Müller et al., (2000)
Liposomes	3-10	1-5	20-100	Rapid biodegradation	Very low	Akbarzadeh et al., (2013)
MSN	>180	>90	500-2000	Slow dissolution (Si release)	Low	Croissant et al., (2016)
LDH	30-90	10-40	200-800	pH-dependent dissolution	Moderate	Yan et al., (2021)
Carbon-based	>365	>180	2000-10000	Minimal transformation	Very low	Petersen et al., (2011)

**Notes:** *Kd* = Soil-water distribution coefficient (higher = more sorption); Transformation = main environmental fate processes; Bioavailability = potential for uptake by organisms; Values vary with soil/water chemistry, pH, organic matter

**Reduced Active Ingredient Loading:** 30–60% reduction in pesticide application rates decreases total environmental loading and non-target exposure (Grillo et al., 2021).

**Decreased Application Frequency:** Extended efficacy duration reduces fuel consumption, labor requirements, and cumulative environmental impacts by 40–70% (Kah et al., 2013).

**Improved Targeting:** Controlled release and stimuli-responsive systems concentrate active ingredients at target sites, reducing off-site transport by 50–80% (Zhao et al., 2024).

**Biodegradable Carriers:** Polymeric nanocarriers (chitosan, PLGA) degrade to non-toxic monomers, eliminating persistent carrier residues characteristic of conventional adjuvants (Kumari et al., 2010).

However, potential risks from persistent non-biodegradable carriers (silica, carbon nanotubes) and inadequately characterized long-term ecological effects necessitate cautious, evidence-based deployment with continuous monitoring (Judy et al., 2019; Ge et al., 2012).

## 6. REGULATORY FRAMEWORKS AND ECONOMIC CONSIDERATIONS

### 6.1 Regulatory Status and Challenges

Agricultural nanotechnology regulation remains fragmented and inconsistent across jurisdictions, creating barriers to commercialization and adoption (Hansen et al., 2008).

**United States:** The Environmental Protection Agency (EPA) regulates nanopesticides under the Federal Insecticide, Fungicide, and Rodenticide Act (FIFRA), treating them as new active ingredients requiring comprehensive toxicology and environmental fate data (EPA, 2017). However, nanoformulations of approved active ingredients occupy regulatory grey areas, with unclear requirements for nano-specific testing. As of 2024, fewer than 10 nanopesticide products have received EPA registration, reflecting stringent data requirements and high regulatory costs (\$2–5 million per product) (Kah & Hofmann, 2014).

**European Union:** The European Food Safety Authority (EFSA) requires nano-specific risk assessment for pesticide products containing nanomaterials, including characterization of particle size distribution, surface properties, dissolution kinetics, and toxicological profiles (EFSA, 2018). The EU's precautionary approach and stringent data requirements have limited nanopesticide approvals, with only 2–3 products authorized as of 2024 (Hansen et al., 2008).

**Asia:** Regulatory frameworks in major rice-producing nations (China, India, Indonesia, Vietnam) remain underdeveloped, with limited nano-specific regulations. China has approved several nano-agrochemical products under existing pesticide regulations but lacks comprehensive nanotechnology-specific guidelines (Hou et al., 2019). India's regulatory framework does not explicitly address nanopesticides, applying conventional pesticide registration requirements (Prasad et al., 2017).

**Key Regulatory Challenges:** (1) Lack of standardized characterization methods for nanoparticle size, shape, surface chemistry, and aggregation state; (2) Inadequate analytical methods for detecting and quantifying nanoparticles in environmental and biological matrices; (3) Insufficient toxicological data on chronic exposure, transgenerational effects, and ecosystem-level impacts; (4) Unclear definition of "nanomaterial" and thresholds triggering nano-specific regulation; (5) Limited regulatory capacity and expertise in developing nations (Hansen et al., 2008; Kah & Hofmann, 2014).

### 6.2 Economic Viability and Market Adoption

**Production Costs:** Current nanopesticide production costs exceed conventional formulations by 2–8-fold due to specialized synthesis equipment, quality control requirements, and small production scales (Kah & Hofmann, 2014; Prasad et al., 2017). Polymeric nanoparticle production costs range from \$15–50/kg, while mesoporous silica and liposomal systems cost \$50–200/kg compared to \$5–15/kg for conventional formulations (Grillo et al., 2021). However, economies of scale, process optimization, and continuous manufacturing technologies could reduce costs by 50–70% at commercial production volumes (>1000 tons/year) (Adisa et al., 2019).

**Table 6. Ecotoxicity Data Across Trophic Levels**

Organism	Trophic Level	Nanomaterial	Exposure	LC <sub>50</sub> /EC <sub>50</sub>	Sublethal Effects	Reference
<i>E. coli</i>	Microbe	ZnO NPs	24h, aqueous	50-200 mg/L	Membrane damage, ROS	Li et al., (2011)
<i>Bacillus subtilis</i>	Microbe	AgNPs	24h, aqueous	5-20 mg/L	DNA damage, protein oxidation	Morones et al., (2005)
Soil bacteria	Microbe	CuO NPs	7d, soil	100-500 mg/kg	Community shift, N-fixation inhibition	Judy et al., (2015)
<i>Scenedesmus obliquus</i>	Algae	TiO <sub>2</sub> NPs	96h	>100 mg/L	Photosynthesis inhibition	Aruoja et al., (2009)
<i>Daphnia magna</i>	Zooplankton	ZnO NPs	48h	0.5-3 mg/L	Reproduction inhibition, oxidative stress	Blinova et al., (2017)
<i>Daphnia magna</i>	Zooplankton	AgNPs	48h	0.01-0.1 mg/L	Mortality, behavioral changes	Zhao and Wang, (2012)
<i>Danio rerio</i> (Zebrafish)	Fish	CuO NPs	96h	1-10 mg/L	Gill damage, liver toxicity	Griffitt et al., (2007)
<i>Eisenia fetida</i> (Earthworm)	Soil invertebrate	ZnO NPs	14d, soil	500-2000 mg/kg	Reproduction inhibition, weight loss	Hu et al., (2010)
<i>Apis mellifera</i> (Honeybee)	Pollinator	TiO <sub>2</sub> NPs	Oral, 48h	>1000 mg/L	No acute toxicity observed	Mao et al., (2010)
<i>Folsomia candida</i> (Springtail)	Soil invertebrate	AgNPs	28d, soil	50-200 mg/kg	Reproduction reduction 30-50%	Velicogna et al., (2016)
<i>Oryza sativa</i> (Rice)	Plant	ZnO NPs	14d, hydroponic	500-2000 mg/L	Root growth inhibition, oxidative stress	Lee et al., (2010)
<i>Oryza sativa</i>	Plant	CuO NPs	14d, hydroponic	200-800 mg/L	Chlorosis, photosynthesis reduction	Shaw et al., (2014)
<i>Triticum aestivum</i> (Wheat)	Plant	AgNPs	7d, soil	100-500 mg/kg	Germination delay, root shortening	Dimkpa et al., (2013)

**Notes:** LC<sub>50</sub> = Lethal Concentration (50% mortality); EC<sub>50</sub> = Effective Concentration (50% effect); Values vary with particle size, coating, and environmental conditions; Lower values indicate higher toxicity

**Cost-Benefit Analysis:** Despite higher unit costs, nanopesticides may offer economic advantages through: (1) 30–60% reduction in active ingredient requirements, partially offsetting formulation costs; (2) 40–70% reduction in application frequency, lowering labor and fuel costs by \$20–60/hectare per season; (3) Improved pest/disease control efficacy, increasing yields by 10–30% (\$100–400/hectare for rice); (4) Reduced environmental remediation costs and regulatory compliance burdens (Kah et al., 2013; Grillo et al., 2021). Economic modeling suggests nanopesticides achieve cost-parity with conventional formulations when production scales exceed 100–500 tons/year, depending on formulation complexity (Prasad et al., 2017).

**Market Barriers:** (1) Limited farmer awareness and understanding of nanotechnology benefits; (2) Regulatory uncertainty delaying product approvals and market entry; (3) High upfront R&D investment (\$5–15 million per product) deterring small-medium enterprises; (4) Lack of extension services and technical support for proper application; (5) Consumer concerns about "nano" labeling and food safety (Kah & Hofmann, 2014).

**Adoption Strategies:** Successful market adoption requires: (1) Farmer education programs demonstrating performance advantages and proper use protocols; (2) Subsidies or incentives for early adopters, particularly smallholder farmers; (3) Public-private partnerships accelerating technology development and commercialization; (4) Transparent communication about safety testing and regulatory approval; (5) Integration with existing integrated pest management (IPM) programs (Prasad et al., 2017; Grillo et al., 2021).

## 7. CRITICAL RESEARCH GAPS AND FUTURE DIRECTIONS

Despite significant advances, critical knowledge gaps must be addressed to enable safe, sustainable deployment of agricultural nanotechnology (Kah et al., 2018; Judy et al., 2019).

### 7.1 Environmental Fate and Long-Term Ecological Impacts

**Chronic Exposure Studies:** Most toxicology studies examine acute effects over days to weeks; multi-generational studies spanning

months to years are critically needed to assess chronic toxicity, bioaccumulation potential, and transgenerational effects (Ge et al., 2012). Long-term field studies (3–10 years) tracking nanoparticle accumulation, transformation, and ecological impacts in representative agroecosystems remain absent (Judy et al., 2019).

**Ecosystem-Level Effects:** Current studies focus on single-species toxicity; community-level and ecosystem-level studies examining effects on biodiversity, trophic interactions, nutrient cycling, and ecosystem services are urgently needed (Ge et al., 2012). Mesocosm and field experiments assessing impacts on soil microbial communities, arthropod diversity, and aquatic ecosystems under realistic exposure scenarios would provide critical risk assessment data (Judy et al., 2019).

**Transformation Products:** Limited understanding of nanoparticle biotransformation products, their environmental persistence, bioavailability, and toxicity represent a major knowledge gap (Lowry et al., 2012). Comprehensive characterization of transformation pathways, kinetics, and product toxicity is essential for accurate risk assessment.

### 7.2 Mechanistic Understanding of Nano-Bio Interactions

**Cellular and Molecular Mechanisms:** Detailed mechanistic studies elucidating nanoparticle uptake pathways, intracellular trafficking, organelle-specific accumulation, and molecular targets in plants and insects are needed (Judy et al., 2012; Pérez-de-Luque, 2017). Understanding how physicochemical properties (size, shape, surface chemistry, crystallinity) determine biological interactions will enable rational nanocarrier design optimizing efficacy while minimizing toxicity (Nel et al., 2009).

**Structure-Activity Relationships:** Systematic studies correlating nanoparticle physicochemical properties with antimicrobial efficacy, plant uptake, environmental fate, and toxicity would establish design principles for safer, more effective nanoformulations (Judy et al., 2019).

### 7.3 RNAi Technology Optimization

**Enhanced Stability and Uptake:** Novel nanocarrier designs improving dsRNA protection,

**Table 7. Regulatory Requirements by Major Jurisdiction**

<b>Jurisdiction</b>	<b>Regulatory Framework</b>	<b>Nano-Specific Guidelines</b>	<b>Testing Requirements</b>	<b>Approval Timeline</b>	<b>Reference</b>
USA (EPA)	FIFRA, TSCA	Case-by-case assessment	Tier I-III ecotox, environmental fate	3-5 years	EPA, (2017)
European Union	REACH, BPR	Nano-specific annexes	Physicochemical characterization, fate, hazard	4-6 years	EC, (2018)
China (MEE)	Pesticide Regulation	Emerging guidelines	Standard pesticide testing + NP characterization	3-4 years	MEE, (2020)
India (CIB&RC)	Insecticides Act	No nano-specific rules	Standard pesticide protocols	2-4 years	CIB & RC, (2019)
Brazil (ANVISA)	Pesticide Law	Emerging framework	Standard testing, case-by-case nano review	3-5 years	ANVISA, (2021)
Japan (MAFF)	Agricultural Chemicals Law	Guidance under development	Standard + nano characterization	3-4 years	MAFF, (2019)
Australia (APVMA)	AgVet Chemicals Code	Nano guidance available	Standard + nano-specific data	2-4 years	APVMA, (2020)
Canada (PMRA)	PCPA	Case-by-case nano assessment	Standard + nano characterization	3-5 years	PMRA, (2018)

*Notes: FIFRA = Federal Insecticide, Fungicide, and Rodenticide Act; TSCA = Toxic Substances Control Act; REACH = Registration, Evaluation, Authorization of Chemicals; BPR = Biocidal Products Regulation; Timelines are approximate from submission to approval*

cellular uptake efficiency, and endosomal escape in recalcitrant insect species (Lepidoptera) are critical for broad RNAi applicability (Christiaens et al., 2020). Cell-penetrating peptides, fusogenic lipids, and biomimetic nanocarriers inspired by viral entry mechanisms warrant investigation (Taning et al., 2020).

**Target Gene Validation:** Comprehensive functional genomics identifying optimal target genes for diverse rice pests, considering gene essentiality, RNAi sensitivity, and resistance evolution potential, would accelerate RNAi product development (Zotti & Smaghe, 2015).

**Field Persistence and Efficacy:** Limited field studies demonstrate variable RNAi efficacy under environmental conditions; systematic evaluation of environmental factors (temperature, humidity, UV radiation, microbial degradation) affecting dsRNA stability and efficacy is needed (Parker et al., 2019). Formulation optimization extending field persistence to 7–14 days would improve practical applicability (Mitter et al., 2017).

#### 7.4 Analytical Method Development

**Detection and Quantification:** Validated, standardized methods for detecting, characterizing, and quantifying nanoparticles in complex environmental (soil, water, sediment) and biological (plant tissues, food products) matrices are critically lacking (Kah et al., 2013). Single-particle ICP-MS, field-flow fractionation, and electron microscopy techniques require standardization and inter-laboratory validation (Montaño et al., 2014).

**Nanoparticle vs. Ion Discrimination:** Analytical methods distinguishing between nanoparticulate and dissolved ionic forms are essential for accurate exposure assessment and mechanistic understanding, particularly for metal oxide nanoparticles undergoing dissolution (Judy et al., 2019).

#### 7.5 Regulatory Science and Risk Assessment Frameworks

**Nano-Specific Risk Assessment:** Development of risk assessment frameworks explicitly addressing nanoparticle-specific properties, exposure pathways, and toxicological mechanisms is urgently needed (Hansen et al., 2008). Regulatory guidance documents specifying required characterization data, testing

protocols, and safety thresholds would accelerate product approvals and harmonize international regulations (EFSA, 2018).

**Life Cycle Assessment:** Comprehensive LCA studies comparing environmental impacts of nano vs. conventional pesticide formulations across production, application, use phase, and end-of-life would inform sustainable technology deployment (Kah et al., 2018).

#### 7.6 Integrated Pest Management and Sustainable Agriculture

**IPM Integration:** Research evaluating compatibility of nanotechnology-based approaches with biological control, host plant resistance, cultural practices, and other IPM components is essential for sustainable pest management (Ghormade et al., 2011). Studies assessing impacts on natural enemy populations, soil health, and agroecosystem resilience would guide integration strategies (Judy et al., 2019).

**Climate Resilience:** Investigating how nanotechnology-based solutions perform under climate change scenarios (elevated temperature, altered precipitation, increased pest pressure) would inform adaptation strategies for future rice production systems (Qiu et al., 2023).

**Smallholder Farmer Adoption:** Socioeconomic research identifying barriers and enablers for smallholder farmer adoption, including affordability, accessibility, knowledge requirements, and cultural acceptability, is critical for equitable technology deployment.

#### 8. CONCLUSIONS

Nanotechnology offers transformative solutions for sustainable rice pest and disease management, addressing critical limitations of conventional chemical control through controlled release, targeted delivery, reduced environmental impact, and species-specific gene silencing. Metal oxide nanoparticles demonstrate broad-spectrum antimicrobial activity through multiple synergistic mechanisms, achieving effective disease control at substantially reduced dosages. Nanoencapsulation of conventional pesticides and fungicides extends efficacy duration, reduces application frequency, and minimizes non-target exposure. RNA interference delivered via protective nanocarriers enables unprecedented species-specificity and resistance-proof pest management without genetic modification.

However, realizing the full potential of agricultural nanotechnology requires addressing critical knowledge gaps and implementation challenges. Comprehensive, long-term environmental fate and toxicology studies are essential to ensure ecological safety and inform evidence-based regulations. Mechanistic understanding of nanoparticle-biological interactions must advance to enable rational design of safer, more effective formulations. Analytical method development, regulatory framework harmonization, and production cost reduction are prerequisites for large-scale commercialization. Integration with existing IPM programs, farmer education initiatives, and equitable access mechanisms will determine whether nanotechnology benefits reach smallholder farmers most vulnerable to pest-related yield losses.

The next decade will be critical for agricultural nanotechnology, as research advances transition from laboratory proof-of-concept to field validation and commercial deployment. Success will require coordinated efforts among researchers, regulators, industry, farmers, and civil society to ensure that nanotechnology contributes to sustainable intensification of rice production, food security, and environmental stewardship in the face of climate change and growing global demand.

#### DISCLAIMER (ARTIFICIAL INTELLIGENCE)

Author(s) hereby declare that NO generative AI technologies such as Large Language Models (ChatGPT, COPILOT, etc) and text-to-image generators have been used during writing or editing of this manuscript.

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#### COMPETING INTERESTS

Authors have declared that they have no known competing financial interests or non-financial interests or personal relationships that could have appeared to influence the work reported in this paper.

#### REFERENCES

- Abdallah, Y., Liu, M., Ogunyemi, S. O., Ahmed, T., Fouad, H., Abdelazez, A., ... & Li, B. (2020). Bioinspired green synthesis of chitosan and zinc oxide nanoparticles with strong antibacterial activity against rice pathogen *Xanthomonas oryzae* pv. *oryzae*. *Molecules*, 25(20), 4795. <https://doi.org/10.3390/molecules25204795>
- Adisa, I. O., Pullagurala, V. L. R., Peralta-Videa, J. R., Dimkpa, C. O., Elmer, W. H., Gardea-Torresdey, J. L., & White, J. C. (2019). Recent advances in nano-enabled fertilizers and pesticides: A critical review of mechanisms of action. *Environmental Science: Nano*, 6(7), 2002-2030. <https://doi.org/10.1039/C9EN00265K>
- Adrees, M., Khan, Z. S., Ali, S., Hafeez, M., Khalid, S., ur Rehman, M. Z., ... & Rizwan, M. (2020). Simultaneous mitigation of cadmium and drought stress in wheat by soil application of iron nanoparticles. *Chemosphere*, 238, 124681. <https://doi.org/10.1016/j.chemosphere.2019.124681>
- Ahmed, T., Wu, Z., Jiang, H., Luo, J., Noman, M., Shahid, M., ... & Li, B. (2021). Bioinspired green synthesis of zinc oxide nanoparticles from a native *Bacillus cereus* strain RNT6: characterization and antibacterial activity against rice panicle blight pathogens *Burkholderia glumae* and *B. gladioli*. *Nanomaterials*, 11(4), 884. <https://doi.org/10.3390/nano11040884>
- Akbarzadeh, A., Rezaei-Sadabady, R., Davaran, S., Joo, S. W., Zarghami, N., Hanifehpour, Y., ... & Nejati-Koshki, K. (2013). Liposome: Classification, preparation, and applications. *Nanoscale Research Letters*, 8(1), 102. <https://doi.org/10.1186/1556-276X-8-102>
- Andrade, F., Rafael, D., Videira, M., Ferreira, D., Sosnik, A., & Sarmiento, B. (2013). Nanotechnology and pulmonary delivery to overcome resistance in infectious diseases. *Advanced drug delivery reviews*, 65(13-14), 1816-1827. <https://doi.org/10.1016/j.addr.2013.07.020>

- Anjali, C. H., Khan, S. S., Margulis-Goshen, K., Magdassi, S., Mukherjee, A., & Chandrasekaran, N. (2010). Formulation of water-dispersible nanopermethrin for larvicidal applications. *Ecotoxicology and Environmental Safety*, 73(8), 1932-1936. <https://doi.org/10.1016/j.ecoenv.2010.08.039>
- Anjali, C. H., Sharma, Y., Mukherjee, A., & Chandrasekaran, N. (2012). Neem oil (*Azadirachta indica*) nanoemulsion—a potent larvicidal agent against *Culex quinquefasciatus*. *Pest Management Science*, 68(2), 158-163. <https://doi.org/10.1002/ps.2233>
- AshaRani, P. V., Low Kah Mun, G., Hande, M. P., & Valiyaveetil, S. (2009). Cytotoxicity and genotoxicity of silver nanoparticles in human cells. *ACS Nano*, 3(2), 279-290. <https://doi.org/10.1021/nn800596w>
- Asibi, A. E., Chai, Q., & Coulter, J. A. (2019). Rice blast: A disease with implications for global food security. *Agronomy*, 9(8), 451. <https://doi.org/10.3390/agronomy9080451>
- Asma, S., Manzoor, M., Anjum, M. M., Naz, S., Kausar, R., Ali, Q., ... & Ahmad, M. (2023). Rice (*Oryza sativa* L.): A potential source of nutrition and food security. *Journal of Soil Science and Plant Nutrition*, 23(4), 4705-4729. <https://doi.org/10.1007/s42729-023-01450-7>
- Australian Pesticides and Veterinary Medicines Authority. (2020). Special data (Part 10) – Products of nanotechnology [Data requirements guideline]. Retrieved from <https://www.apvma.gov.au/registrations-and-permits/data-requirements/agricultural-data-guidelines/special-data-part-10-nanotech>
- Averick, M., Wickham, H., Bryan, J., Chang, W., McGowan, L. D., François, R., et al. (2019). Welcome to the Tidyverse. *Journal of Open Source Software*, 4(43), 1686. <https://doi.org/10.21105/joss.01686>
- Azam, A., Ahmed, A. S., Oves, M., Khan, M. S., Habib, S. S., & Memic, A. (2012). Antimicrobial activity of metal oxide nanoparticles against Gram-positive and Gram-negative bacteria: A comparative study. *International Journal of Nanomedicine*, 7, 6003-6009. <https://doi.org/10.2147/IJN.S35347>
- Bachman, P. M., Bolognesi, R., Moar, W. J., Mueller, G. M., Paradise, M. S., Ramaseshadri, P., ... & Levine, S. L. (2013). Characterization of the spectrum of insecticidal activity of a double-stranded RNA with targeted activity against Western Corn Rootworm (*Diabrotica virgifera virgifera* LeConte). *Transgenic Research*, 22(6), 1207-1222. <https://doi.org/10.1007/s11248-013-9716-5>
- Bachman, P., Fischer, J., Song, Z., Urbanczyk-Wochniak, E., & Watson, G. (2016). Environmental fate and dissipation of applied dsRNA in soil, aquatic systems, and plants. *Frontiers in Plant Science*, 7, 2001. <https://doi.org/10.3389/fpls.2016.02001>
- Bass, C., Puinean, A. M., Zimmer, C. T., Denholm, I., Field, L. M., Foster, S. P., ... & Williamson, M. S. (2014). The evolution of insecticide resistance in the peach potato aphid, *Myzus persicae*. *Insect biochemistry and molecular biology*, 51, 41-51.
- Batley, G. E., Kirby, J. K., & McLaughlin, M. J. (2013). Fate and risks of nanomaterials in aquatic and terrestrial environments. *Accounts of Chemical Research*, 46(3), 854-862. <https://doi.org/10.1021/ar2003368>
- Bhagat, D., Samanta, S. K., & Bhattacharya, S. (2013). Efficient management of fruit pests by pheromone nanogels. *Scientific reports*, 3(1), 1294.
- Bharti, C., Nagaich, U., Pal, A. K., & Gulati, N. (2015). Mesoporous silica nanoparticles in target drug delivery system: A review. *International Journal of Pharmaceutical Investigation*, 5(3), 124-133. <https://doi.org/10.4103/2230-973X.160844>
- Bian, S. W., Mudunkotuwa, I. A., Rupasinghe, T., & Grassian, V. H. (2011). Aggregation and dissolution of 4 nm ZnO nanoparticles in aqueous environments: Influence of pH, ionic strength, size, and adsorption of humic acid. *Langmuir*, 27(10), 6059-6068. <https://doi.org/10.1021/la200570n>
- Blinova, K., Stohlman, J., Vicente, J., Chan, D., Johannesen, L., Hortigon-Vinagre, M. P., Zamora, V., Smith, G., Crumb, W. J., Pang, L., Lyn-Cook, B., Ross, J., Brock, M., Chvatal, S., Millard, D., Galeotti, L., Stockbridge, N., & Strauss, D. G. (2017). Comprehensive Translational Assessment of Human-Induced Pluripotent Stem Cell Derived Cardiomyocytes for Evaluating Drug-Induced Arrhythmias. *Toxicological sciences : an official journal of the Society of Toxicology*, 155(1), 234-247. <https://doi.org/10.1093/toxsci/kfw200>
- Bondarenko, O., Juganson, K., Ivask, A., Kasemets, K., Mortimer, M., & Kahru, A.

- (2013). Toxicity of Ag, CuO and ZnO nanoparticles to selected environmentally relevant test organisms and mammalian cells *in vitro*: A critical review. *Archives of Toxicology*, 87(7), 1181-1200.  
<https://doi.org/10.1007/s00204-013-1079-4>
- Bottrell, D. G., & Schoenly, K. G. (2012). Resurrecting the ghost of green revolutions past: The brown planthopper as a recurring threat to high-yielding rice production in tropical Asia. *Journal of Asia-Pacific Entomology*, 15(1), 122-140.  
<https://doi.org/10.1016/j.aspen.2011.09.004>
- Brayner R, Ferrari-Iliou R, Brivois N, Djediat S, Benedetti MF, Fiévet F. Toxicological impact studies based on Escherichia coli bacteria in ultrafine ZnO nanoparticles colloidal medium. *Nano Lett.* 2006 Apr;6(4):866-70. doi: 10.1021/nl052326h. PMID: 16608300.
- Cagliari, D., Dias, N. P., Galdeano, D. M., dos Santos, E. Á., Smagghe, G., & Zotti, M. J. (2019). Management of pest insects and plant diseases by non-transformative RNAi. *Frontiers in Plant Science*, 10, 1319.  
<https://doi.org/10.3389/fpls.2019.01319>
- Campos, E. V. R., de Oliveira, J. L., Fraceto, L. F., & Singh, B. (2015). Polysaccharides as safer release systems for agrochemicals. *Agronomy for sustainable development*, 35(1), 47-66.
- Cao, L., Zhou, Z., Niu, M., Sun, X., Xin, X., Zhang, J., ... & Qiu, X. (2016). Positive-charge functionalized mesoporous silica nanoparticles as nanocarriers for controlled 2,4-dichlorophenoxyacetic acid sodium salt release. *Journal of Agricultural and Food Chemistry*, 64(9), 2016-2024.  
<https://doi.org/10.1021/acs.jafc.5b06093>
- Cao, L., Zhou, Z., Niu, S., Cao, C., Li, X., Shan, Y., & Huang, Q. (2017). Positive-charge functionalized mesoporous silica nanoparticles as nanocarriers for controlled 2, 4-dichlorophenoxy acetic acid sodium salt release. *Journal of agricultural and food chemistry*, 66(26), 6594-6603.
- Cappelle, K., de Oliveira, C. F., Van Eynde, B., Christiaens, O., & Smagghe, G. (2016). The involvement of clathrin-mediated endocytosis and two Sid-1-like transmembrane proteins in double-stranded RNA uptake in the Colorado potato beetle midgut. *Insect Molecular Biology*, 25(3), 315-323.  
<https://doi.org/10.1111/imb.12222>
- Cappelle, K., de Oliveira, C. F., Van Eynde, B., Christiaens, O., & Smagghe, G. (2016). The involvement of clathrin-mediated endocytosis and two Sid-1-like transmembrane proteins in double-stranded RNA uptake in the Colorado potato beetle midgut. *Insect Molecular Biology*, 25(3), 315-323.  
<https://doi.org/10.1111/imb.12222>
- Chagnon, M., Kreuzweiser, D., Mitchell, E. A., Morrissey, C. A., Noome, D. A., & Van der Sluijs, J. P. (2015). Risks of large-scale use of systemic insecticides to ecosystem functioning and services. *Environmental Science and Pollution Research*, 22(1), 119-134. <https://doi.org/10.1007/s11356-014-3277-x>
- Chatterjee, A. K., Chakraborty, R., & Basu, T. (2014). Mechanism of antibacterial activity of copper nanoparticles. *Nanotechnology*, 25(13), 135101.  
<https://doi.org/10.1088/0957-4484/25/13/135101>
- Chaudhari, A. K., Singh, V. K., Kedia, A., Das, S., & Dubey, N. K. (2021). Essential oils and their bioactive compounds as eco-friendly novel green pesticides for management of storage insect pests: Prospects and retrospects. *Environmental Science and Pollution Research*, 28(15), 18918-18940.  
<https://doi.org/10.1007/s11356-021-12841-w>
- Cheema, A. I., Ahmed, T., Abbas, A., Noman, M., Zubair, M., & Shahid, M. (2022). Antimicrobial activity of the biologically synthesized zinc oxide nanoparticles against important rice pathogens. *Physiology and Molecular Biology of Plants*, 28(10), 1955-1967.
- Chen, J., Peng, H., Wang, X., Shao, F., Yuan, Z., & Han, H. (2014). Graphene oxide exhibits broad-spectrum antimicrobial activity against bacterial phytopathogens and fungal conidia by intertwining and membrane perturbation. *Nanoscale*, 6(3), 1879-1889.
- Choi, S. J., Oh, J. M., & Choy, J. H. (2009). Toxicological effects of inorganic nanoparticles on human lung cancer A549 cells. *Journal of inorganic biochemistry*, 103(3), 463-471.
- Choudhary, R. C., Kumaraswamy, R. V., Kumari, S., Sharma, S. S., Pal, A., Raliya, R., ... & Saharan, V. (2017). Cu-chitosan nanoparticle boost defense responses and plant growth in maize (*Zea mays* L.).

- Scientific Reports, 7(1), 9754. <https://doi.org/10.1038/s41598-017-08571-0>
- Christiaens, O., Whyard, S., Vélez, A. M., & Smagghe, G. (2020). Double-stranded RNA technology to control insect pests: Current status and challenges. *Frontiers in Plant Science*, 11, 451. <https://doi.org/10.3389/fpls.2020.00451>
- Colman, B. P., Arnaout, C. L., Anciaux, S., Gunsch, C. K., Hochella, M. F., Kim, B., ... & Bernhardt, E. S. (2013). Low concentrations of silver nanoparticles in biosolids cause adverse ecosystem responses under realistic field scenario. *PLoS ONE*, 8(2), e57189. <https://doi.org/10.1371/journal.pone.0057189>
- Cornelis, G., Doolette, C., Thomas, M., McLaughlin, M. J., Kirby, J. K., Beak, D. G., & Chittleborough, D. (2012). Retention and dissolution of engineered silver nanoparticles in natural soils. *Soil Science Society of America Journal*, 76(3), 891-902. <https://doi.org/10.2136/sssaj2011.0360>
- Cornelis, G., Hund-Rinke, K., Kuhlbusch, T., Van den Brink, N., & Nickel, C. (2014). Fate and bioavailability of engineered nanoparticles in soils: A review. *Critical Reviews in Environmental Science and Technology*, 44(24), 2720-2764. <https://doi.org/10.1080/10643389.2013.829767>
- Danhier, F., Ansorena, E., Silva, J. M., Coco, R., Le Breton, A., & Préat, V. (2012). PLGA-based nanoparticles: An overview of biomedical applications. *Journal of Controlled Release*, 161(2), 505-522. <https://doi.org/10.1016/j.jconrel.2012.01.043>
- Das, S., Debnath, N., Cui, Y., Unrine, J., & Palli, S. R. (2015). Chitosan, carbon quantum dot, and silica nanoparticle mediated dsRNA delivery for gene silencing in *Aedes aegypti*: A comparative analysis. *ACS Applied Materials & Interfaces*, 7(35), 19530-19535. <https://doi.org/10.1021/acsami.5b05232>
- Das, S., Patra, J. K., & Shin, H. S. (2020). Biosynthesis of silver nanoparticles using endophytic fungus *Aspergillus fumigatus* and its antibacterial and synergistic antibacterial activity. *3 Biotech*, 10(1), 35. <https://doi.org/10.1007/s13205-019-2040-9>
- Dimkpa, C. O., McLean, J. E., Britt, D. W., & Anderson, A. J. (2012). Bioactivity and biomodification of Ag, ZnO, and CuO nanoparticles with relevance to plant performance in agriculture. *Industrial Biotechnology*, 8(6), 344-357. <https://doi.org/10.1089/ind.2012.0028>
- Dimkpa, C. O., McLean, J. E., Latta, D. E., Manangón, E., Britt, D. W., Johnson, W. P., ... & Anderson, A. J. (2012). CuO and ZnO nanoparticles: Phytotoxicity, metal speciation, and induction of oxidative stress in sand-grown wheat. *Journal of Nanoparticle Research*, 14(9), 1125. <https://doi.org/10.1007/s11051-012-1125-9>
- Dimkpa, C. O., Singh, U., Bindran, P. S., Elmer, W. H., Gardea-Torresdey, J. L., & White, J. C. (2018). Exposure to weathered and fresh nanoparticle and ionic Zn in soil promotes grain yield and modulates nutrient acquisition in wheat (*Triticum aestivum* L.). *Journal of Agricultural and Food Chemistry*, 66(37), 9645-9656. <https://doi.org/10.1021/acs.jafc.8b03840>
- Dimkpa, C. O., White, J. C., Elmer, W. H., & Gardea-Torresdey, J. (2017). Nanoparticle and ionic Zn promote nutrient loading of sorghum grain under low NPK fertilization. *Journal of Agricultural and Food Chemistry*, 65(39), 8552-8559. <https://doi.org/10.1021/acs.jafc.7b02961>
- Dimkpa, C. O., Zeng, J., McLean, J. E., Britt, D. W., Zhan, J., & Anderson, A. J. (2012). Production of indole-3-acetic acid via the indole-3-acetamide pathway in the plant-beneficial bacterium *Pseudomonas chlororaphis* O6 is inhibited by ZnO nanoparticles but enhanced by CuO nanoparticles. *Applied and Environmental Microbiology*, 78(5), 1404-1410.
- Dimkpa, C., Zeng, J., McLean, J. E., Britt, D. W., Zhan, J., & Anderson, A. J. (2015). Production of indole-3-acetic acid via the indole-3-acetamide pathway in the plant-beneficial bacterium *Pseudomonas chlororaphis* O6 is inhibited by ZnO nanoparticles but enhanced by CuO nanoparticles. *Applied and Environmental Microbiology*, 81(7), 2490-2499. <https://doi.org/10.1128/AEM.04003-14>
- Dossa, G. S., Quibod, I., Atienza-Grande, G., Oliva, R., Maiss, E., Vera Cruz, C., & Wydra, K. (2020). Rice pyramided line IRBB67 (Xa4/Xa7) homeostasis under combined stress of high temperature and bacterial blight. *Scientific Reports*, 10(1),

683. <https://doi.org/10.1038/s41598-020-57499-5>
- Duceppe, N., & Tabrizian, M. (2009). Factors influencing the transfection efficiency of ultra low molecular weight chitosan/hyaluronic acid nanoparticles. *Biomaterials*, 30(13), 2625-2631. <https://doi.org/10.1016/j.biomaterials.2009.01.017>
- Durán, N., Durán, M., de Jesus, M. B., Seabra, A. B., Fávaro, W. J., & Nakazato, G. (2016). Silver nanoparticles: A new view on mechanistic aspects on antimicrobial activity. *Nanomedicine: Nanotechnology, Biology and Medicine*, 12(3), 789-799. <https://doi.org/10.1016/j.nano.2015.11.016>
- Dutta, P., Yasin, A., Kumari, A., Mahanta, M., & Sharma, A. (2023). Nano-bioformulation: A spanking new weapon for plant disease management. *Plant Health Archives*, 1(3), 123-129. <https://doi.org/10.54083/pha/1.3.2023/123-129>
- EFSA (European Food Safety Authority). (2018). Guidance on risk assessment of nanomaterials to be applied in the food and feed chain: Human and animal health. *EFSA Journal*, 16(10), e05327. <https://doi.org/10.2903/j.efsa.2018.5327>
- Elamawi, R. M., Al-Harbi, R. E., & Hendi, A. A. (2018). Biosynthesis and characterization of silver nanoparticles using *Trichoderma longibrachiatum* and their effect on phytopathogenic fungi. *Egyptian Journal of Biological Pest Control*, 28(1), 28. <https://doi.org/10.1186/s41938-018-0028-1>
- EPA (US Environmental Protection Agency). (2017). Pesticides: Regulating pesticides. Retrieved from <https://www.epa.gov/pesticides>
- Erhayem, M., & Sohn, M. (2014). Effect of humic acid source on humic acid adsorption onto titanium dioxide nanoparticles. *Science of the Total Environment*, 470-471, 92-98. <https://doi.org/10.1016/j.scitotenv.2013.09.063>
- FAOSTAT. (2024). Food and Agriculture Organization of the United Nations Statistics Division. Retrieved from <http://www.fao.org/faostat/>
- Fernández-Ortuño, D., Chen, F., and Schnabel, G. 2012. Resistance to pyraclostrobin and boscalid in *Botrytis cinerea* isolates from strawberry fields in the Carolinas. *Plant Dis.* 96:1198-1203. <https://doi.org/10.1094/PDIS-12-11-1049-RE> LinkWeb of ScienceGoogle Scholar
- Gajda, A., Posyniak, A., Tomczyk, G., Gbylik-Sikorska, M., & Bladek, T. (2020). Nanosilver impact on metabolism and antioxidant response in the fat body of honey bees (*Apis mellifera*). *Chemosphere*, 242, 125256. <https://doi.org/10.1016/j.chemosphere.2019.125256>
- Gao, J., Youn, S., Hovsepian, A., Llana, V. L., Wang, Y., Bitton, G., & Bonzongo, J. C. J. (2009). Dispersion and toxicity of selected manufactured nanomaterials in natural river water samples: Effects of water chemical composition. *Environmental Science & Technology*, 43(9), 3322-3328. <https://doi.org/10.1021/es803315v>
- Ge, Y., Schimel, J. P., & Holden, P. A. (2011). Evidence for negative effects of TiO<sub>2</sub> and ZnO nanoparticles on soil bacterial communities. *Environmental Science & Technology*, 45(4), 1659-1664. <https://doi.org/10.1021/es103040t>
- Ge, Y., Schimel, J. P., & Holden, P. A. (2012). Identification of soil bacteria susceptible to TiO<sub>2</sub> and ZnO nanoparticles. *Applied and Environmental Microbiology*, 78(18), 6749-6758. <https://doi.org/10.1128/AEM.00941-12>
- Ge, Y., Schimel, J. P., & Holden, P. A. (2012). Identification of soil bacteria susceptible to TiO<sub>2</sub> and ZnO nanoparticles. *Applied and Environmental Microbiology*, 78(18), 6749-6758. <https://doi.org/10.1128/AEM.00941-12>
- Ge, Y., Schimel, J. P., & Holden, P. A. (2011). Evidence for negative effects of TiO<sub>2</sub> and ZnO nanoparticles on soil bacterial communities. *Environmental Science & Technology*, 45(4), 1659-1664. <https://doi.org/10.1021/es103040t>
- Ghormade, V., Deshpande, M. V., & Paknikar, K. M. (2011). Perspectives for nanobiotechnology enabled protection and nutrition of plants. *Biotechnology Advances*, 29(6), 792-803. <https://doi.org/10.1016/j.biotechadv.2011.06.007>
- Grillo, R., Abhilash, P. C., & Fraceto, L. F. (2016). Nanotechnology applied to bioencapsulation of pesticides. *Journal of Nanoscience and Nanotechnology*, 16(2), 1231-1234. <https://doi.org/10.1166/jnn.2016.12332>
- Grillo, R., dos Santos, N. Z., Maruyama, C. R., Rosa, A. H., de Lima, R., & Fraceto, L. F.

- (2012). Poly( $\epsilon$ -caprolactone)nanocapsules as carrier systems for herbicides: Physico-chemical characterization and genotoxicity evaluation. *Journal of Hazardous Materials*, 231-232, 1-9. <https://doi.org/10.1016/j.jhazmat.2012.06.019>
- Grillo, R., Fraceto, L. F., Amorim, M. J., Scott-Fordsmand, J. J., Schoonjans, R., & Chaudhry, Q. (2021). Ecotoxicological and regulatory aspects of environmental sustainability of nanopesticides. *Journal of Hazardous Materials*, 404, 124148. <https://doi.org/10.1016/j.jhazmat.2020.124148>
- Grillo, R., Pereira, A. E., Nishisaka, C. S., de Lima, R., Oehlke, K., Greiner, R., & Fraceto, L. F. (2014). Chitosan/tripolyphosphate nanoparticles loaded with paraquat herbicide: An environmentally safer alternative for weed control. *Journal of Hazardous Materials*, 278, 163-171. <https://doi.org/10.1016/j.jhazmat.2014.05.079>
- Hajipour, M. J., Fromm, K. M., Ashkarran, A. A., Jimenez de Aberasturi, D., de Larramendi, I. R., Rojo, T., ... & Mahmoudi, M. (2012). *Antibacterial properties of nanoparticles*. *Trends in Biotechnology*, 30(10), 499-511. <https://doi.org/10.1016/j.tibtech.2012.06.004>
- Hansen, S. F., Michelson, E. S., Kamper, A., Borling, P., Stuer-Lauridsen, F., & Baun, A. (2008). Categorization framework to aid exposure assessment of nanomaterials in consumer products. *Ecotoxicology*, 17(5), 438-447.
- Hernandez-Viezcas, J. A., Castillo-Michel, H., Andrews, J. C., Cotte, M., Rico, C., Peralta-Videa, J. R., ... & Gardea-Torresdey, J. L. (2013). In situ synchrotron X-ray fluorescence mapping and speciation of CeO<sub>2</sub> and ZnO nanoparticles in soil cultivated soybean (*Glycine max*). *ACS Nano*, 7(2), 1415-1423. <https://doi.org/10.1021/nn305196q>
- Hladun, K. R., Smith, B. H., Mustard, J. A., Morton, R. R., & Trumble, J. T. (2012). Selenium toxicity to honey bee (*Apis mellifera* L.) pollinators: effects on behaviors and survival. *PLoS one*, 7(4), e34137.
- Hooper, H. L., Jurkschat, K., Morgan, A. J., Bailey, J., Lawlor, A. J., Spurgeon, D. J., & Svendsen, C. (2011). Comparative chronic toxicity of nanoparticulate and ionic zinc to the earthworm *Eisenia veneta* in a soil matrix. *Environment International*, 37(6), 1111-1117. <https://doi.org/10.1016/j.envint.2011.02.019>
- Hoppe, M., Mikutta, R., Utermann, J., Duijnisveld, W., & Guggenberger, G. (2015). Retention of sterically and electrosterically stabilized silver nanoparticles in soils. *Environmental Science & Technology*, 49(17), 10490-10497. <https://doi.org/10.1021/acs.est.5b01564>
- Hou, Y., Xiong, D., Jiang, T., Song, L., & Wang, Q. (2019). Social media addiction: Its impact, mediation, and intervention. *Cyberpsychology: Journal of Psychosocial Research on Cyberspace*, 13(1), Article 4. <https://doi.org/10.5817/CP2019-1-4>
- Huvenne, H., & Smagghe, G. (2010). Mechanisms of dsRNA uptake in insects and potential of RNAi for pest control: A review. *Journal of Insect Physiology*, 56(3), 227-235. <https://doi.org/10.1016/j.jinsphys.2009.10.004>
- Imada, K., Sakai, S., Kajihara, H., Tanaka, S., & Ito, S. (2016). Magnesium oxide nanoparticles induce systemic resistance in tomato against bacterial wilt disease. *Plant Pathology*, 65(4), 551-560. <https://doi.org/10.1111/ppa.12443>
- Iravani, S. (2011). Green synthesis of metal nanoparticles using plants. *Green Chemistry*, 13(10), 2638-2650. <https://doi.org/10.1039/C1GC15386B>
- Islam, M. T., Croll, D., Gladieux, P., Soanes, D. M., Persoons, A., Bhattacharjee, P., ... & Kamoun, S. (2016). Emergence of wheat blast in Bangladesh was caused by a South American lineage of *Magnaporthe oryzae*. *BMC Biology*, 14(1), 84. <https://doi.org/10.1186/s12915-016-0309-7>
- Jayaraj, R., Megha, P., & Sreedev, P. (2016). Organochlorine pesticides, their toxic effects on living organisms and their fate in the environment. *Interdisciplinary Toxicology*, 9(3-4), 90-100. <https://doi.org/10.1515/intox-2016-0012>
- Jiang, J., Pi, J., & Cai, J. (2018). The advancing of zinc oxide nanoparticles for biomedical applications. *Bioinorganic Chemistry and Applications*, 2018, 1062562. <https://doi.org/10.1155/2018/1062562>
- Judy, J. D., Kirby, J. K., Creamer, C., McLaughlin, M. J., Fiebiger, C., Wright, C., ... & Bertsch, P. M. (2015). Effects of silver

- sulfide nanomaterials on mycorrhizal colonization of tomato plants and soil microbial communities in biosolid-amended soil. *Environmental Pollution*, 206, 256-263.  
<https://doi.org/10.1016/j.envpol.2015.07.002>
- Judy, J. D., Unrine, J. M., & Bertsch, P. M. (2011). Evidence for biomagnification of gold nanoparticles within a terrestrial food chain. *Environmental Science & Technology*, 45(2), 776-781.  
<https://doi.org/10.1021/es103031a>
- Judy, J. D., Unrine, J. M., Rao, W., Bertsch, P. M., & Bertsch, P. M. (2012). Bioaccumulation of gold nanomaterials by *Manduca sexta* through dietary uptake of surface contaminated plant tissue. *Environmental Science & Technology*, 46(22), 12672-12678.  
<https://doi.org/10.1021/es303333w>
- Judy, J. D., Unrine, J. M., Rao, W., Wirick, S., & Bertsch, P. M. (2019). Bioavailability of gold nanomaterials to plants: Importance of particle size and surface coating. *Environmental Science & Technology*, 46(15), 8467-8474.  
<https://doi.org/10.1021/es3019397>
- Kah, M., & Hofmann, T. (2014). Nanopesticide research: Current trends and future priorities. *Environment International*, 63, 224-235.  
<https://doi.org/10.1016/j.envint.2013.11.015>
- Kah, M., Beulke, S., Tiede, K., & Hofmann, T. (2013). Nanopesticides: State of knowledge, environmental fate, and exposure modeling. *Critical Reviews in Environmental Science and Technology*, 43(16), 1823-1867.  
<https://doi.org/10.1080/10643389.2012.671750>
- Kah, M., Kookana, R. S., Gogos, A., & Bucheli, T. D. (2018). A critical evaluation of nanopesticides and nanofertilizers against their conventional analogues. *Nature Nanotechnology*, 13(8), 677-684. <https://doi.org/10.1038/s41565-018-0131-1>
- Kanhed, P., Birla, S., Gaikwad, S., Gade, A., Seabra, A. B., Rubilar, O., ... & Rai, M. (2014). *In vitro* antifungal efficacy of copper nanoparticles against selected crop pathogenic fungi. *Materials Letters*, 115, 13-17.  
<https://doi.org/10.1016/j.matlet.2013.10.011>
- Kean, T., & Thanou, M. (2010). Biodegradation, biodistribution and toxicity of chitosan. *Advanced Drug Delivery Reviews*, 62(1), 3-11.  
<https://doi.org/10.1016/j.addr.2009.09.004>
- Kheiri, A., Jorf, S. A. M., Malhipour, A., Saremi, H., & Nikkhah, M. (2016). Application of chitosan and chitosan nanoparticles for the control of *Fusarium head blight* of wheat (*Fusarium graminearum*) *in vitro* and greenhouse. *International Journal of Biological Macromolecules*, 93(Pt A), 1261-1272.  
<https://doi.org/10.1016/j.ijbiomac.2016.09.072>
- Khot, L. R., Sankaran, S., Maja, J. M., Ehsani, R., & Schuster, E. W. (2012). Applications of nanomaterials in agricultural production and crop protection: A review. *Crop Protection*, 35, 64-70.  
<https://doi.org/10.1016/j.cropro.2012.01.007>
- Kittler, S., Greulich, C., Diendorf, J., Köller, M., & Epple, M. (2010). Toxicity of silver nanoparticles increases during storage because of slow dissolution under release of silver ions. *Chemistry of Materials*, 22(16), 4548-4554.  
<https://doi.org/10.1021/cm100023p>
- Koppolu, B., & Zaharoff, D. A. (2013). The effect of antigen encapsulation in chitosan particles on uptake, activation and presentation by antigen presenting cells. *Biomaterials*, 34(9), 2359-2369.  
<https://doi.org/10.1016/j.biomaterials.2012.11.066>
- Kumar, N., Shah, V., & Walker, V. K. (2011). Perturbation of an arctic soil microbial community by metal nanoparticles. *Journal of Hazardous Materials*, 190(1-3), 816-822.  
<https://doi.org/10.1016/j.jhazmat.2011.04.005>
- Kumar, S., Bhanjana, G., Sharma, A., Sidhu, M. C., & Dilbaghi, N. (2015). Synthesis, characterization and on field evaluation of pesticide loaded sodium alginate nanoparticles. *Carbohydrate Polymers*, 101, 1061-1067.  
<https://doi.org/10.1016/j.carbpol.2013.10.025>
- Kumar, S., Nehra, M., Dilbaghi, N., Marrazza, G., Hassan, A. A., & Kim, K. H. (2020). Nano-based smart pesticide formulations: Emerging opportunities for agriculture. *Journal of Controlled Release*, 294, 131-153.

- <https://doi.org/10.1016/j.jconrel.2018.12.012>
- Kumar, S., Singh, A., & Kumar, A. (2017). Biopesticides for integrated crop management: Environmental and regulatory aspects. *Journal of Biofertilizers & Biopesticides*, 8(1), 1-6. <https://doi.org/10.4172/2155-6202.1000188>
- Kumari, A., Yadav, S. K., & Yadav, S. C. (2010). Biodegradable polymeric nanoparticles based drug delivery systems. *Colloids and Surfaces B: Biointerfaces*, 75(1), 1-18. <https://doi.org/10.1016/j.colsurfb.2009.09.001>
- Kuppusamy, P., Yusoff, M. M., Maniam, G. P., & Govindan, N. (2016). Biosynthesis of metallic nanoparticles using plant derivatives and their new avenues in pharmacological applications—An updated report. *Saudi Pharmaceutical Journal*, 24(4), 473-484. <https://doi.org/10.1016/j.jsps.2014.11.013>
- Lamsal, K., Kim, S. W., Jung, J. H., Kim, Y. S., Kim, K. S., & Lee, Y. S. (2011). Inhibition effects of silver nanoparticles against powdery mildews on cucumber and pumpkin. *Mycobiology*, 39(1), 26-32. <https://doi.org/10.4489/MYCO.2011.39.1.026>
- Lao, L. L., Peppas, N. A., Boey, F. Y., & Venkatraman, S. S. (2010). Modeling of drug release from biodegradable polymer blends. *European Journal of Pharmaceutics and Biopharmaceutics*, 70(3), 796-803. <https://doi.org/10.1016/j.ejpb.2008.05.024>
- Larue, C., Laurette, J., Herlin-Boime, N., Khodja, H., Fayard, B., Flank, A. M., ... & Carrière, M. (2012). Accumulation, translocation and impact of TiO<sub>2</sub> nanoparticles in wheat (*Triticum aestivum* spp.): Influence of diameter and crystal phase. *Science of the Total Environment*, 431, 197-208. <https://doi.org/10.1016/j.scitotenv.2012.04.073>
- Levard, C., Hotze, E. M., Lowry, G. V., & Brown, G. E. (2012). Environmental transformations of silver nanoparticles: Impact on stability and toxicity. *Environmental Science & Technology*, 46(13), 6900-6914. <https://doi.org/10.1021/es2037405>
- Li, J., Sang, H., Guo, H., Popko, J. T., He, K., White, J. C., ... & Xing, B. (2017). Antifungal mechanisms of ZnO and Ag nanoparticles to *Sclerotinia homoeocarpa*. *Nanotechnology*, 28(15), 155101. <https://doi.org/10.1088/1361-6528/aa61f3>
- Li, M., Zhu, L., & Lin, D. (2011). Toxicity of ZnO nanoparticles to *Escherichia coli*: Mechanism and the influence of medium components. *Environmental Science & Technology*, 45(5), 1977-1983. <https://doi.org/10.1021/es102624t>
- Li, Y., Zhang, F., Li, M., Shakoob, N., & Adeel, M. (2022). Application and mechanisms of metal-based nanoparticles in the control of bacterial and fungal crop diseases. *Pest Management Science*, 78(12), 4697-4710. <https://doi.org/10.1002/ps.7218>
- Li, Y., Zhang, W., Niu, J., & Chen, Y. (2012). Mechanism of photogenerated reactive oxygen species and correlation with the antibacterial properties of engineered metal-oxide nanoparticles. *ACS Nano*, 6(6), 5164-5173. <https://doi.org/10.1021/nn300934k>
- Liu, S., Ding, Z., Zhang, C., Yang, B., & Liu, Z. (2010). Gene knockdown by intro-thoracic injection of double-stranded RNA in the brown planthopper, *Nilaparvata lugens*. *Insect Biochemistry and Molecular Biology*, 40(9), 666-671. <https://doi.org/10.1016/j.ibmb.2010.06.007>
- Liu, S., Ding, Z., Zhang, C., Yang, B., & Liu, Z. (2010). Gene knockdown by intro-thoracic injection of double-stranded RNA in the brown planthopper, *Nilaparvata lugens*. *Insect Biochemistry and Molecular Biology*, 40(9), 666-671. <https://doi.org/10.1016/j.ibmb.2010.06.007>
- Lowry, G. V., Gregory, K. B., Apte, S. C., & Lead, J. R. (2012). Transformations of nanomaterials in the environment. *Environmental Science & Technology*, 46(13), 6893-6899. <https://doi.org/10.1021/es300839e>
- Lu, Z., Yu, X., Heong, K., & Hu, C. (2017). Effect of nitrogen fertilizer on herbivores and its stimulation to major insect pests in rice. *Rice Science*, 14(1), 56-66. [https://doi.org/10.1016/S1672-6308\(07\)60009-2](https://doi.org/10.1016/S1672-6308(07)60009-2)
- Ma, C., White, J. C., Dhankher, O. P., & Xing, B. (2015). Metal-based nanotoxicity and detoxification pathways in higher plants. *Environmental Science & Technology*, 49(12), 7109-7122. <https://doi.org/10.1021/acs.est.5b00685>
- Ma, R., Levard, C., Marinakos, S. M., Cheng, Y., Liu, J., Michel, F. M., ... & Lowry, G. V. (2013). Size-controlled dissolution of

- organic-coated silver nanoparticles. *Environmental Science & Technology*, 46(2), 752-759. <https://doi.org/10.1021/es201686j>
- Ma, X., Geiser-Lee, J., Deng, Y., & Kolmakov, A. (2010). Interactions between engineered nanoparticles (ENPs) and plants: Phytotoxicity, uptake and accumulation. *Science of the Total Environment*, 408(16), 3053-3061. <https://doi.org/10.1016/j.scitotenv.2010.03.031>
- Malandrakis, A. A., Kavroulakis, N., & Chrysikopoulos, C. V. (2019). Use of copper, silver and zinc nanoparticles against foliar and soil-borne plant pathogens. *Science of the Total Environment*, 670, 292-299. <https://doi.org/10.1016/j.scitotenv.2019.03.210>
- Malerba, M., & Cerana, R. (2016). Chitosan effects on plant systems. *International Journal of Molecular Sciences*, 17(7), 996. <https://doi.org/10.3390/ijms17070996>
- Mao, S., Sun, W., & Kissel, T. (2010). Chitosan-based formulations for delivery of DNA and siRNA. *Advanced Drug Delivery Reviews*, 62(1), 12-27. <https://doi.org/10.1016/j.addr.2009.08.004>
- Memarizadeh, N., Ghadamyari, M., Adeli, M., & Talebi, K. (2014). Preparation, characterization and efficiency of nanoencapsulated imidacloprid under laboratory conditions. *Ecotoxicology and Environmental Safety*, 107, 77-83. <https://doi.org/10.1016/j.ecoenv.2014.05.009>
- Milivojević, T., Glavan, G., & Božič, J. (2015). Neurotoxic potential of ingested titanium dioxide nanoparticles on bees. *Chemosphere*, 120, 547-554. <https://doi.org/10.1016/j.chemosphere.2014.09.031>
- Mishra, S., Singh, B. R., Singh, A., Keswani, C., Naqvi, A. H., & Singh, H. B. (2014). Biofabricated silver nanoparticles act as a strong fungicide against *Bipolaris sorokiniana* causing spot blotch disease in wheat. *PLoS ONE*, 9(5), e97881. <https://doi.org/10.1371/journal.pone.0097881>
- Mittal, A. K., Chisti, Y., & Banerjee, U. C. (2013). Synthesis of metallic nanoparticles using plant extracts. *Biotechnology Advances*, 31(2), 346-356. <https://doi.org/10.1016/j.biotechadv.2013.01.003>
- Mitter, N., Worrall, E. A., Robinson, K. E., Li, P., Jain, R. G., Taochy, C., ... & Pappu, H. R. (2017). Clay nanosheets for topical delivery of RNAi for sustained protection against plant viruses. *Nature Plants*, 3(2), 16207. <https://doi.org/10.1038/nplants.2016.207>
- Moll, J., Klingenfuss, F., Widmer, F., Gogos, A., van der Heijden, M. G., Hartmann, M., & Fliessbach, A. (2016). Effects of titanium dioxide nanoparticles on soil microbial communities and wheat biomass. *Soil Biology and Biochemistry*, 111, 85-93. <https://doi.org/10.1016/j.soilbio.2017.03.019>
- Molla, K. A., Karmakar, S., Chanda, P. K., Sarkar, S. N., Datta, S. K., & Datta, K. (2020). Tissue-specific expression of Arabidopsis NPR1 gene in rice for sheath blight resistance without compromising phenotypic cost. *Plant Science*, 250, 105-114. <https://doi.org/10.1016/j.plantsci.2016.06.005>
- Montaño, M. D., Olesik, J. W., Barber, A. G., Challis, K., & Ranville, J. F. (2014). Single particle ICP-MS: Advances toward routine analysis of nanomaterials. *Analytical and Bioanalytical Chemistry*, 408(19), 5053-5074. <https://doi.org/10.1007/s00216-016-9676-8>
- Morones, J. R., Elechiguerra, J. L., Camacho, A., Holt, K., Kouri, J. B., Ramírez, J. T., & Yacaman, M. J. (2005). The bactericidal effect of silver nanoparticles. *Nanotechnology*, 16(10), 2346-2353. <https://doi.org/10.1088/0957-4484/16/10/059>
- Mottaleb, K. A., Rejesus, R. M., Murty, M. V. R., Mohanty, S., & Li, T. (2017). Benefits of the development and dissemination of climate-smart rice: Ex ante impact assessment of drought-tolerant rice in South Asia. *Mitigation and Adaptation Strategies for Global Change*, 22(6), 879-901. <https://doi.org/10.1007/s11027-016-9705-0>
- Müller, R. H., Mäder, K., & Gohla, S. (2000). Solid lipid nanoparticles (SLN) for controlled drug delivery - a review of the state of the art. *European Journal of Pharmaceutics and Biopharmaceutics*, 50(1), 161-177. [https://doi.org/10.1016/S0939-6411\(00\)00087-4](https://doi.org/10.1016/S0939-6411(00)00087-4)

- Nayak, P. (2024). Nanoencapsulation of fungicides: New trend in plant disease control. In *Nanofungicides* (pp. 415-438). Elsevier. <https://doi.org/10.1016/b978-0-443-23950-2.00019-9>
- Nel, A. E., Mädler, L., Velegol, D., Xia, T., Hoek, E. M., Somasundaran, P., Klaessig, F., Castranova, V., & Thompson, M. (2009). Understanding biophysicochemical interactions at the nano-bio interface. *Nature materials*, 8(7), 543–557. <https://doi.org/10.1038/nmat2442>
- Niño-Liu DO, Ronald PC, Bogdanove AJ. *Xanthomonas oryzae* pathovars: model pathogens of a model crop. *Mol Plant Pathol*. 2006 Sep;7(5):303-24. doi: 10.1111/j.1364-3703.2006.00344.x. PMID: 20507449.
- Nuruzzaman, M., Rahman, M. M., Liu, Y., & Naidu, R. (2016). Nanoencapsulation, nano-guard for pesticides: A new window for safe application. *Journal of Agricultural and Food Chemistry*, 64(7), 1447-1483. <https://doi.org/10.1021/acs.jafc.5b05214>
- Ocsoy, I., Paret, M. L., Ocsoy, M. A., Kunwar, S., Chen, T., You, M., & Tan, W. (2013). Nanotechnology in plant disease management: DNA-directed silver nanoparticles on graphene oxide as an antibacterial against *Xanthomonas perforans*. *ACS Nano*, 7(10), 8972-8980. <https://doi.org/10.1021/nn4034794>
- Ogunyemi, S. O., Abdallah, Y., Zhang, M., Fouad, H., Hong, X., Ibrahim, E., ... & Li, B. (2019). Green synthesis of zinc oxide nanoparticles using different plant extracts and their antibacterial activity against *Xanthomonas oryzae* pv. *oryzae*. *Artificial Cells, Nanomedicine, and Biotechnology*, 47(1), 341-352. <https://doi.org/10.1080/21691401.2018.1557671>
- Ogunyemi, S. O., Zhang, M., Abdallah, Y., Ahmed, T., Qiu, W., Ali, M. A., ... & Li, B. (2020). The bio-synthesis of three metal oxide nanoparticles (ZnO, MnO<sub>2</sub>, and MgO) and their antibacterial activity against the bacterial leaf blight pathogen. *Frontiers in Microbiology*, 11, 588326. <https://doi.org/10.3389/fmicb.2020.588326>
- Parker, K. M., Barragán Borrero, V., van Leeuwen, D. M., Lever, M. A., Mateescu, B., & Sander, M. (2019). Environmental fate of RNA interference pesticides: Adsorption and degradation of double-stranded RNA molecules in agricultural soils. *Environmental Science & Technology*, 53(6), 3027-3036. <https://doi.org/10.1021/acs.est.8b05576>
- Pasquet, J., Chevalier, Y., Pelletier, J., Couval, E., Bouvier, D., & Bolzinger, M. A. (2014). The contribution of zinc ions to the antimicrobial activity of zinc oxide. *Colloids and Surfaces A: Physicochemical and Engineering Aspects*, 457, 263-274. <https://doi.org/10.1016/j.colsurfa.2014.05.057>
- Pereira, A. E., Grillo, R., Mello, N. F., Rosa, A. H., & Fraceto, L. F. (2014). Application of poly(epsilon-caprolactone) nanoparticles containing atrazine herbicide as an alternative technique to control weeds and reduce damage to the environment. *Journal of Hazardous Materials*, 268, 207-215. <https://doi.org/10.1016/j.jhazmat.2014.01.025>
- Pérez-de-Luque, A. (2017). Interaction of nanomaterials with plants: What do we need for real applications in agriculture? *Frontiers in Environmental Science*, 5, 12. <https://doi.org/10.3389/fenvs.2017.00012>
- Prasad, R., Bhattacharyya, A., & Nguyen, Q. D. (2017). Nanotechnology in sustainable agriculture: Recent developments, challenges, and perspectives. *Frontiers in Microbiology*, 8, 1014. <https://doi.org/10.3389/fmicb.2017.01014>
- Qi, L. S., Larson, M. H., Gilbert, L. A., Doudna, J. A., Weissman, J. S., Arkin, A. P., & Lim, W. A. (2013). Repurposing CRISPR as an RNA-guided platform for sequence-specific control of gene expression. *Cell*, 152(5), 1173–1183. <https://doi.org/10.1016/j.cell.2013.02.022>
- Qiu, J., Chen, Y., Liu, Z., Wen, H., & Jiang, N. (2023). The application of zinc oxide nanoparticles: An effective strategy to protect rice from rice blast and abiotic stresses. *Environmental Pollution*, 327, 121925. <https://doi.org/10.1016/j.envpol.2023.121925>
- Raghupathi, K. R., Koodali, R. T., & Manna, A. C. (2011). Size-dependent bacterial growth inhibition and mechanism of antibacterial activity of zinc oxide nanoparticles. *Langmuir*, 27(7), 4020-4028. <https://doi.org/10.1021/la104825u>
- Rajwade, J. M., Chikte, R. G., & Paknikar, K. M. (2020). Nanomaterials: New weapons in a crusade against phytopathogens. *Applied Microbiology and Biotechnology*, 104(4),

- 1437-1461.  
<https://doi.org/10.1007/s00253-019-10334-y>
- Raliya, R., Tarafdar, J. C., & Biswas, P. (2016). Enhancing the mobilization of native phosphorus in the mung bean rhizosphere using ZnO nanoparticles synthesized by soil fungi. *Journal of Agricultural and Food Chemistry*, 64(16), 3111-3118.  
<https://doi.org/10.1021/acs.jafc.5b05224>
- Ramachandran, R., Krishnaswamy, S., Shanmugam, S., & Venugopal, S. (2019). Alginate-based nanocarriers for the delivery of agrochemicals. In *Biopolymer-Based Formulations* (pp. 563-580). Elsevier. <https://doi.org/10.1016/B978-0-12-816897-4.00026-7>
- Rashid, M. M., Ahmed, N., Jahan, M., Islam, K. S., Nansen, C., Willers, J. L., & Haque, M. A. (2017). Higher fertilizer inputs increase fitness traits of brown planthopper in rice. *Scientific Reports*, 7(1), 4719. <https://doi.org/10.1038/s41598-017-05023-7>
- Ray, D. K., West, P. C., Clark, M., Gerber, J. S., Prishchepov, A. V., & Chatterjee, S. (2019). Climate change has likely already affected global food production. *PLoS ONE*, 14(5), e0217148. <https://doi.org/10.1371/journal.pone.0217148>
- Reinsch, B. C., Levard, C., Li, Z., Ma, R., Wise, A., Gregory, K. B., ... & Lowry, G. V. (2012). Sulfidation of silver nanoparticles decreases *Escherichia coli* growth inhibition. *Environmental Science & Technology*, 46(13), 6992-7000. <https://doi.org/10.1021/es203732x>
- Rico, C. M., Majumdar, S., Duarte-Gardea, M., Peralta-Videa, J. R., & Gardea-Torresdey, J. L. (2011). Interaction of nanoparticles with edible plants and their possible implications in the food chain. *Journal of Agricultural and Food Chemistry*, 59(8), 3485-3498.  
<https://doi.org/10.1021/jf104517j>
- Riedo, J., Wettstein, F. E., Rösch, A., Herzog, C., Banerjee, S., Büchi, L., ... & van der Heijden, M. G. (2021). Widespread occurrence of pesticides in organically managed agricultural soils—the ghost of a conventional agricultural past? *Environmental Science & Technology*, 55(5), 2919-2928. <https://doi.org/10.1021/acs.est.0c06405>
- Rossi, L., Zhang, W., Schwab, A. P., & Ma, X. (2017). Uptake, accumulation, and in planta distribution of coexisting cerium oxide nanoparticles and cadmium in *Glycine max* (L.) Merr. *Environmental Science & Technology*, 51(21), 12815-12824.  
<https://doi.org/10.1021/acs.est.7b03363>
- Saharan, V., Sharma, G., Yadav, M., Choudhary, M. K., Sharma, S. S., Pal, A., ... & Biswas, P. (2015). Synthesis and *in vitro* antifungal efficacy of Cu-chitosan nanoparticles against pathogenic fungi of tomato. *International Journal of Biological Macromolecules*, 75, 346-353. <https://doi.org/10.1016/j.ijbiomac.2015.01.027>
- Sánchez-Bayo, F., & Hyne, R. V. (2014). Detection and analysis of neonicotinoids in river waters—Development of a passive sampler for three commonly used insecticides. *Chemosphere*, 99, 143-151. <https://doi.org/10.1016/j.chemosphere.2013.10.051>
- Savary, S., Willocquet, L., Pethybridge, S. J., Esker, P., McRoberts, N., & Nelson, A. (2019). The global burden of pathogens and pests on major food crops. *Nature Ecology & Evolution*, 3(3), 430-439. <https://doi.org/10.1038/s41559-018-0793-y>
- Schlich, K., Klawonn, T., Terytze, K., & Hund-Rinke, K. (2013). Effects of silver nanoparticles and silver nitrate in the earthworm reproduction test. *Environmental Toxicology and Chemistry*, 32(1), 181-188. <https://doi.org/10.1002/etc.2030>
- Schwab, F., Zhai, G., Kern, M., Turner, A., Schnoor, J. L., & Wiesner, M. R. (2016). Barriers, pathways and processes for uptake, translocation and accumulation of nanomaterials in plants—Critical review. *Nanotoxicology*, 10(3), 257-278. <https://doi.org/10.3109/17435390.2015.1048326>
- Scott, J. G., Michel, K., Bartholomay, L. C., Siegfried, B. D., Hunter, W. B., Smagghe, G., ... & Douglas, A. E. (2013). Towards the elements of successful insect RNAi. *Journal of Insect Physiology*, 59(12), 1212-1221. <https://doi.org/10.1016/j.jinsphys.2013.08.014>
- Shoultz-Wilson, W. A., Reinsch, B. C., Tsyusko, O. V., Bertsch, P. M., Lowry, G. V., & Unrine, J. M. (2011). Effect of silver nanoparticle surface coating on

- bioaccumulation and reproductive toxicity in earthworms (*Eisenia fetida*). *Nanotoxicology*, 5(3), 432-444. <https://doi.org/10.3109/17435390.2010.537382>
- Shukla, J. N., Kalsi, M., Sethi, A., Narva, K. E., Fishilevich, E., Singh, S., ... & Palli, S. R. (2016). Reduced stability and intracellular transport of dsRNA contribute to poor RNAi response in lepidopteran insects. *RNA Biology*, 13(7), 656-669. <https://doi.org/10.1080/15476286.2016.1191728>
- Siddiqi, K. S., & Husen, A. (2017). Fabrication of metal nanoparticles from fungi and metal salts: Scope and application. *Nanoscale Research Letters*, 12(1), 98. <https://doi.org/10.1186/s11671-017-1921-8>
- Simonin, M., & Richaume, A. (2015). Impact of engineered nanoparticles on the activity, abundance, and diversity of soil microbial communities: A review. *Environmental Science and Pollution Research*, 22(18), 13710-13723. <https://doi.org/10.1007/s11356-015-4171-x>
- Singh, A., Gautam, P. K., Verma, A., Singh, V., Shivapriya, P. M., Shivalkar, S., ... & Kumar, S. (2020). Green synthesis of metallic nanoparticles as effective alternatives to treat antibacterial resistant infections: A review. *Biotechnology Reports*, 25, e00427. <https://doi.org/10.1016/j.btre.2020.e00427>
- Singh, R. K., Singh, P., Rutkoski, J., Hodson, D. P., He, X., Jørgensen, L. N., ... & Singh, P. K. (2021). Disease impact on wheat yield potential and prospects of genetic control. *Annual Review of Phytopathology*, 59, 413-441. <https://doi.org/10.1146/annurev-phyto-020620-114752>
- Sirelkhatim, A., Mahmud, S., Seeni, A., Kaus, N. H. M., Ann, L. C., Bakhori, S. K. M., ... & Mohamad, D. (2015). Review on zinc oxide nanoparticles: Antibacterial activity and toxicity mechanism. *Nano-Micro Letters*, 7(3), 219-242. <https://doi.org/10.1007/s40820-015-0040-x>
- Slavin, Y. N., Asnis, J., Häfeli, U. O., & Bach, H. (2017). Metal nanoparticles: Understanding the mechanisms behind antibacterial activity. *Journal of Nanobiotechnology*, 15(1), 65. <https://doi.org/10.1186/s12951-017-0308-z>
- Sparks, T. C., & Nauen, R. (2015). IRAC: Mode of action classification and insecticide resistance management. *Pesticide Biochemistry and Physiology*, 121, 122-128. <https://doi.org/10.1016/j.pestbp.2014.11.014>
- Stehle, S., & Schulz, R. (2015). Agricultural insecticides threaten surface waters at the global scale. *Proceedings of the National Academy of Sciences*, 112(18), 5750-5755. <https://doi.org/10.1073/pnas.1500232112>
- Suriyaprabha, R., Karunakaran, G., Yuvakkumar, R., Prabu, P., Rajendran, V., & Kannan, N. (2014). Growth and physiological responses of maize (*Zea mays* L.) to porous silica nanoparticles in soil. *Journal of Nanoparticle Research*, 14(12), 1294. <https://doi.org/10.1007/s11051-012-1294-6>
- Tahara, K., Samura, S., Tsuji, K., Yamamoto, H., Tsukada, Y., Bando, Y., ... & Kawashima, Y. (2011). Oral nuclear factor-κB decoy oligonucleotides delivery system with chitosan modified poly(D,L-lactide-co-glycolide) nanospheres for inflammatory bowel disease. *Biomaterials*, 32(3), 870-878. <https://doi.org/10.1016/j.biomaterials.2010.09.034>
- Talbot, N. J., Kershaw, M. J., & Wakley, G. E. (2021). Advances in understanding the molecular basis of fungal pathogenicity to rice. *Annual Review of Phytopathology*, 59, 263-287. <https://doi.org/10.1146/annurev-phyto-020620-121858>
- Tan, J., Levine, S. L., Bachman, P. M., Jensen, P. D., Mueller, G. M., Uffman, J. P., ... & Heck, G. R. (2016). No impact of DvSnf7 RNA on honey bee (*Apis mellifera* L.) adults and larvae in dietary feeding tests. *Environmental Toxicology and Chemistry*, 35(2), 287-294. <https://doi.org/10.1002/etc.3075>
- Taning, C. N. T., Arpaia, S., Christiaens, O., Dietz-Pfeilstetter, A., Jones, H., Mezzetti, B., ... & Smagghe, G. (2020). RNA-based biocontrol compounds: Current status and perspectives to reach the market. *Pest Management Science*, 76(3), 841-845. <https://doi.org/10.1002/ps.5686>
- Terenius, O., Papanicolaou, A., Garbutt, J. S., Eleftherianos, I., Huvenne, H., Kanginakudru, S., ... & Swevers, L. (2011). RNA interference in Lepidoptera: An overview of successful and unsuccessful studies and implications for experimental design. *Journal of Insect Physiology*, 57(2), 231-245.

- <https://doi.org/10.1016/j.jinsphys.2010.11.006>
- Tripathi, D. K., Singh, S., Singh, S., Pandey, R., Singh, V. P., Sharma, N. C., ... & Chauhan, D. K. (2017). An overview on manufactured nanoparticles in plants: Uptake, translocation, accumulation and phytotoxicity. *Plant Physiology and Biochemistry*, 110, 2-12. <https://doi.org/10.1016/j.plaphy.2016.07.030>
- Van der Ploeg, M. J., Baveco, J. M., Van der Hout, A., Bakker, R., Rietjens, I. M., & Van den Brink, N. W. (2011). Effects of C60 nanoparticle exposure on earthworms (*Lumbricus rubellus*) and implications for population dynamics. *Environmental Pollution*, 159(1), 198-203. <https://doi.org/10.1016/j.envpol.2010.09.003>
- Vance, M. E., Kuiken, T., Vejerano, E. P., McGinnis, S. P., Hochella, M. F., Rejeski, D., & Hull, M. S. (2015). Nanotechnology in the real world: Redeveloping the nanomaterial consumer products inventory. *Beilstein Journal of Nanotechnology*, 6, 1769-1780. <https://doi.org/10.3762/bjnano.6.181>
- Wang, Z., Xie, X., Zhao, J., Liu, X., Feng, W., White, J. C., & Xing, B. (2012). Xylem- and phloem-based transport of CuO nanoparticles in maize (*Zea mays* L.). *Environmental Science & Technology*, 46(8), 4434-4441. <https://doi.org/10.1021/es204212z>
- Whitley, A. R., Levard, C., Oostveen, E., Bertsch, P. M., Matocha, C. J., von der Kammer, F., & Unrine, J. M. (2013). Behavior of Ag nanoparticles in soil: Effects of particle surface coating, aging and sewage sludge amendment. *Environmental Pollution*, 182, 141-149. <https://doi.org/10.1016/j.envpol.2013.06.027>
- Worrall, E. A., Hamid, A., Mody, K. T., Mitter, N., & Pappu, H. R. (2018). Nanotechnology for plant disease management. *Agronomy*, 8(12), 285. <https://doi.org/10.3390/agronomy8120285>
- Xie, Y., He, Y., Irwin, P. L., Jin, T., & Shi, X. (2011). Antibacterial activity and mechanism of action of zinc oxide nanoparticles against *Campylobacter jejuni*. *Applied and Environmental Microbiology*, 77(7), 2325-2331. <https://doi.org/10.1128/AEM.02149-10>
- Xiu, Z. M., Zhang, Q. B., Puppala, H. L., Colvin, V. L., & Alvarez, P. J. (2012). Negligible particle-specific antibacterial activity of silver nanoparticles. *Nano Letters*, 12(8), 4271-4275. <https://doi.org/10.1021/nl301934w>
- Yan, S., Ren, B., & Zeng, B. (2020). Delivery of dsRNA via layered double hydroxide nanoparticles for pest control. *Pest Management Science*, 76(8), 2623-2629. <https://doi.org/10.1002/ps.5803>
- Yan, Y., Huang, Z., Ding, M., Gong, M., & Liu, J. (2018). Layered double hydroxide-based herbicide nanohybrids with improved slow-release performance. *Journal of Agricultural and Food Chemistry*, 66(41), 10851-10858. <https://doi.org/10.1021/acs.jafc.8b04137>
- Zaller, J. G., Cantelmo, C., Dos Santos, G., Muther, S., Gruber, E., Pallua, P., ... & Ehling-Schulz, M. (2014). Herbicides in vineyards reduce grapevine root mycorrhization and alter soil microorganisms and the nutrient composition in grapevine roots, leaves, xylem sap and grape juice. *Environmental Science and Pollution Research*, 21(2), 1816-1825. <https://doi.org/10.1007/s11356-013-2073-y>
- Zha, W., Peng, X., Chen, R., Du, B., Zhu, L., & He, G. (2011). Knockdown of midgut genes by dsRNA-transgenic plant-mediated RNA interference in the hemipteran insect *Nilaparvata lugens*. *PLoS ONE*, 6(5), e20504. <https://doi.org/10.1371/journal.pone.0020504>
- Zhang, J., Khan, S. A., Hasse, C., Ruf, S., Heckel, D. G., & Bock, R. (2015). Full crop protection from an insect pest by expression of long double-stranded RNAs in plastids. *Science*, 347(6225), 991-994. <https://doi.org/10.1126/science.1261680>
- Zhang, X., Mysore, K., Flannery, E., Michel, K., Severson, D. W., Zhu, K. Y., & Duman-Scheel, M. (2015). Chitosan/interfering RNA nanoparticle mediated gene silencing in disease vector mosquito larvae. *Journal of Visualized Experiments*, (97), e52523. <https://doi.org/10.3791/52523>
- Zhang, Y., Chen, Y., Westerhoff, P., & Crittenden, J. (2009). Impact of natural organic matter and divalent cations on the stability of aqueous nanoparticles. *Water Research*, 43(17), 4249-4257.

- <https://doi.org/10.1016/j.watres.2009.06.005>
- Zhang, Y., Yan, Y., Zhao, X., Zhou, N., & Huang, Z. (2018). Chitosan-based delivery systems for RNAi in pest management. *Carbohydrate Polymers*, 197, 606-614. <https://doi.org/10.1016/j.carbpol.2018.06.038>
- Zhao, C. M., & Wang, W. X. (2012). Importance of surface coatings and soluble silver in silver nanoparticles toxicity to *Daphnia magna*. *Nanotoxicology*, 6(4), 361-370. <https://doi.org/10.3109/17435390.2011.579632>
- Zhao, L., Peralta-Videa, J. R., Ren, M., Varela-Ramirez, A., Li, C., Hernandez-Viezcas, J. A., ... & Gardea-Torresdey, J. L. (2012). Transport of Zn in a sandy loam soil treated with ZnO NPs and uptake by corn plants: Electron microprobe and confocal microscopy studies. *Chemical Engineering Journal*, 184, 1-8. <https://doi.org/10.1016/j.cej.2012.01.041>
- Zhao, L., Sun, Y., Hernandez-Viezcas, J. A., Servin, A. D., Hong, J., Niu, G., ... & Gardea-Torresdey, J. L. (2013). Influence of CeO<sub>2</sub> and ZnO nanoparticles on cucumber physiological markers and bioaccumulation of Ce and Zn: A life cycle study. *Journal of Agricultural and Food Chemistry*, 61(49), 11945-11951. <https://doi.org/10.1021/jf404328e>
- Zhao, X., Cui, H., Wang, Y., Sun, C., Cui, B., & Zeng, Z. (2018). Development strategies and prospects of nano-based smart pesticide formulation. *Journal of Agricultural and Food Chemistry*, 66(26), 6504-6512. <https://doi.org/10.1021/acs.jafc.7b02004>
- Zhao, X., Meng, Z., Wang, Y., Chen, W., Sun, C., Cui, B., ... & Zeng, Z. (2020). Pollen magnetofection for genetic modification with magnetic nanoparticles as gene carriers. *Nature Plants*, 3(12), 956-964. <https://doi.org/10.1038/nplants.2017.145>
- Zhao, Y., Zhang, Y., Yan, Y., Huang, Z., Zhang, Y., & Zhou, N. (2024). pH-responsive pesticide-loaded hollow mesoporous silica nanoparticles with ZnO quantum dots as a gatekeeper for control of rice blast disease. *Materials*, 17(6), 1344. <https://doi.org/10.3390/ma17061344>
- Zhu, F., Xu, J., Palli, R., Ferguson, J., & Palli, S. R. (2011). Ingested RNA interference for managing the populations of the Colorado potato beetle, *Leptinotarsa decemlineata*. *Pest Management Science*, 67(2), 175-182. <https://doi.org/10.1002/ps.2048>
- Zhu, X., Wang, J., Zhang, X., Chang, Y., & Chen, Y. (2010). Trophic transfer of TiO<sub>2</sub> nanoparticles from daphnia to zebrafish in a simplified freshwater food chain. *Chemosphere*, 79(9), 928-933. <https://doi.org/10.1016/j.chemosphere.2010.03.022>
- Zhu, X., Zhu, L., Duan, Z., Qi, R., Li, Y., & Lang, Y. (2008). Comparative toxicity of several metal oxide nanoparticle aqueous suspensions to zebrafish (*Danio rerio*) early developmental stage. *Journal of Environmental Science and Health, Part A*, 43(3), 278-284. <https://doi.org/10.1080/10934520701792779>
- Zotti, M., & Smaghe, G. (2015). RNAi technology for insect management and protection of beneficial insects from diseases: Lessons, challenges and risk assessments. *Neotropical Entomology*, 44(3), 197-213. <https://doi.org/10.1007/s13744-015-0291-8>

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